



Università di Napoli Federico II
Dipartimento di Agraria

Stato dell'arte sull'uso degli acidi nucleici nel biocontrollo

Prof. Stefano Mazzoleni



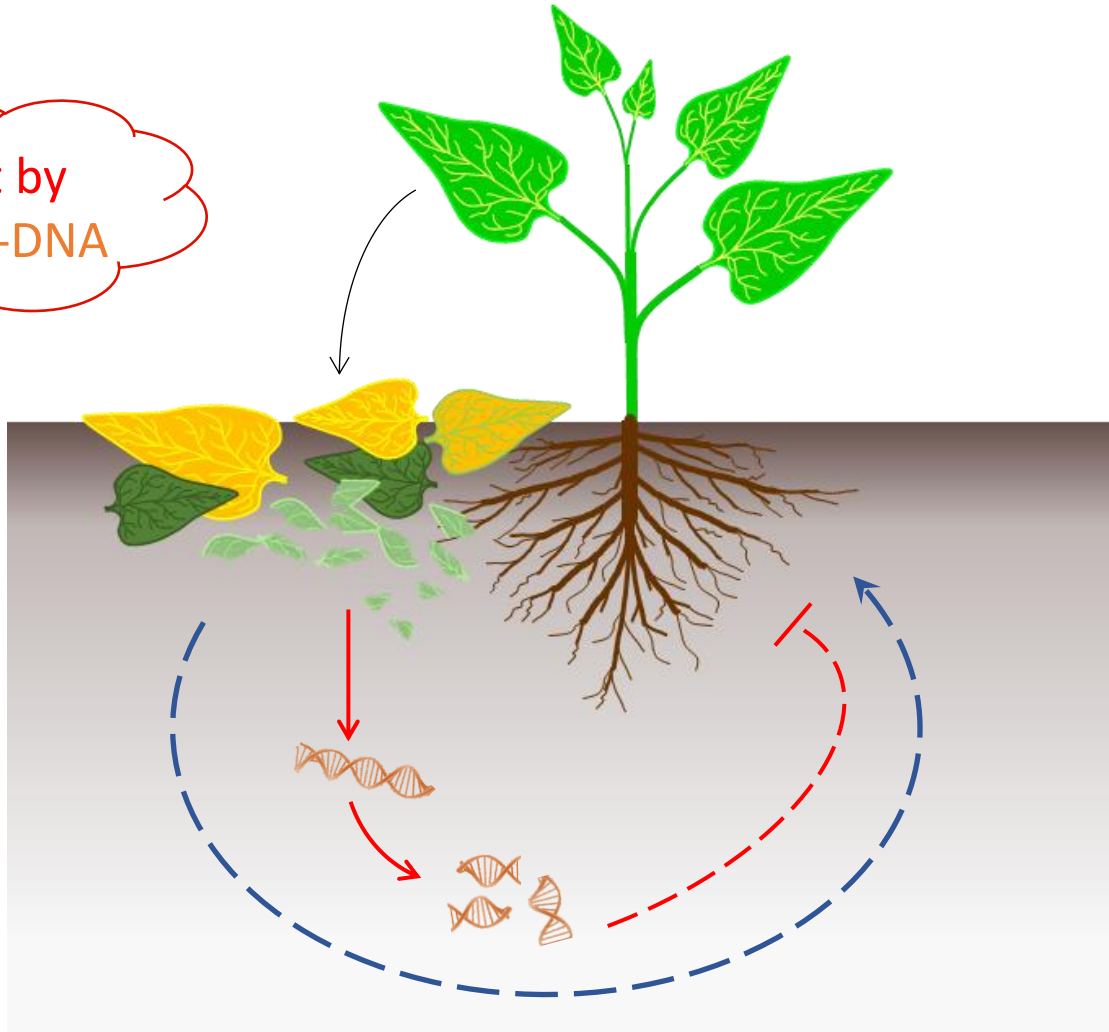
Napoli 28 Ottobre 2025

Decomposizione della sostanza organica: un processo chiave nel ciclo dei nutrienti



Model idea

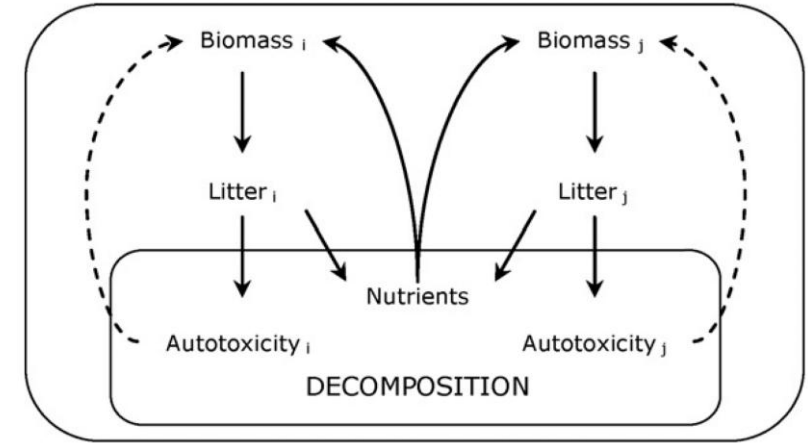
Inhibitory effect by
extracellular SELF-DNA



Nutrient cycling and organic matter decomposition

Modelling the effects of litter decomposition on tree diversity patterns

Mazzoleni et al. (2010) *Ecological Modelling*

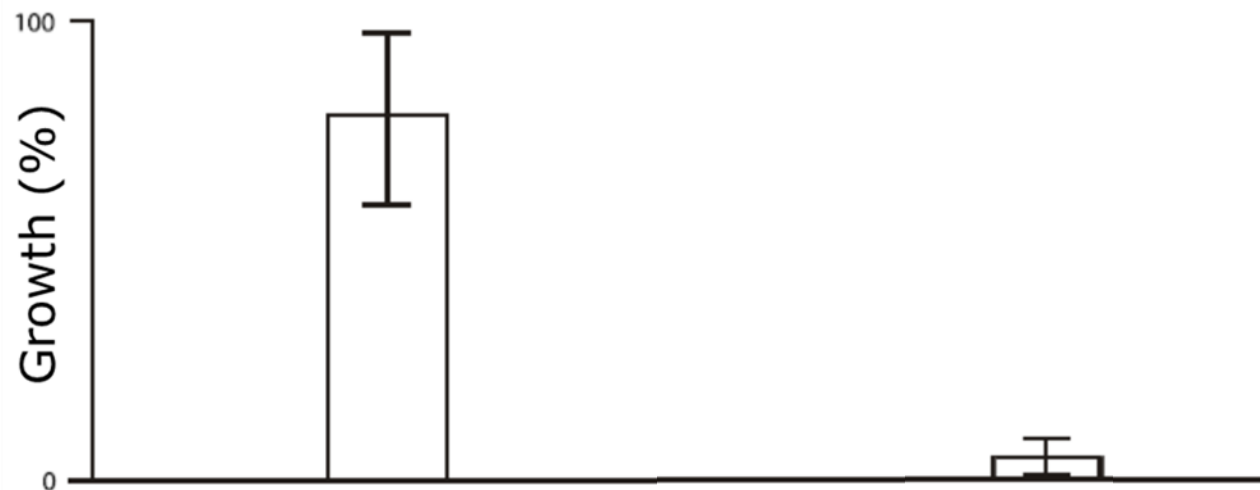


Inibizione da self-DNA nelle piante: dimostrata in > 30 species



Inhibitory and toxic effects of extracellular self-DNA in litter:
a mechanism for negative plant-soil feedbacks?

Mazzoleni et al. New Phytologist 2015 a

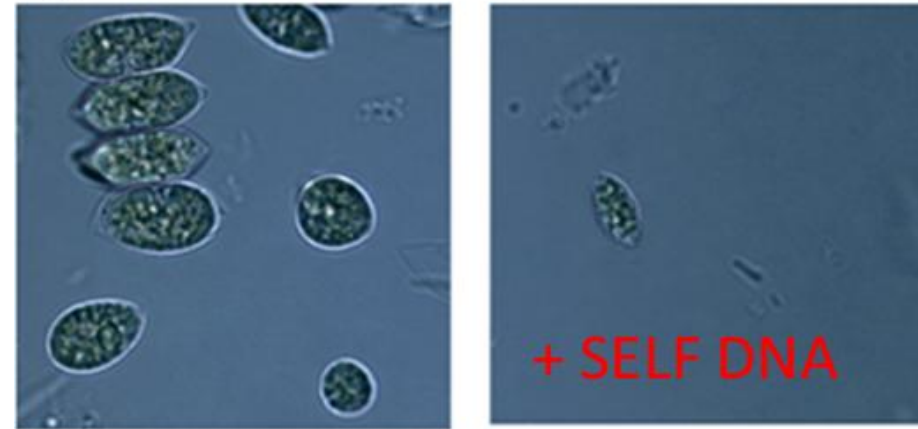


Effetto auto-inibitorio non solo per le piante: **é un fenomeno biologico generale!**

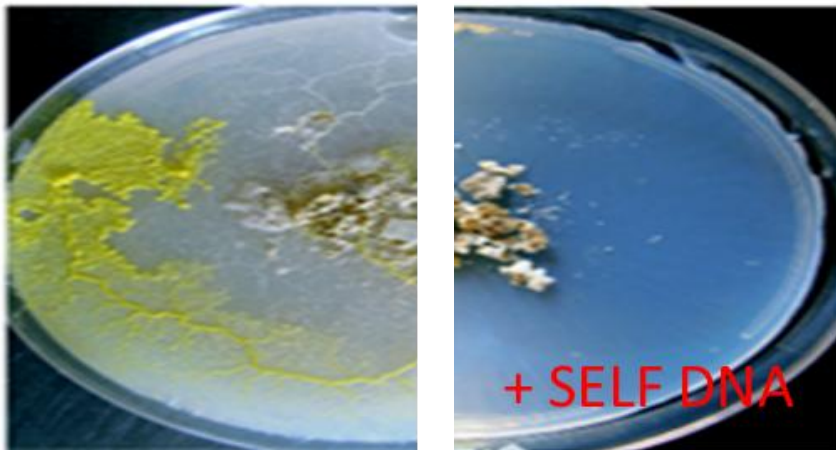
Bacteria: *Bacillus subtilis*



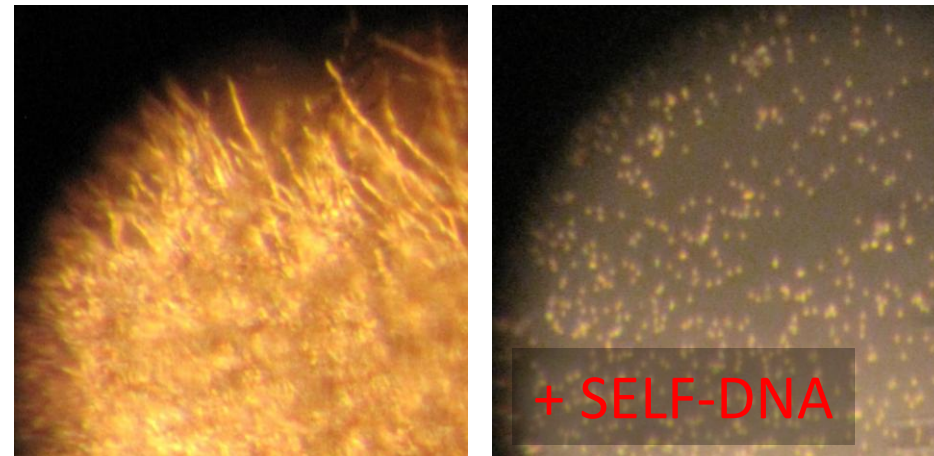
Algae: *Scenedesmus obliquus*



Protozoa: *Physarum polycephalum*



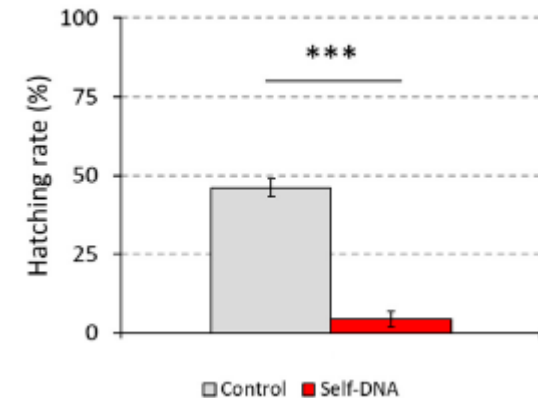
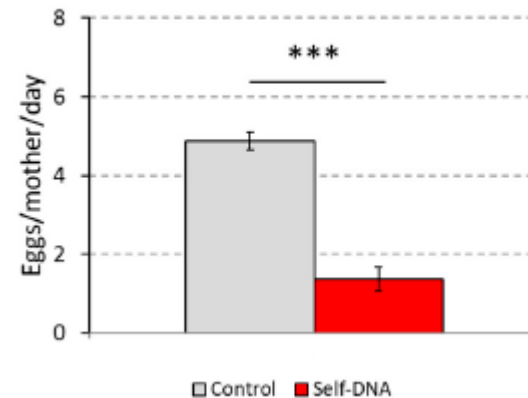
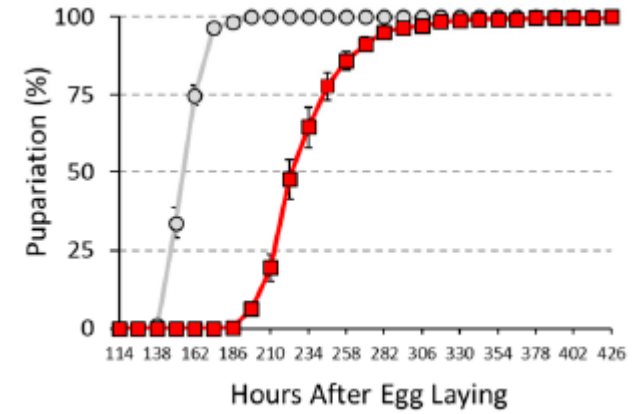
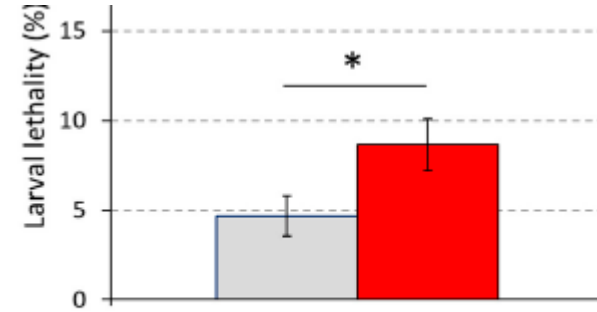
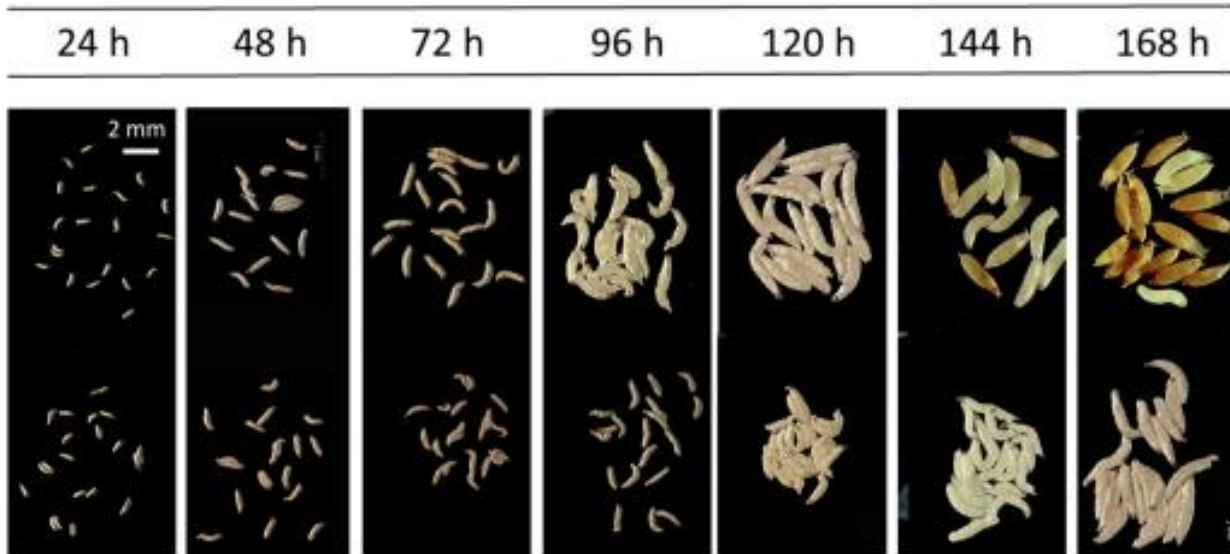
Fungi: *Trichoderma harzianum*



Experimental evidence: studies in model organisms



Article
Self-DNA Inhibition in *Drosophila melanogaster* Development: Metabolomic Evidence of the Molecular Determinants



Experimental evidence: studies in model organisms

α RAD-51



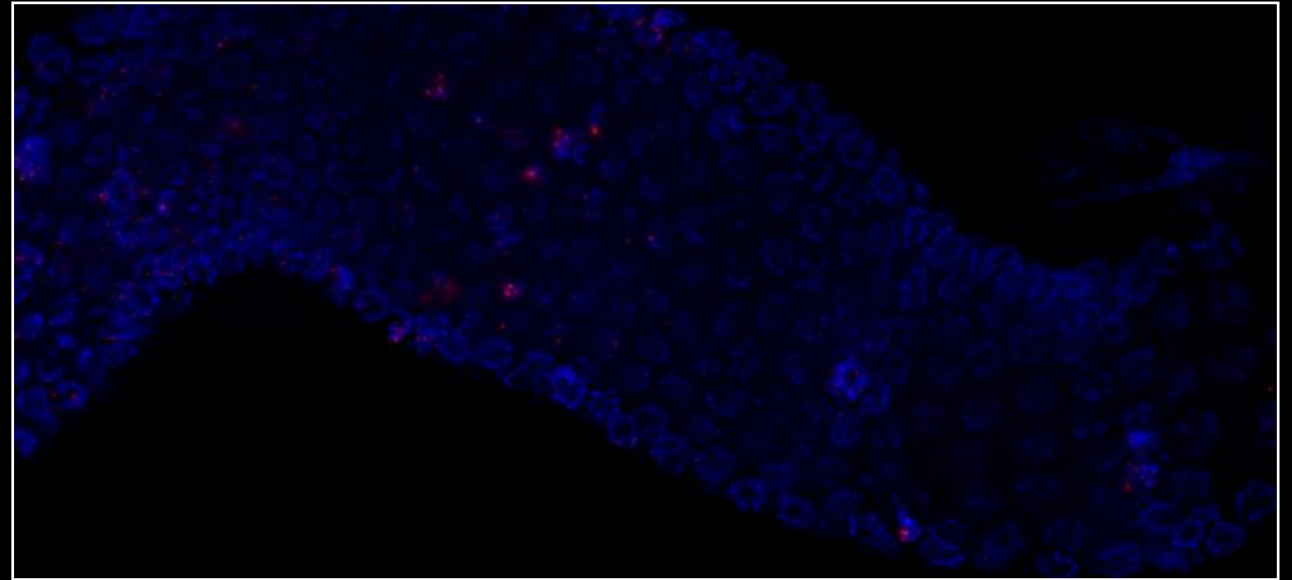
biology



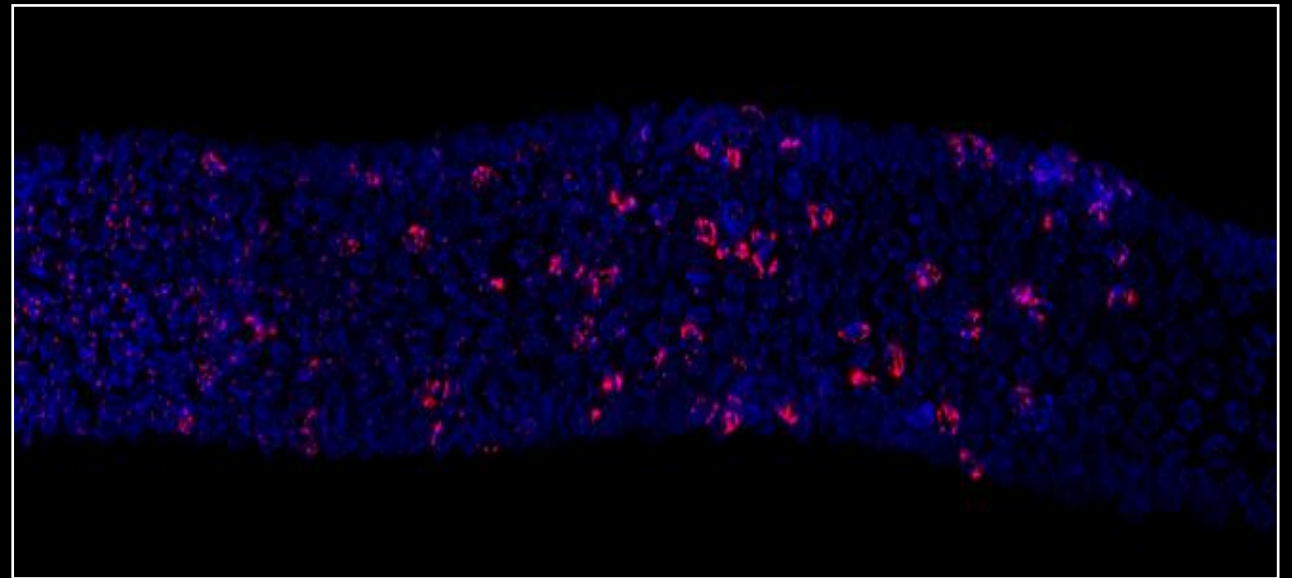
Article

Self-DNA Exposure Induces Developmental Defects and Germline DNA Damage Response in *Caenorhabditis elegans*

nonsel-DNA



self-DNA



Molecular mechanisms



Contents lists available at ScienceDirect

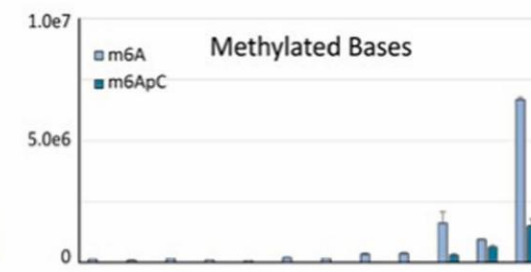
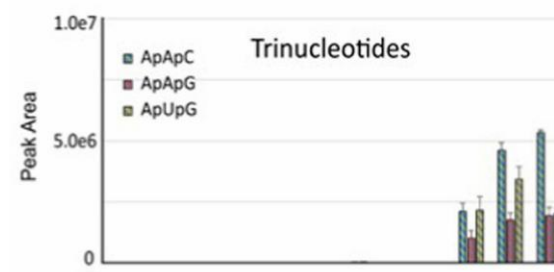
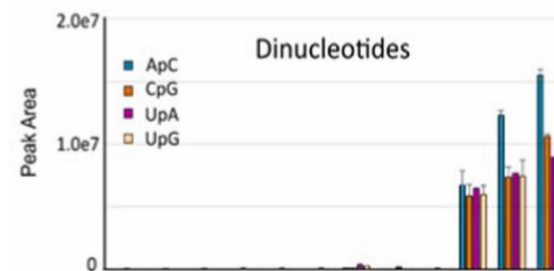
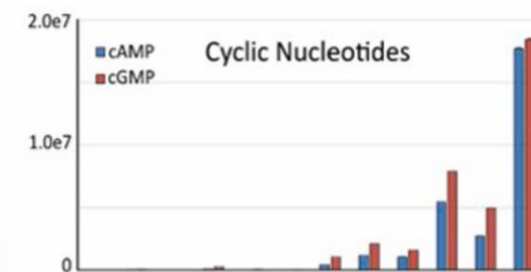
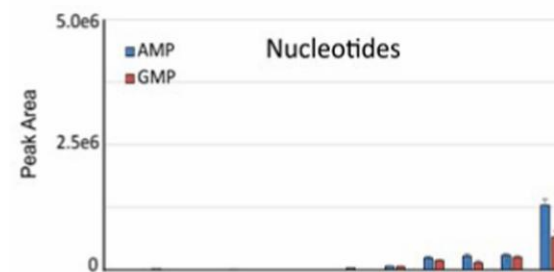
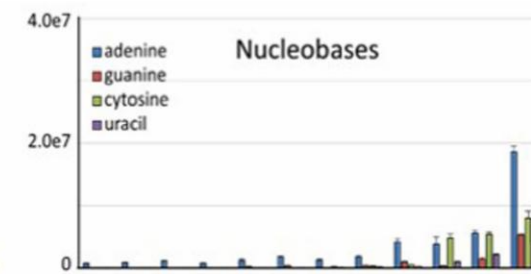
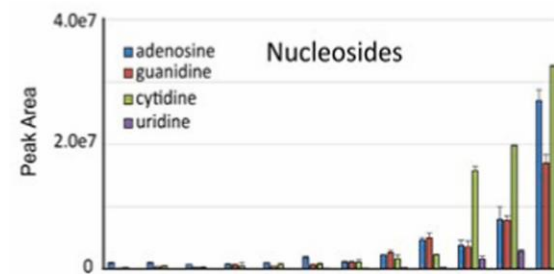
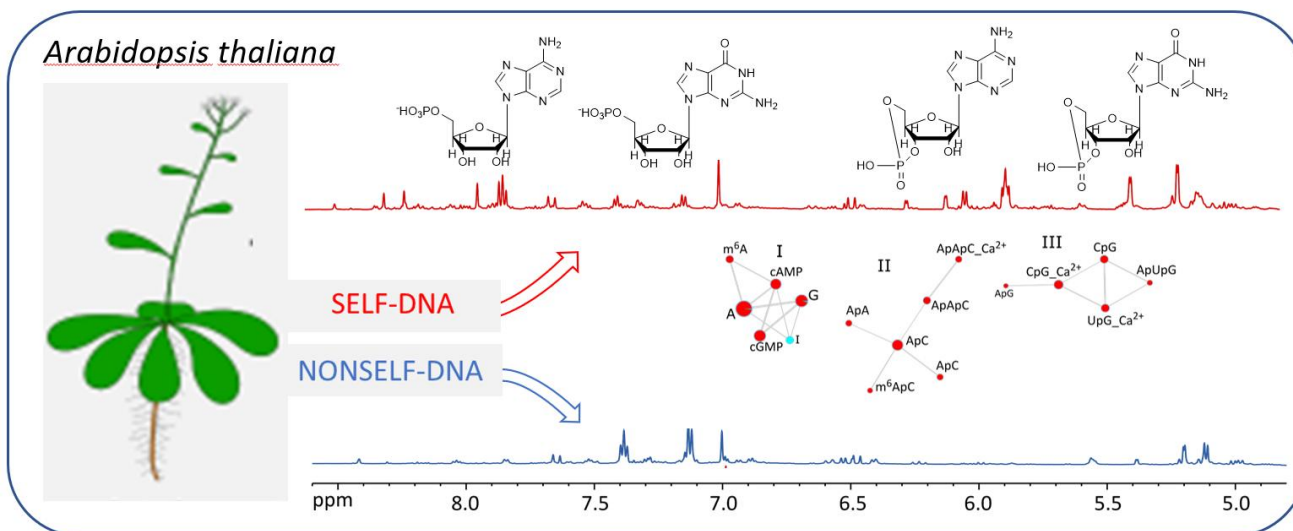
Phytochemistry

journal homepage: www.elsevier.com/locate/phytochem



Metabolomics and molecular networking analyses in *Arabidopsis thaliana* show that extracellular self-DNA affects nucleoside/nucleotide cycles with accumulation of cAMP, cGMP and N6-methyl-AMP

Virginia Lanzotti^{a,*}, Laura Grauso^a, Alfonso Mangoni^b, Pasquale Termolino^c, Emanuela Palomba^c, Attilio Anzano^a, Guido Incerti^d, Stefano Mazzoleni^{a,*}



1 10 15 1 10 15 1 10 15 1 10 15 1 10 15
C N₂ N_C S C N₂ N_C S

Molecular mechanisms

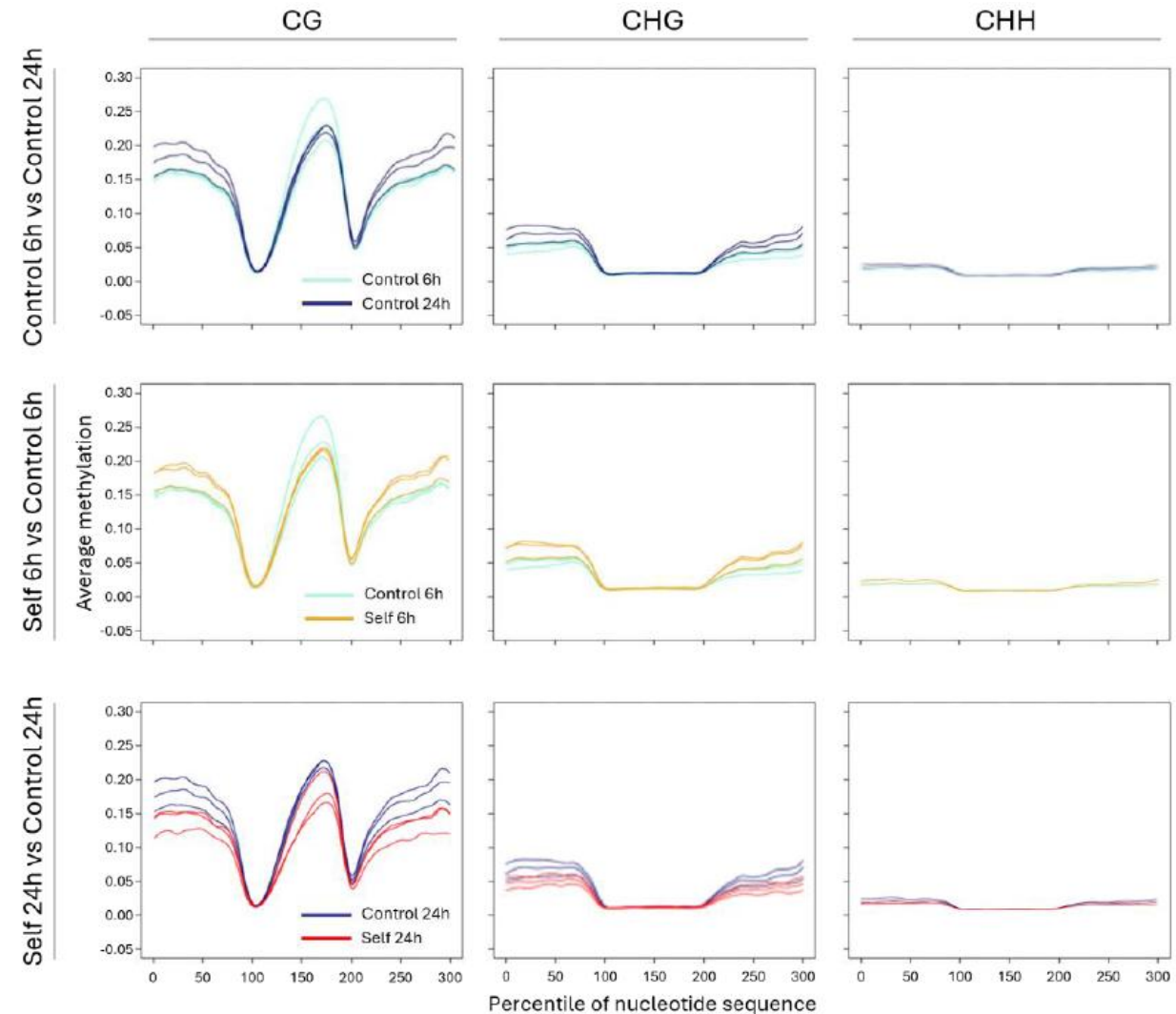
Article

Arabidopsis thaliana Roots Exposed to Extracellular Self-DNA: Evidence of Epigenetic Effects

Alessia Ronchi ^{1,2}, Guido Incerti ^{1,*}, Emanuele De Paoli ¹, Speranza Claudia Panico ¹, Giovanni Luca Sciabbarrasi ¹, Pasquale Termolino ³, Fabrizio Carteni ⁴, Mariachiara Langella ⁴, Maria Luisa Chiusano ⁴ and Stefano Mazzoleni ^{4,*}

Epigenomes 2025, 9, 13

General description	GO term	6h		6-24h		24h	
		E	# DEGs	E	# DEGs	E	# DEGs
Response to stress	response to stimulus		33				55
	response to stress		26				37
	response to abiotic stimulus		20				
	response to chemical		18				
	response to heat		18		7		30
	extracellular region						23
	response to toxic substance						9
	detoxification						8
	cellular response to decreased oxygen levels		8		4		
Response to stress	response to stress		12				
	response to chemical		11				
	cellular response to chemical stimulus		9				
	cellular response to decreased oxygen levels		6				
	cellular response to extracellular stimulus						7
	cellular response to iron ion starvation						4
Transport	zinc ion transmembrane transport		2		2		18
	transition metal ion transmembrane transporter				2		11
	transmembrane transport						8
	inorganic ion transmembrane transport						7
	metal ion transport						6
Photosynthesis associated	plastid envelope						23
	thylakoid						18
	photosynthesis						14
	generation of precursor metabolites and energy						13
	photosynthesis, light reaction						11
	pigment biosynthetic process						6
photosynthetic electron transport chain						5	



Debate and supporting evidence by other teams

A Review: Is Cinderella's story of self-DNA extracellular effect towards plant growth real?

R Purnamasari¹, U Sudadi² and D A Santosa^{2,3*}

¹Soil Science Study Program, Graduate School, IPB University


²Department of Soil Science and Land Resource, Faculty of Agriculture, IPB University

³Biotechnology Center, Institute for Research and Community Empowerment, IPB University

Abstract. Research related to the extracellular self-DNA effect in plants has been widely conducted during the last decade. Researchers reported the impact of extracellular self-DNA inhibition on plant growth, assuming that extracellular self-DNA could enter plant tissue and thus stimulate an inhibitory response. Environmental conditions have a big role to play in supporting this inhibition, especially with the significant climate changes that have occurred in recent years. Climate changes such as rainfall, humidity, light intensity, and air temperature have a positive effect on the decomposition process of plant litter. Especially if there is a significant climate change accompanied by a monoculture cropping pattern will further trigger the accumulation of extracellular DNA. Dissolved DNA concentration in the soil with the probability of occurrence of the self-DNA effect is directly proportional. Although several studies have confirmed the inhibitory results, self-DNA's mechanism can enter plant tissue and stimulate an inhibitory response has not been widely discussed. It has sparked debate regarding the actual effect of self-DNA and the extent of its potential to inhibit plant growth. Therefore, this paper intends to collect various hypotheses and research results regarding self-DNA's impact on plant growth and reconstruct more comprehensive assumptions regarding the effect of self-DNA, its inhibitory potential, and its ecological implications in agriculture. Exhaustive assumptions about self-DNA are expected to be the basis for generating new research designs that include new variables that have not been considered in previous research.

Review

Extracellular DNA: Insight of a Signal Molecule in Crop Protection




Ileri Alejandra Carbajal-Valenzuela ¹, Gabriela Medina-Ramos ^{2,*}, Laura Helena Caicedo-Lopez ¹ , Alejandra Jiménez-Hernández ¹, Adrian Esteban Ortega-Torres ¹, Luis Miguel Contreras-Medina ¹, Irineo Torres-Pacheco ¹ and Ramón Gerardo Guevara-González ^{1,*}

- ¹ C. A. Biosystems Engineering, Campus Amazcala, Autonomous University of Queretaro, Carr. Chichimequillas-Amazcala Km 1 S/N, C.P., El Marques, Querétaro 76265, Mexico; jancarval@hotmail.com (I.A.C.-V.); inglauraclo23@gmail.com (L.H.C.-L.); ale.jhtsu@gmail.com (A.J.-H.); adrianesotorres@gmail.com (A.E.O.-T.); mcontreras.uaq@gmail.com (L.M.C.-M.); irineo.torres@uaq.mx (I.T.-P.)
- ² Molecular Plant Pathology Laboratory, Polytechnic University of Guanajuato, Cortazar 38496, Mexico
- * Correspondence: gmedina@upgto.edu.mx (G.M.-R.); ramongg66@gmail.com or ramon.guevara@uaq.mx (R.G.G.-G.); Tel.: +52-1-461-441-4300 (G.M.-R.); +52-1-442-192-1200 (ext. 6093) (R.G.G.-G.)

Biology 2021, 10, 1022.

Article

Extracellular DNA as a Strategy to Manage Vascular Wilt Caused by *Fusarium oxysporum* in Tomato (*Solanum lycopersicum* L.) Based on Its Action as a Damage-Associated Molecular Pattern (DAMP) or Pathogen-Associated Molecular Pattern (PAMP)

Alejandra Jiménez-Hernández ^{1,†}, Ileri Alejandra Carbajal-Valenzuela ^{1,†}, Irineo Torres-Pacheco ¹ , Enrique Rico-García ¹, Rosalía V. Ocampo-Velazquez ¹, Ana Angélica Feregrino-Pérez ²  and Ramón Gerardo Guevara-González ^{1,*} 

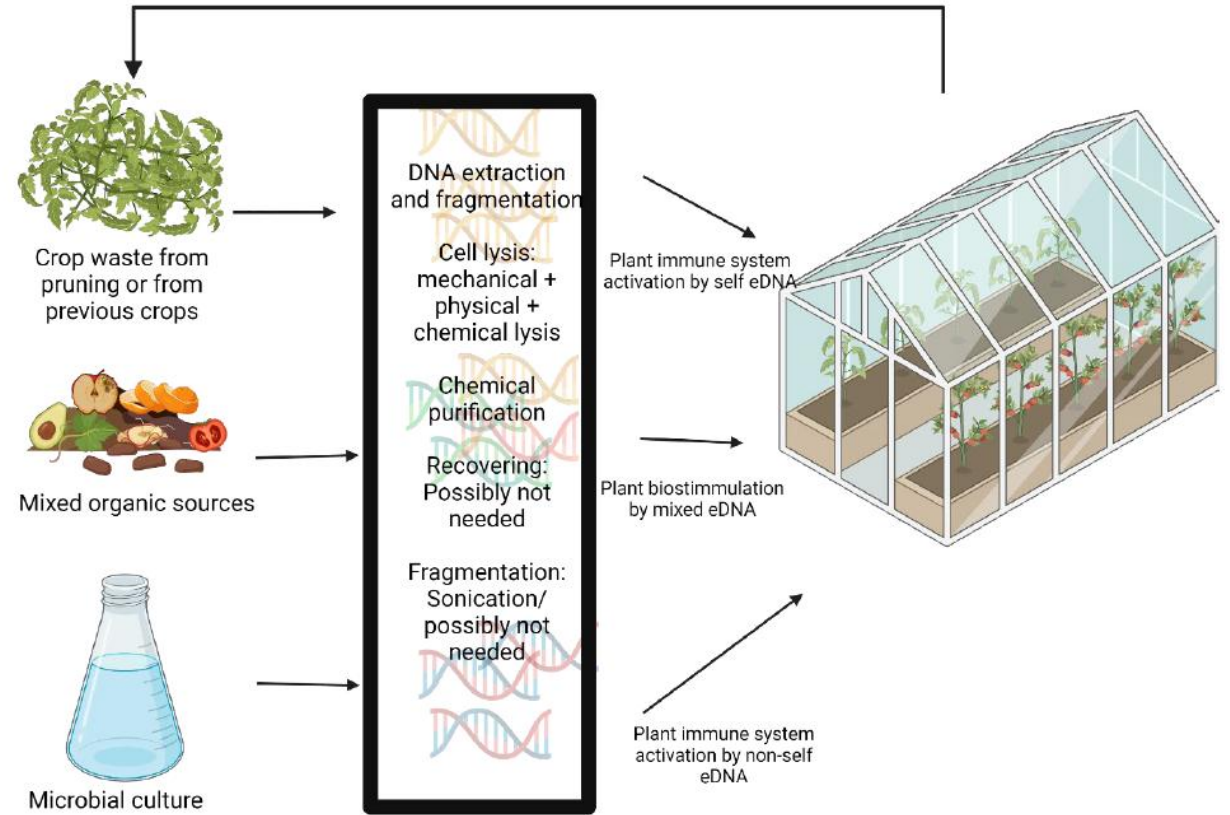


Figure 1. A proposed system of eDNA as an agriculture treatment with circular economy agricultural eDNA treatments.

Extracellular Fragmented Self-DNA Is Involved in Plant Responses to Biotic Stress

Francesca Barbero¹, Michela Guglielmotto², Monirul Islam^{3†} and Massimo E. Maffei^{3*}

¹ Department of Life Sciences and Systems Biology, University of Turin, Turin, Italy, ² Neuroscience Institute of Cavalieri Ottolenghi Foundation, University of Turin, Turin, Italy, ³ Plant Physiology Unit, Department of Life Sciences and Systems Biology, University of Turin, Turin, Italy

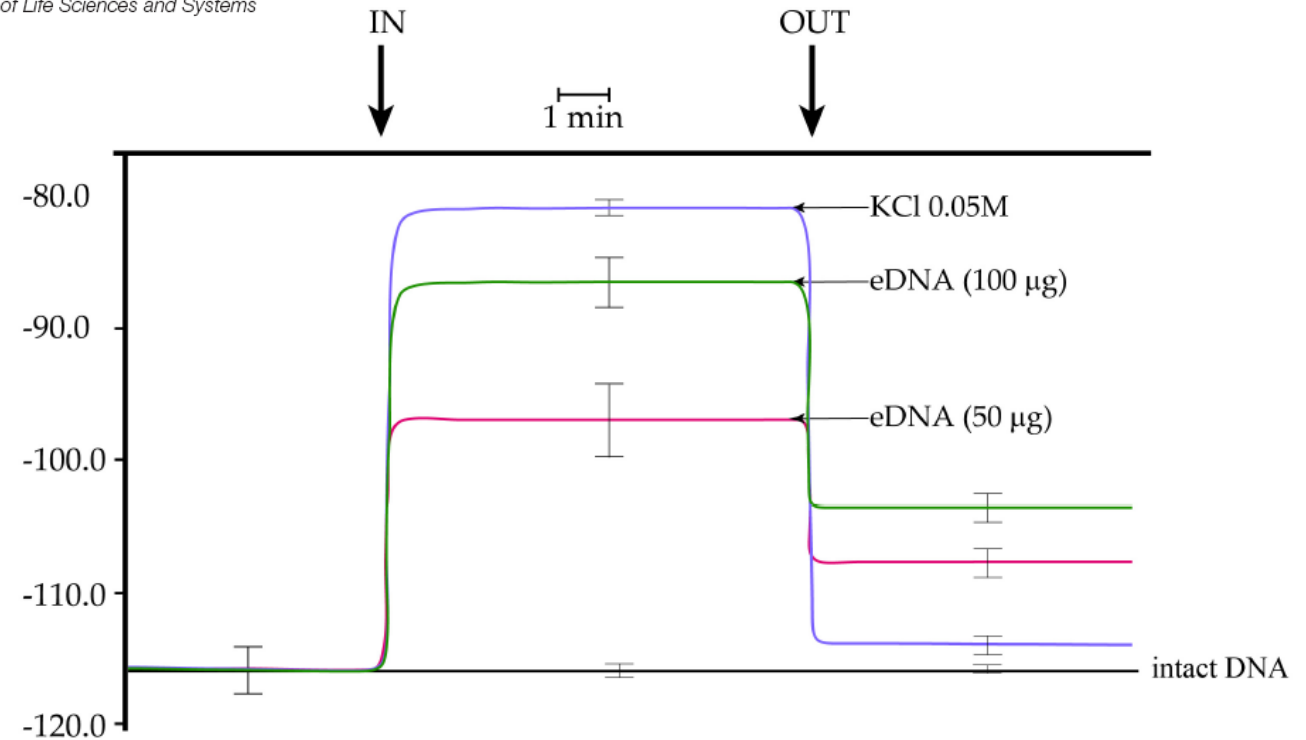


International Journal of
Molecular Sciences

Article

Extracellular Self-DNA (esDNA), but Not Heterologous Plant or Insect DNA (etDNA), Induces Plasma Membrane Depolarization and Calcium Signaling in Lima Bean (*Phaseolus lunatus*) and Maize (*Zea mays*)

Francesca Barbero¹, Michela Guglielmotto², Andrea Capuzzo³ and Massimo E. Maffei^{1,*}



Extracellular self-DNA induces local inhibition of growth, regulates production of reactive oxygen species, and gene expression in rice roots

T. O. S. TJIA¹ , K. MEITHA^{1,*} , P. SEPTIANI¹ , R. AWALUDIN¹ , and D. SUMARDI¹ 

School of Life Sciences and Technology, Bandung Institute of Technology, Bandung, 40116, Indonesia

<https://doi.org/10.1093/plphys/kiad252>

PLANT PHYSIOLOGY 2023: 192: 2233–2234

Plant Physiology®

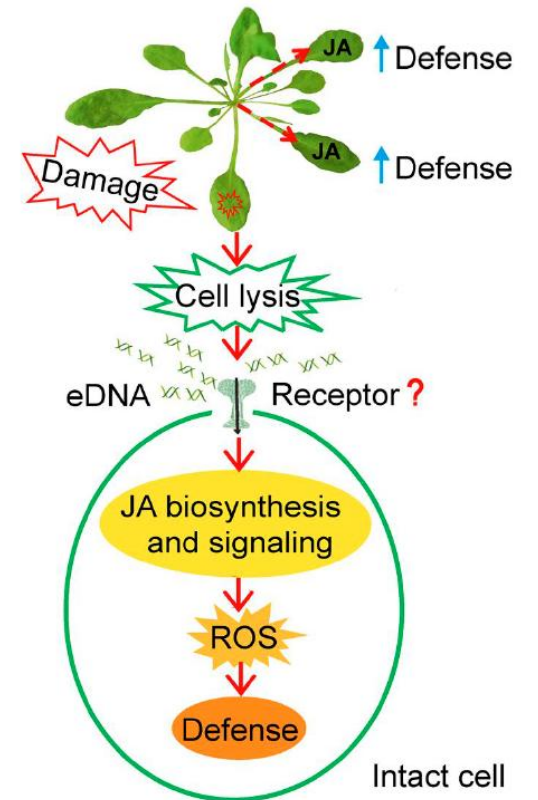
It's not you, it's me: Extracellular self-DNA signals through the jasmonic acid pathway

Guadalupe L. Fernández-Milmanda ^{1,2,3,*}

¹ Assistant Features Editor, Plant Physiology, American Society of Plant Biologists, USA

² Department of Plant Biotechnology and Bioinformatics, Ghent University, 9052 Ghent, Belgium

³ VIB, Center for Plant Systems Biology, 9052 Ghent, Belgium





Contents lists available at [ScienceDirect](https://www.sciencedirect.com)

Postharvest Biology and Technology

journal homepage: www.elsevier.com/locate/postharvbio



Extracellular self-DNA induced a PTI-related local defence against *Rhizopus* rot in postharvest peach fruit

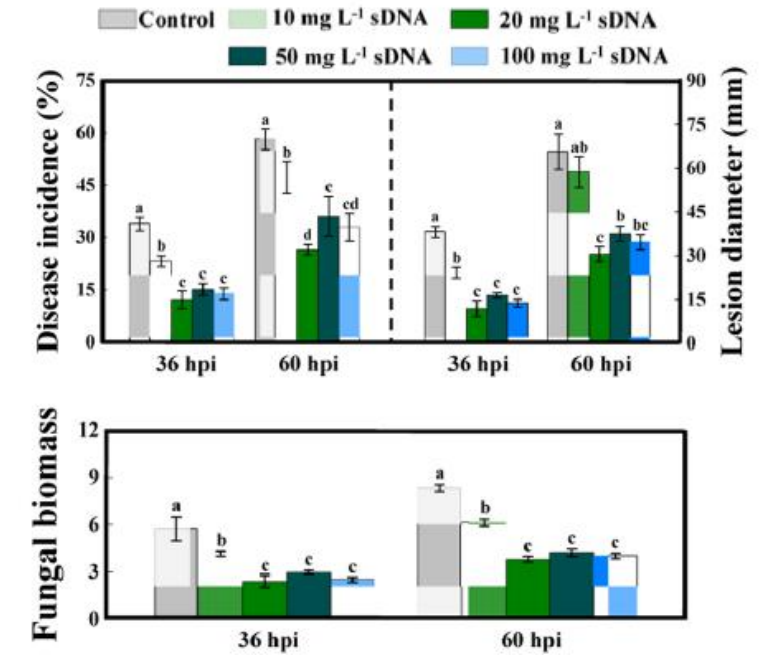
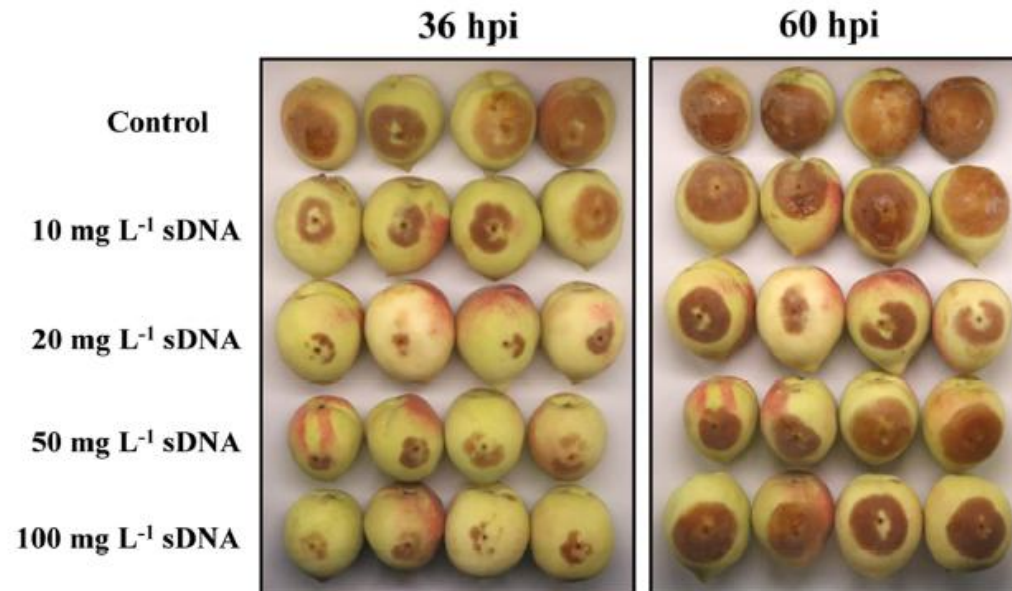
Chunhong Li^{a,b,c}, Kaituo Wang^{a,b,c,*}, Yanyu Zou^{a,b,c}, Changyi Lei^b, Zongxiang Chen^d, Yonghua Zheng^c

^a Laboratory of Fruit Function and Disease Management, Department of Public Health and Management, Chongqing Three Gorges Medical College, Chongqing 404000, China

^b College of Biology and Food Engineering, Chongqing Three Gorges University, Chongqing 404000, China

^c College of Food Science and Technology, Nanjing Agricultural University, Nanjing 210095 Jiangsu, China

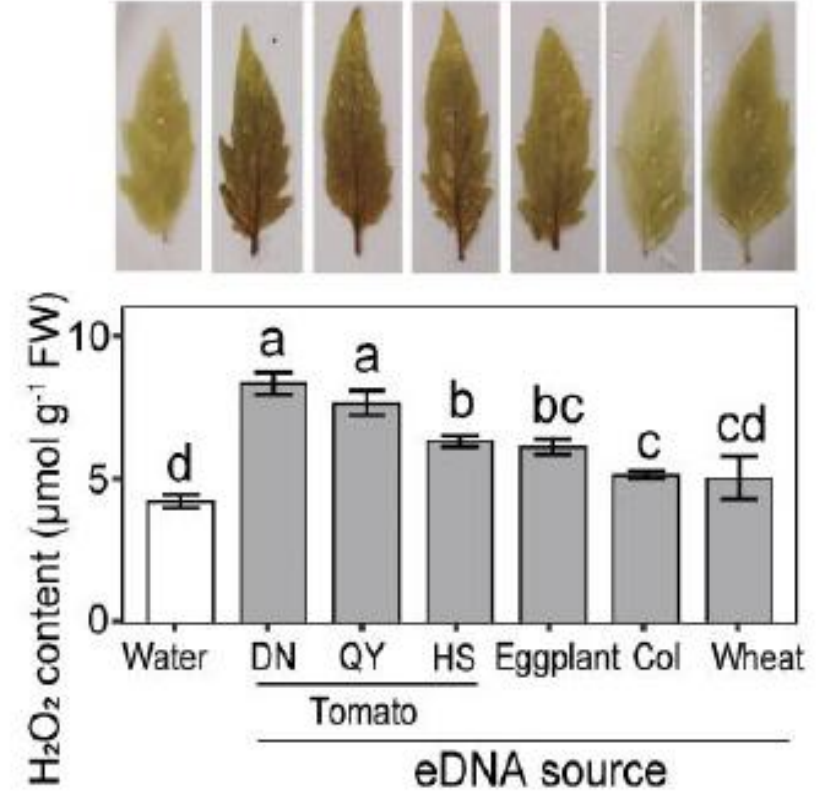
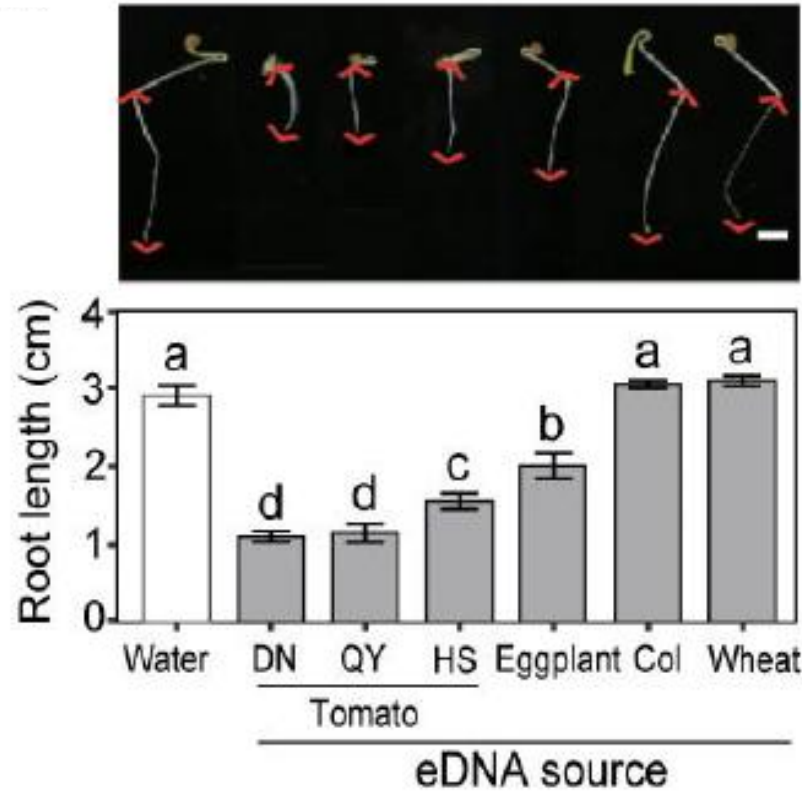
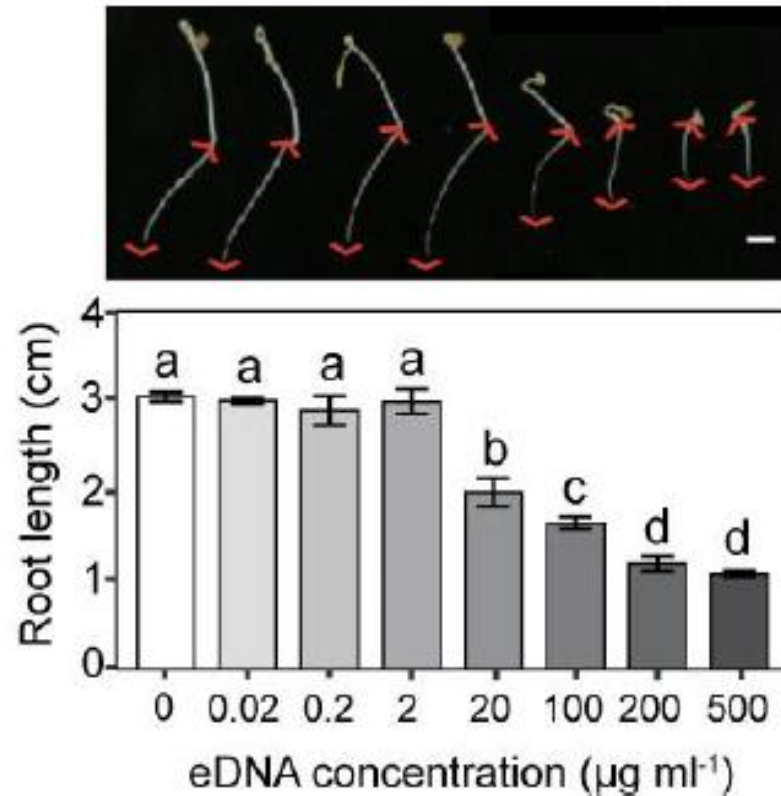
^d Chongqing Customs Technology Center, Chongqing Customs, Chongqing 401147, China





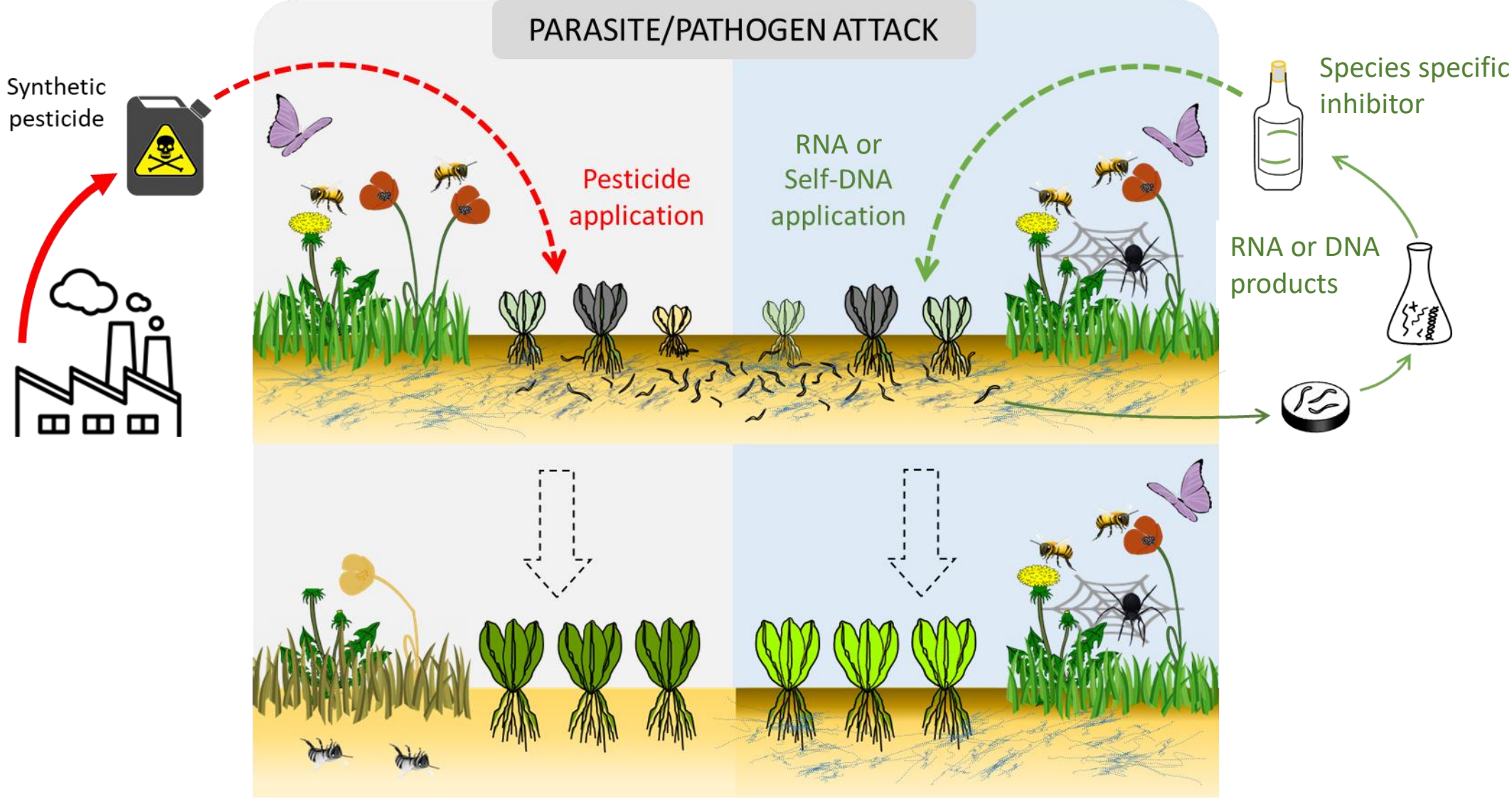
Plant extracellular self-DNA inhibits growth and induces immunity via the jasmonate signaling pathway

Xingang Zhou ,¹ Huan Gao ,^{1,2} Xianhong Zhang ,¹ Muhammad Khashi u Rahman ,¹ Stefano Mazzoleni ,³ Minmin Du * and Fenzhi Wu 1,*



NUOVO SCENARIO PER IL BIOCONTROLLO IN AGRICOLTURA: Acidi nucleici?

Past versus Future





Key Mechanistic Principles and Considerations Concerning RNA Interference

Petr Svoboda*

Institute of Molecular Genetics of the Czech Academy of Sciences, Prague, Czechia

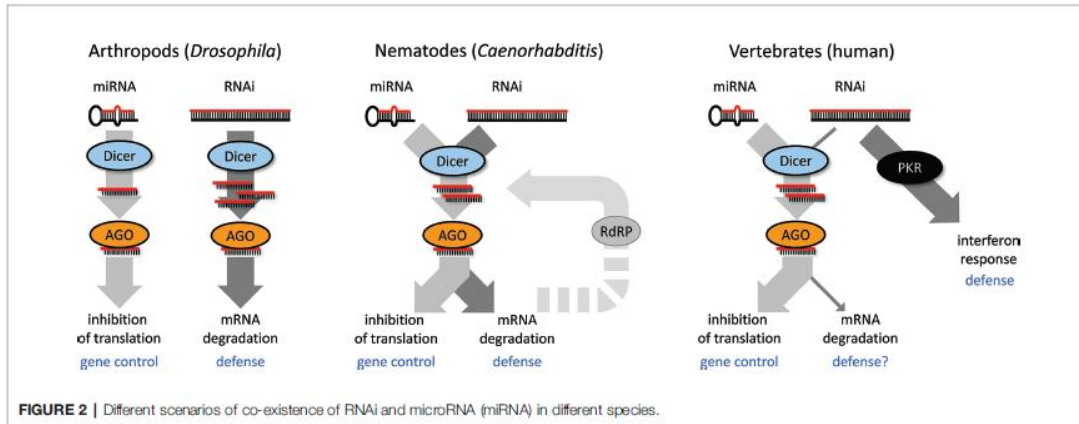


FIGURE 2 | Different scenarios of co-existence of RNAi and microRNA (miRNA) in different species.

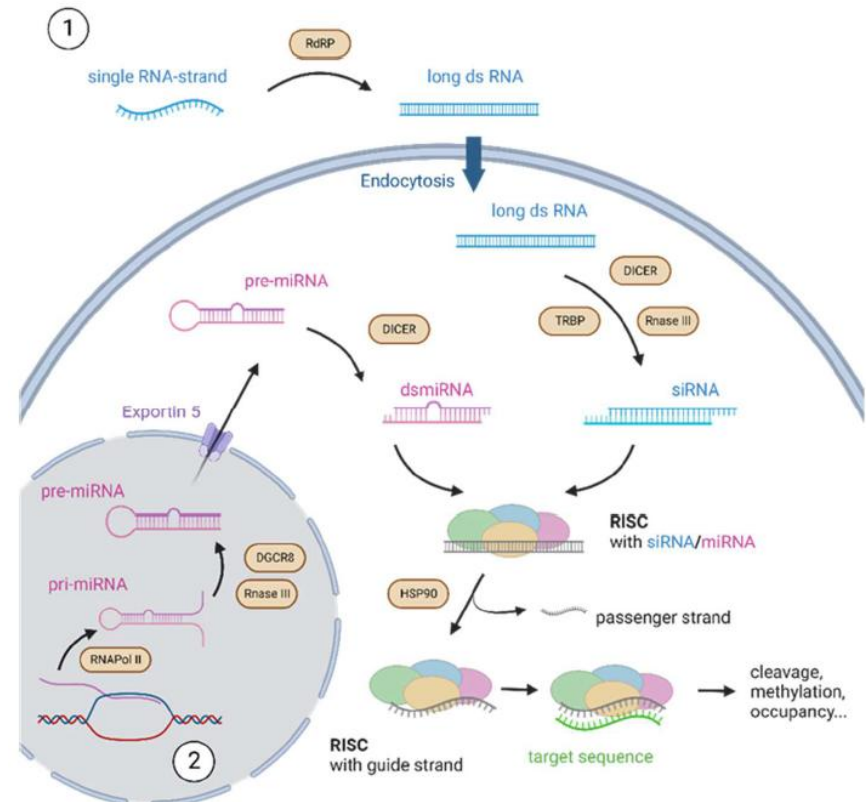
Review

Basic Principles of RNA Interference: Nucleic Acid Types and In Vitro Intracellular Delivery Methods

Marie Isenmann^{1,2}, Martin James Stoddart^{1,2}, Rainer Schmelzeisen¹, Christian Gross¹, Elena Della Bella^{2,†} and René Marcel Rothweiler^{1,2,*}

¹ Department of Oral and Maxillofacial Surgery, Faculty of Medicine, University of Freiburg, Hugstetterstrasse 55, 79106 Freiburg, Germany

² AO Research Institute Davos, Clavadelstrasse 8, 7270 Davos, Switzerland



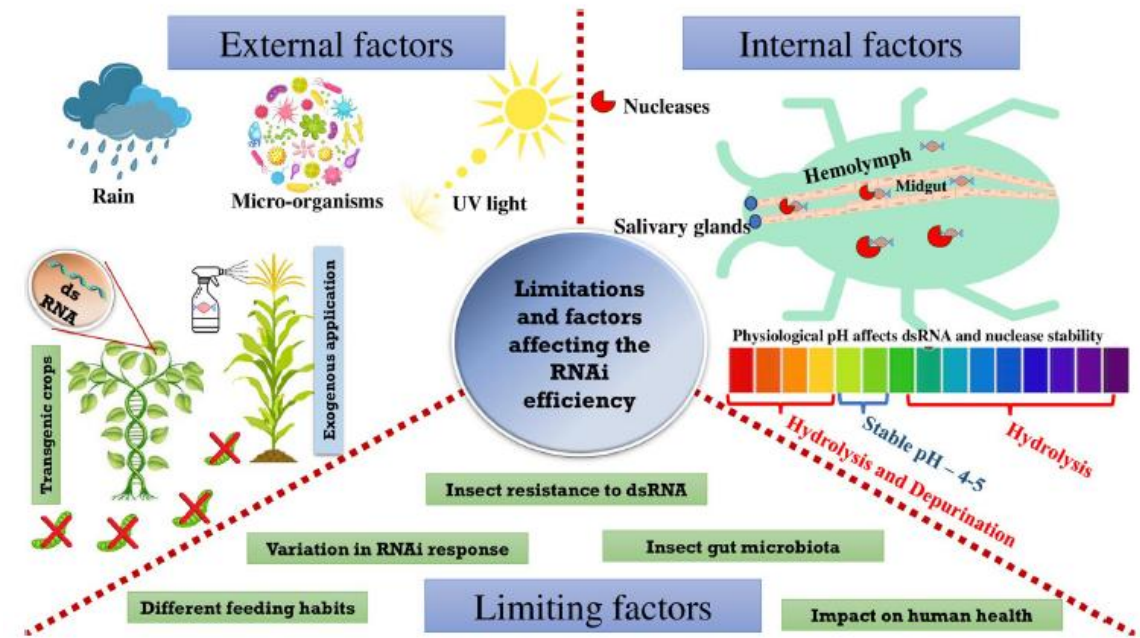
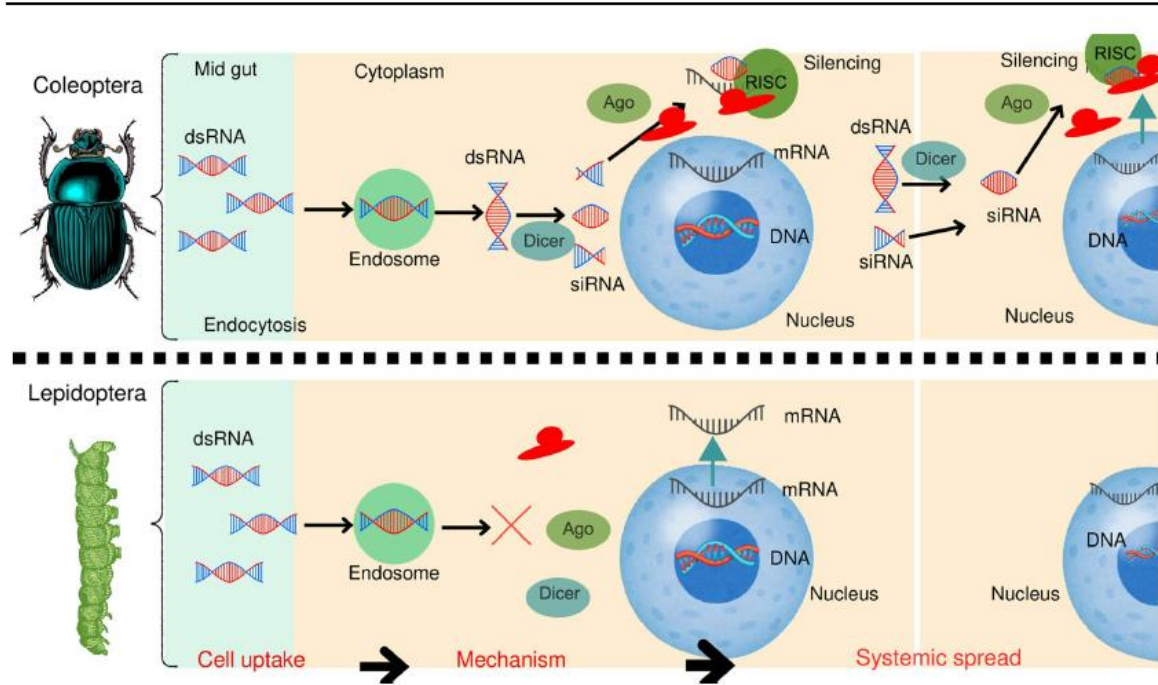


RNA interference (RNAi) for insect pest management: understanding mechanisms, strategies, challenges and future prospects

Deepak Kumar Mahanta¹ · J. Komal² · Tanmaya Kumar Bhoi³ · Ipsita Samal⁴ · Sangeeta Dash⁵ · Sumit Jangra⁶

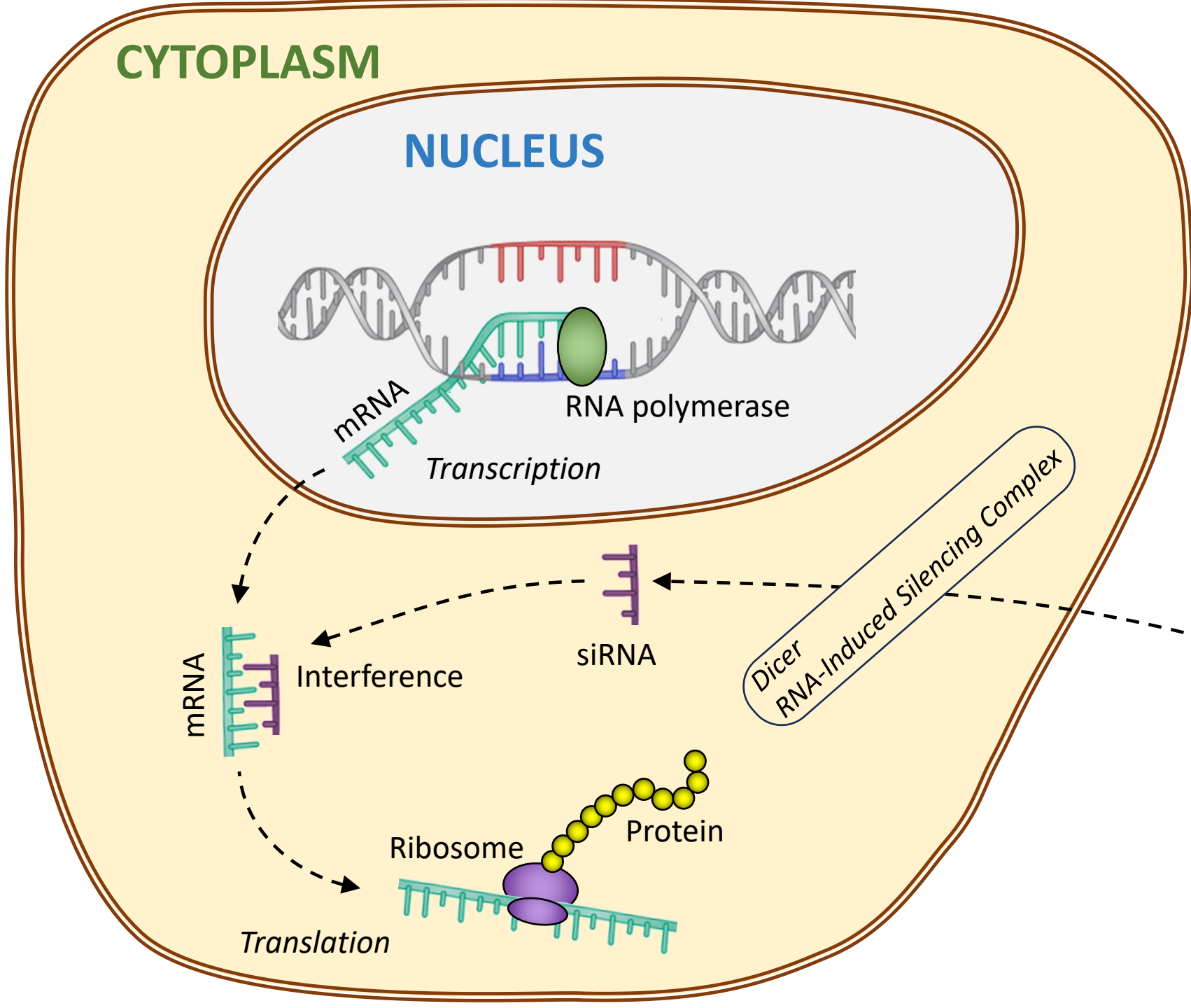
Forest Entomology Discipline, Forest Protection Division,
Indian Council of Forestry Research and Education

Biologia Futura



RNA interference

(Single gene target)



CYTOPLASM

NUCLEUS

mRNA
RNA polymerase
Transcription

mRNA
Interference
siRNA
Ribosome
Protein
Translation

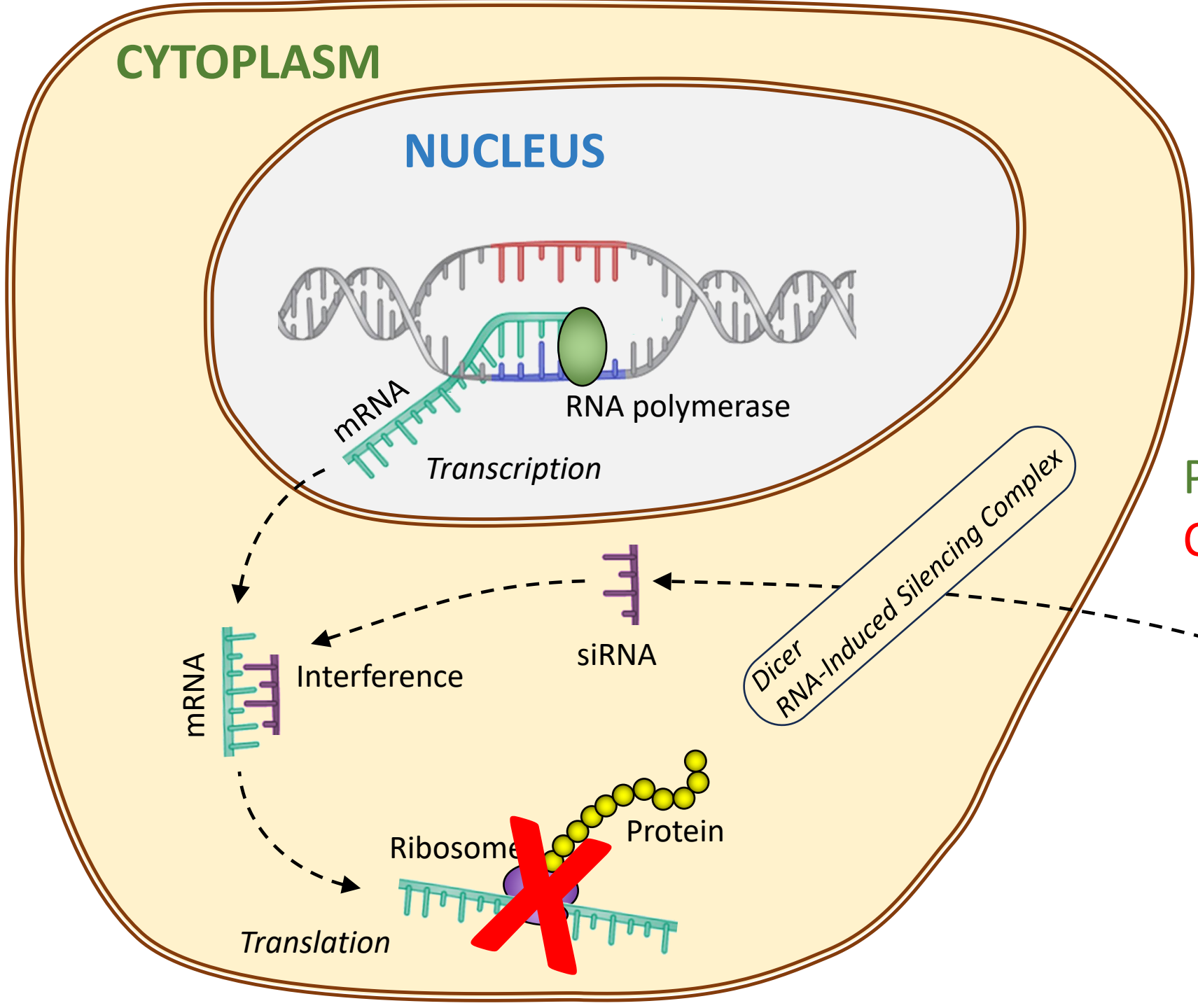
Dicer
RNA-Induced Silencing Complex

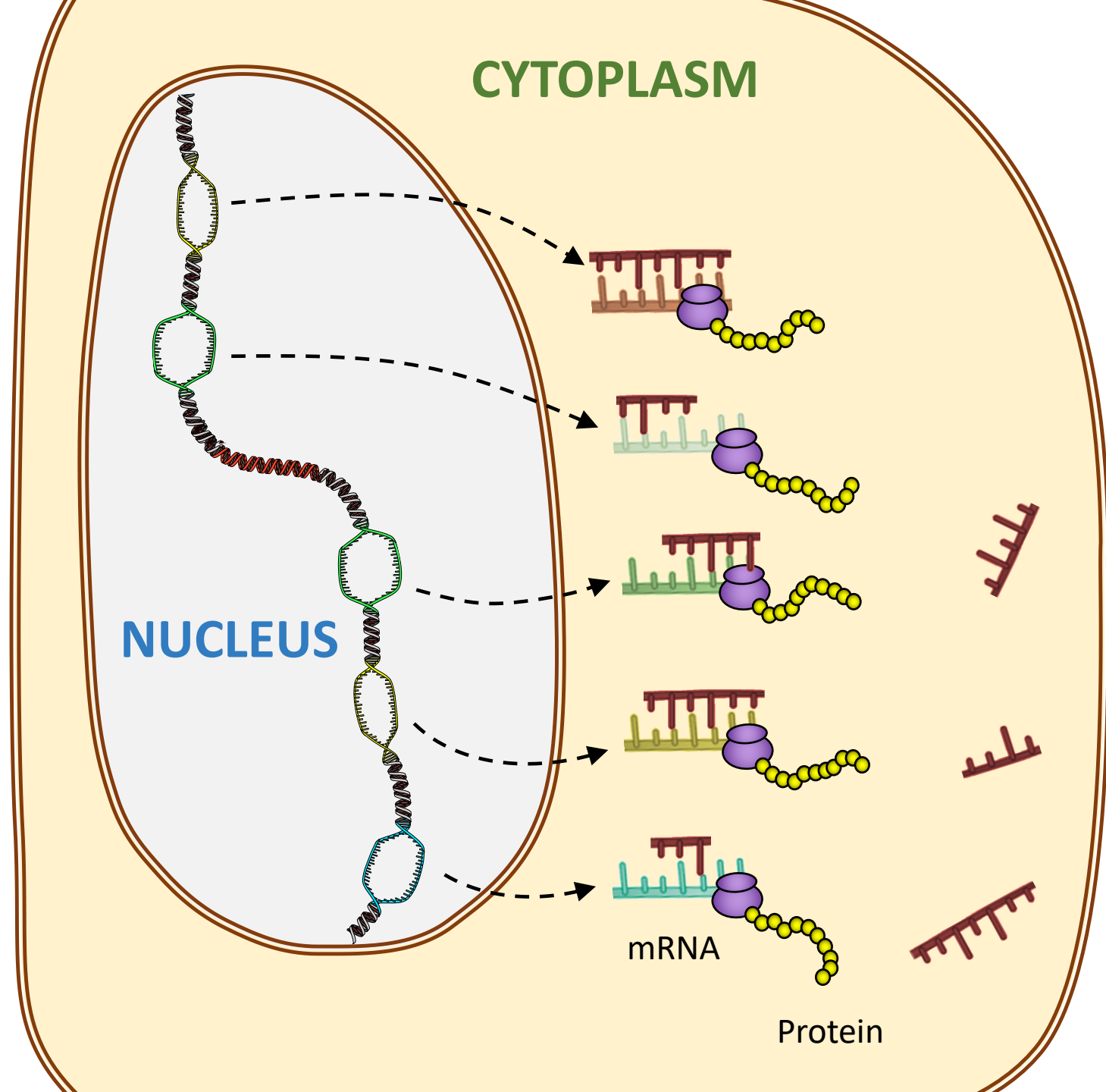
dsRNA

RNA interference

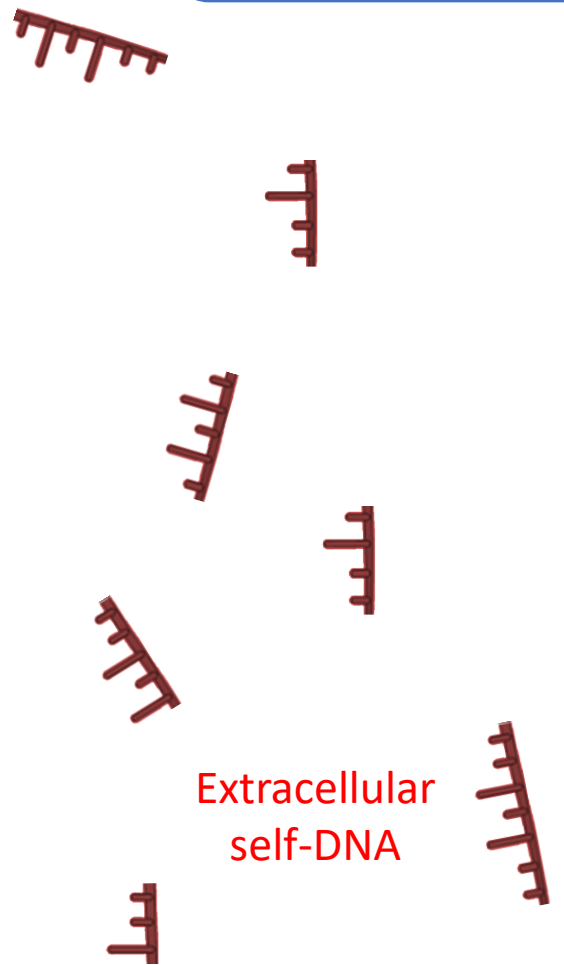
(Single gene target)

Pro: high precision
Con: induction of resistance

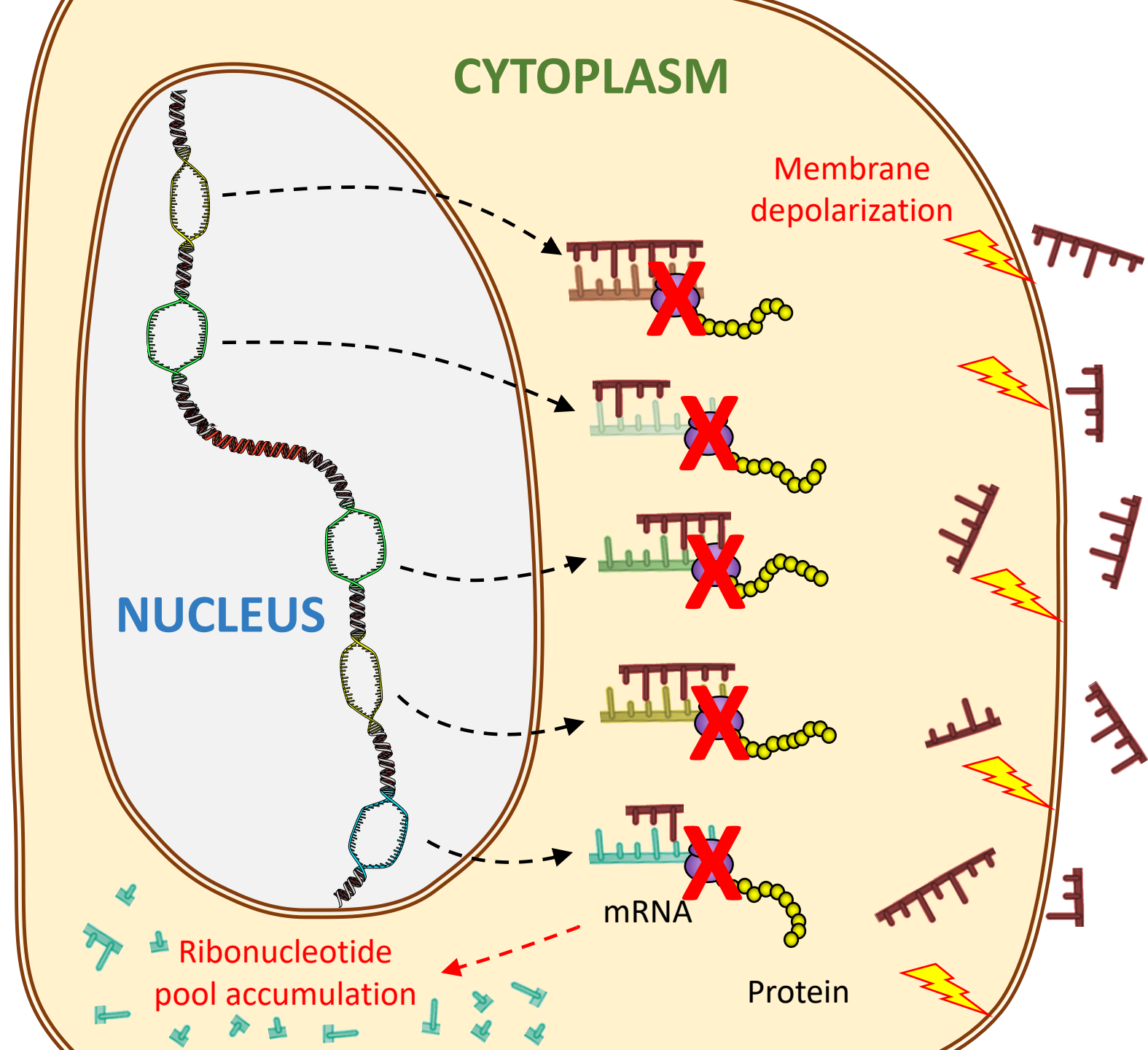




self-DNA inhibition
(generalized targets)



self-DNA inhibition
(generalized targets)



Pro: generalised action
(no resistance)

Con: ?

Uso di acidi nucleici possibile:

- Vantaggi del DNA rispetto al RNA (*AND technology*)
- Problematiche regolatorie

Microrganismi benefici → Biostimolanti



Spirulina (Arthrospira platensis)



Trichoderma harzianum



syngenta
Italia

SC01-SEED

ULTIMO AGGIORNAMENTO: 03.06.2025

BIOSTIMOLANTI-SPECIALITÀ NUTRIZIONALI, CONCANTI
NUTRIZIONE DELLE COLTURE

Composizione:
8% Azoto (N) organico

Formulazione:
Polvere disperdibile

→ Biocontrollo

Bacillus thuringiensis



Beauveria bassiana



Microrganismi benefici

Biostimolanti

Biocontrollo

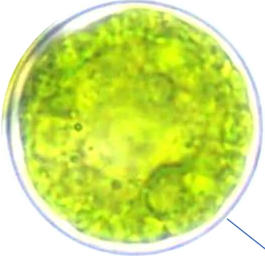
+ DNA_{*i*} → Biocontrollo + efficace su specie *i*

«AND Biologicals» technology

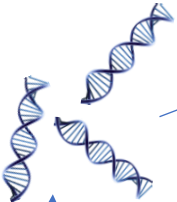
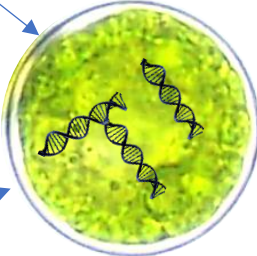


AND technology

Microalgae cell culture



Natural uptake



DNA from Pathogen/Parasite/Weed



Lab Scale-Up



Industrial Production

Biocontrol



Natural carriers
Biomass



AND product



Biocontrol

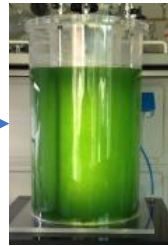
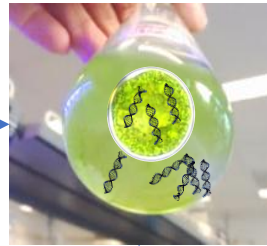
Fusariosi lattuga – esempio di applicazione AND



Fusariosi Lattuga

biocontrol by *Fusarium self-DNA*

Arthrospira platensis
(Spirulina)
starting culture



NAT

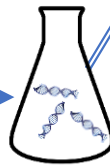


Control

Fusarium oxysporum
culture



Fungal biomass
decomposition



Fusarium DNA

Microalgae
uptake

Scale-up production
in photo-bioreactors

Noself product

Lettuce planting



Foliar application



Fusariosi Lattuga

biocontrol by *Fusarium self-DNA*



Control



NAT

CONCLUSIONI

Nuova funzione del DNA ambientale

Importanza del bilancio self/noself DNA

