

Journal of Materials Chemistry A

Materials for energy and sustainability

Accepted Manuscript

This article can be cited before page numbers have been issued, to do this please use: R. Umapathi, H. Kim, D. Kim, A. Gaur, V. Chaudhary, M. Safarkhani, S. Shin, T. J. Park, G. Mohana Rani, K. Shin and Y. S. Huh, *J. Mater. Chem. A*, 2026, DOI: 10.1039/D5TA09692H.



This is an Accepted Manuscript, which has been through the Royal Society of Chemistry peer review process and has been accepted for publication.

Accepted Manuscripts are published online shortly after acceptance, before technical editing, formatting and proof reading. Using this free service, authors can make their results available to the community, in citable form, before we publish the edited article. We will replace this Accepted Manuscript with the edited and formatted Advance Article as soon as it is available.

You can find more information about Accepted Manuscripts in the [Information for Authors](#).

Please note that technical editing may introduce minor changes to the text and/or graphics, which may alter content. The journal's standard [Terms & Conditions](#) and the [Ethical guidelines](#) still apply. In no event shall the Royal Society of Chemistry be held responsible for any errors or omissions in this Accepted Manuscript or any consequences arising from the use of any information it contains.

Innovative Nanozymes: Emerging Platforms for Efficient Pesticide Detection

Reddicherla Umapathi^{1,a}, Hanseung Kim^{1,a}, Donghyeon Kim¹, Ashish Gaur,² Vishal Chaudhary,³ Moein Safarkhani⁴, Sujeong Shin⁵, Tae Jung Park⁶, Gokana Mohana Rani^{4,*}, Kwangsoo Shin^{7,*}, Yun Suk Huh^{1,*}

¹NanoBio High-Tech Materials Research Center, Department of Biological Sciences and Bioengineering, Inha University, Incheon 22212, Republic of Korea

²Department of Energy Science, Sungkyunkwan University, Suwon, 16419 Republic of Korea

³Centre for Theoretical Physics and Natural Philosophy, Nakhonsawan Studiorum for Advanced Studies, Mahidol University, Nakhonsawan 60130, Thailand

⁴Department of Energy and Materials Engineering, Dongguk University-Seoul, Seoul, 04620, Republic of Korea

⁵New Hazardous Substances Team, Department of Food Safety Evaluation, National Institute of Food and Drug Safety Evaluation, Ministry of Food and Drug Safety, Cheongju-si 28159, Korea

⁶Department of Chemistry, Research Institute of Chem-Bio Diagnostic Technology, Chung-Ang University, 84 Heukseok-ro, Dongjak-gu, Seoul 06974, Republic of Korea

⁷Department of Polymer Science and Engineering and Program in Environmental and Polymer Engineering, Inha University, Incheon 22212, Republic of Korea

*Corresponding authors:

Prof. Yun Suk Huh (yunsuk.huh@inha.ac.kr)

Prof. Kwangsoo Shin (kwangsoo.shin@inha.ac.kr)

Dr. Gokana Mohana Rani (gmohanarani@gmail.com)

^aEqually contributed authors (RU and HK)



1 Abstract

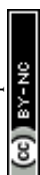
2 Effective and timely detection of pesticide residues is highly crucial as these toxic
3 chemical residues cause several effects on humans and environment. Increasing necessity
4 towards progression of detection systems has flashed considerable curiosity in analytical
5 sensing technologies. Substantial research advancements have been made in analytical
6 chemistry, nanozymes, and materials chemistry, for detection of pesticides. A relatively
7 innovative sensing strategy that has gained widespread attention is based on natural enzyme
8 mimicking catalytic property of nanomaterials, commonly referred to as nanozyme activity.
9 Nanozymes properties make them highly promising sensing materials for precisely
10 determining the pesticides with low detection limits, wide linear ranges, excellent selectivity,
11 and appreciable sensitivity. Apart from increasing the sensing behaviors, these characteristics
12 will facilitate in designing versatile sensing platforms that integrate surface enhanced Raman
13 scattering (SERS), colorimetric, electrochemical, and fluorescence transduction procedures. In
14 this review, we provided a critical overview on nanozymes for pesticides detection,
15 encompassing their material compositions, strategies for activity enhancement, output signals,
16 and methodologies. Explicitly discussed the advances made in nanozyme-based colorimetric,
17 electrochemical, SERS, and fluorescence sensing strategies for the substantial quantification
18 of pesticides. Outlined the strategic advantages and detailed mechanisms for the detection of
19 pesticides. Highlighted the working principle and salient features of various nanozyme-based
20 sensors, to enhance the sensing efficiency. Finally, challenges and associated limitations were
21 discussed to realize full potential of nanozyme-based optical and electrochemical sensors.

22 **Keywords:** Nanozymes, Pesticides, Colorimetric Sensors, Fluorescence Sensors,
23 Electrochemical Sensors, SERS Sensors, Emerging Contaminants.



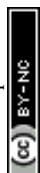
1 1. Introduction

2 Pesticides denote a broad category of chemical substances utilized chiefly for the
3 management of weeds, pathogens, pests, and diseases in livestock production, and forestry.
4 Pesticides can be classified as rodenticides, acaricides, insecticides, plant growth regulators,
5 fungicides, and herbicides depending on their desired application. Over the past few years,
6 ongoing population expansion and rising demand for food products have rendered the
7 importance of pesticides in aquaculture, animal husbandry, agriculture, and the protection of
8 industrial items from pests and molds. All these instances lead to an increase in the
9 consumption of pesticides.¹⁻⁴ The essential application of pesticides for improving agricultural
10 output has become unavoidable. The excessive consumption, improper utilization, and abuse
11 of pesticides, along with their exponential expansion in use, has led to significant pesticide
12 residues that threaten human well-being and ecosystems. Unfortunately, the indiscriminate and
13 unregulated use of these pesticides transcends the designated target areas, with pesticide
14 residues persisting for months or even years. Pesticides released into the surrounding
15 atmosphere permeate soil and aquatic ecosystems through air circulation, where they are
16 frequently absorbed and accumulated by the root systems and medicinal plants, ultimately
17 accumulating in the bodies of humans through the food chain.⁵⁻¹⁰ The lack of precise
18 information has incited extensive insect infestations, presenting considerable health hazards to
19 people and other organisms because of residual pesticides in food. The existence of even
20 minimal quantities of these residual pesticides has become a significant obstacle to
21 international trading in the food commodities. Prolonged exposure of pesticides will result in
22 reproductive toxicity, immunotoxicity, neurotoxicity, hepatotoxicity, and severe epithelial
23 toxicity, moreover, in extreme cases it will lead to the development of cardiovascular diseases,
24 cancer, fetal malformations, and atherosclerosis. Younger populations will be more prone to

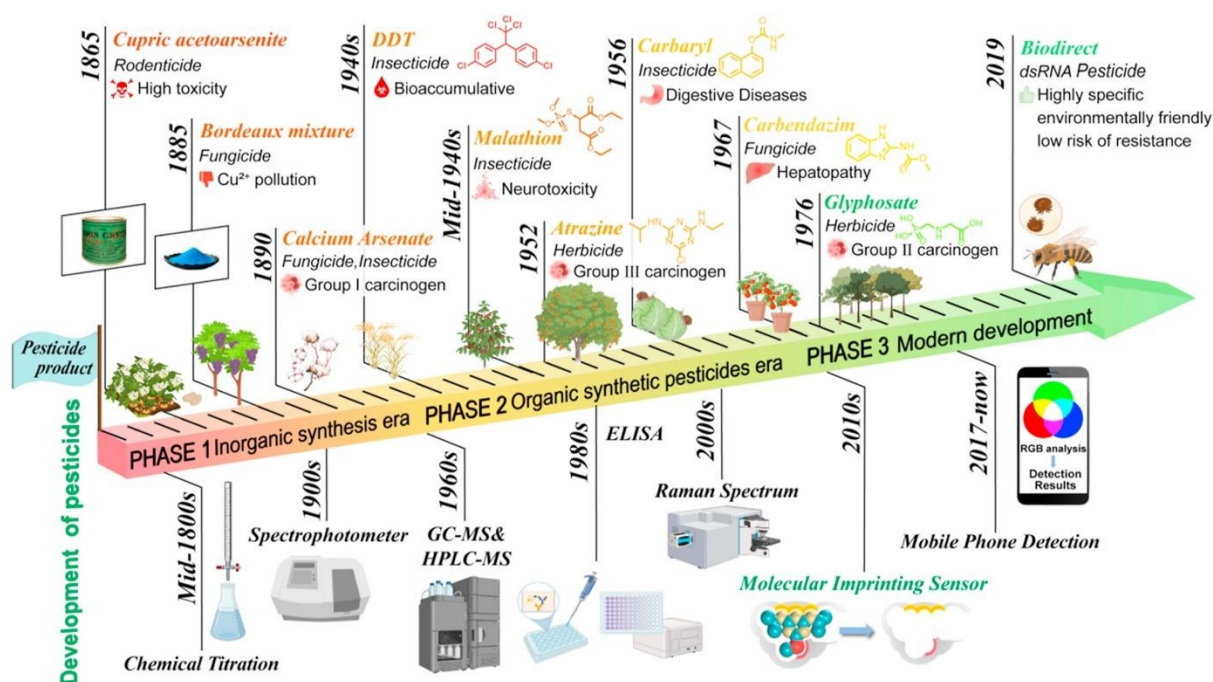


1 these toxic consequences. As a result, pesticide residues have emerged as a major potential
2 hazard to human health. In response to this issue, many nations and globally recognized
3 organizations have set maximum residue limits for pesticides across different matrices and
4 have enacted prompt restrictions and bans on high-risk pesticides, progressively substituting
5 them with low-toxicity synthetic substitutes. Modern medicine encounters intrinsic constraints
6 in therapy due to the emergence of novel diseases caused by the pesticides. Hence, it is
7 imperative to limit the usage and spraying of pesticides to the plants and to timely quantify the
8 pesticides in the surrounding environment.¹¹⁻¹⁵

9
10 Conventional pesticide quantification techniques such as mass spectrometry (MS), high-
11 performance liquid chromatography (HPLC), gas chromatography (GC), and their combined
12 methods HPLC-MS, and GC-MS requires pumps with high pressures to enable interactions
13 between the analytes and functional components of the mobile and stationary phases,
14 permitting the quantification of target compounds based on the peak area. Moreover, mass
15 spectrometry is utilized to ionize sample molecules, facilitating qualitative investigation of the
16 molecular structures. Despite achieving exceptional sensitivity and accuracy, these systems are
17 constrained by inherent restrictions, such as the need for skilled operators, complex and costly
18 apparatus, intricate pretreatment processes, the requirement for trained staff, and the logistical
19 challenges of on-site detection. Recent literature from the past ten years indicates that the swift
20 advancement of sophisticated sensing systems has emerged as one of the most effective
21 solutions to the issues and challenges posed by conventional pesticide quantification
22 techniques.^{5-10, 14, 15} Currently, intensified demand for straightforward, rapid, sensitive, and
23 selective detection platforms has been noticed. Among these technologies, sensors exhibiting



1 excellent sensitivity and thihg selectivity have been identified as viable platforms for pesticide
 2 residue analysis, owing to their capacity for user-friendly high-throughput, low-cost, and on-
 3 site detection.^{5-10, 14, 15}



5 **Scheme 1** The development history of pesticides and their related detection method.

6 Reproduced from reference⁴ with permission from Elsevier, copyright 2025.

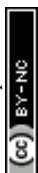
7
 8
 9 For the quantification of the residual pesticides, different types of sensors have been
 10 employed depending on their conversion mechanism including, electrochemical, optical,
 11 acoustic, and thermal sensors, etc. Among these, electrochemical and optical sensors play a
 12 prominent role in detecting contaminants.^{1-11, 14-18} Optical sensors are engineered to respond to
 13 variations in light signals resulting from the interaction between bio-recognition elements and
 14 analytes. Compared to conventional pesticide quantification techniques, optical and
 15 electrochemical sensing methodologies provides a cost effective, fast, and facile approach for



1 the sensitive quantification of the pesticides based on surface-enhanced Raman scattering
2 (SERS), colorimetric, and fluorescence signal variations. Electrochemical (bio)sensors
3 demonstrate high cost-effectiveness due to their simplistic operation, affordability, and
4 efficient in situ detection potential, which makes them particularly valuable for real-time
5 monitoring of pesticide residues. Scheme 1 delineates pesticides developmental trajectory and
6 the advancement of the related sensing technologies.⁴

7
8 Sensors are distinguished by their portability, affordability, and minimal utilization of
9 samples and reagents. A variety of sensors have been designed for the identification of residual
10 pesticides in highly complex matrix systems. Nevertheless, the extensive utilization of the
11 traditional sensors in complex samples has been substantially restricted owing to the intrinsic
12 limitations of recognition elements, including low specificity and inadequate physicochemical
13 stability. For example, sensors based on urease, peroxidase, and acetylcholinesterase (AChE),
14 are prone to the inactivation, this can be due to the deformations in structures caused by reactive
15 oxygen species, pH fluctuations, and elevated temperatures produced during reactions in
16 aquatic or soil environments. Similarly, DNA sensors, microbial sensors, and immunosensors
17 are highly prone to environmental factors, resulting in the degradation of probe. Sensing
18 platforms demonstrate a degree of stability in complicated environments, nevertheless, their
19 ability to select certain pesticides is limited. To address these challenges, nanozymes have been
20 utilized as potential sensing platforms owing to their sensitivity, prospective stability, ease of
21 preparation, surface tunability, and compatibility with the target analytes.^{2, 3, 19-22}

22



1 Unlike enzymes and natural enzymes, nanozymes offer significant potential for practical
2 applications by reducing the overall costs and by enhancing the sensing performance. To date,
3 substantial research efforts has been made in designing various types of nanozymes for the
4 detection of pesticides. The nanozyme nanomaterials with enzyme-mimicking catalytic
5 properties have demonstrated substantial potential with excellent stability under harsh
6 conditions, intrinsic catalytic activity, tunable surface chemistry, and structural diversity. These
7 properties make them highly promising sensing materials for precisely determining the
8 pesticides with low detection limits, wide linear ranges, excellent selectivity, and appreciable
9 sensitivity.^{2, 3, 19-22} Apart from increasing the sensing behaviors, these characteristics will
10 facilitate in designing versatile sensing platforms that integrate SERS, colorimetric,
11 electrochemical, and fluorescence transduction procedures. These systems have substantial
12 advantages including rapid response, portability, low cost, and capability for on-site detection,
13 which are crucial for environmental protection and ensuring food security and food safety.
14 Although recent progress in nanozymes have facilitated excellent pesticide recognition, there
15 remains a lack of comprehensive reviews, predominantly discussing strategic advantages and
16 design principles of nanozyme-based colorimetric, electrochemical, SERS, and fluorescence
17 platforms. This study aims to summarize the advances made in pesticide detection through
18 various transduction approaches, considering significant breakthroughs in advanced methods
19 and existing lack of thorough evaluations. The review particularly emphasizes major pesticide
20 families, including organophosphorus, neonicotinoids, carbamates, fungicides, and herbicides,
21 which are used widely in agriculture and are among the most often detected pesticide residues
22 in food and environmental samples, thereby providing a comprehensive assessment of current
23 detection approaches. Comprehensively investigated nanozymes based colorimetric,
24 fluorescence, SERS, and electrochemical sensors for pesticide detection. Strategic advantages



1 of nanozyme-based detection methods have been explicitly elucidated. Outlined the detailed
2 mechanisms for detection of pesticides. Thoroughly discussed the current limitations and key
3 challenges of each sensing modality and suggested future research trends and perspectives that
4 may accelerate the fabrication of advanced nanozyme-based pesticide sensors. This review
5 aims to offer a holistic overview and comprehensive understanding of state-of-the-art
6 nanozyme-based sensors and directed future initiatives for the systematic fabrication of next
7 generation smart sensing strategies for the on-site pesticide monitoring. The relevant literature
8 was searched systematically using the major scientific databases including “Web of Science”,
9 “Scopus”, and “Google Scholar”. The search was carried out using combinations of keywords
10 such as “nanozymes,” “pesticides,” “colorimetric sensors,” “fluorescence sensors,”
11 “electrochemical sensors,” and “SERS sensors.”

13 **2. An overview of nanozymes**

14 Enzymes are a category of biological macromolecular catalysts that are crucial for both
15 the industrial and biological transformation procedures. Natural enzymes have consistently
16 influenced biosensor technologies owing to their exceptional target selectivity. Nonetheless,
17 the extensive utilization of naturally occurring enzymes have been restricted, particularly in
18 challenging environments, owing to their inherent limitations, such as susceptibility to external
19 factors, low operational stability, elevated costs, limited lifespans, specific requirements for
20 storage, reduced stability under atmospheric conditions, and difficult yet costly production and
21 extraction procedures.^{23, 24} Hence, substantial research attempts have been performed to
22 address these limitations. This has resulted in the design and development of several
23 noteworthy enzyme-mimicking platforms. Among these, particularly nanozymes have



1 received a great deal of interest during the last decade. Substantial progress in nanoscience,
2 protein engineering, and nanozyme engineering has broadened the scope and strategies to
3 address the challenges associated with the natural enzymes.^{25, 26} The identification of
4 oxidoreductase-like and hydrolase-like nanozymes is a notable advancement in
5 nanobiotechnology. Nanozymes, or artificial enzymes, can surpass natural enzymes owing to
6 their versatility, catalytic efficiency, stability in harsh environments, superior shelf-life, ease
7 of modification, enzyme-mimicking capabilities, high stability, cost-effectiveness,
8 biocompatibility, distinctive surface chemistry, tunable surfaces, and robustness. These
9 characteristics allowed them to use in wide array of nanotechnology applications, including
10 food safety, biosensors, environmental protection, antioxidant agents, disease diagnosis, drug
11 delivery, antibacterial agents, imaging and treatment, and tissue engineering. Moreover,
12 nanozymes are potentially using in environmental sciences, chemical engineering, bio-medical
13 field, industrial settings, agricultural sectors, and food safety.²⁰⁻²² Nanozymes are often
14 categorized as nanomaterials exhibiting enzyme-mimicking activity as catalysts. So far, several
15 nanomaterials have been synthesized, comprising but not limited to noble metal
16 nanoparticles, carbon-based nanomaterials, metal oxides, polymetallic nanostructures, and
17 metal-organic frameworks (MOFs), have been studied for enzyme-mimicking activities.^{27, 28}
18 The unique physicochemical characteristics of nanomaterials, which can be customized to
19 fulfill particular application needs, are intricately associated with their enzyme like activities.
20 Enzymatic activity characteristics like substrate multienzyme-like behavior catalytic efficiency,
21 and selectivity, are dependent upon the physicochemical features of nanomaterials, which
22 encompass shape, self-assembly, size, surface modification, and crystallographic state.^{29, 30}
23 Furthermore, temperature and pH are critical parameters that can significantly influence the
24 catalytic function of nanozymes. Nanozymes, as novel materials, have exhibited significant



1 potential in the field of electroanalytical chemistry for the detection of potential environmental
2 contaminants.²⁰⁻²²

3 In addition to electroanalytical performance, the fabrication of nanozyme-based
4 pesticide sensors should be aligned with the sustainability and green chemistry principles.
5 Nanozymes synthesized through green chemistry principles represent a transformative
6 perspective integrating sustainability with the advanced electrocatalytic and nano-catalytic
7 functionalities.³¹ Recent investigations substantially emphasize low-energy and
8 environmentally benign synthesis routes to avoid the usage of toxic reagents and to reduce
9 energy consumption. Appropriate selection of materials also plays an essential role in
10 sustainability. Noble metal-based nanozymes (e.g., gold, platinum, palladium) exhibit
11 exceptional catalytic activity, yet their high cost and limited availability raise concerns
12 regarding material criticality and large-scale deployment.³² On the other hand, nanozymes
13 based on earth-abundant materials, including transition metal oxides, carbon-based
14 nanostructures, and metal organic framework (MOF)-derived systems, provide more cost-
15 effective and sustainable alternatives. From the application point of view, the fabrication of
16 disposable or reusable sensing platforms must consider lifecycle and environmental impact.
17 Nanozymes immobilization on the solid support facilitates reusability and reduces material
18 waste, whereas the flexible or paper-based substrates offer biodegradable, disposable, and low-
19 cost sensing frameworks suitable for the field applications.³³ Moreover, the advancement of
20 field-deployable sensing technologies, including portable electrochemical devices,
21 smartphone-integrated colorimetric assays, and on-site SERS sensing platforms, facilitates
22 low-resource detection in agricultural and ecological environments without requiring complex
23 instrumentation.²² Despite these advancements, comprehensive lifecycle assessments and

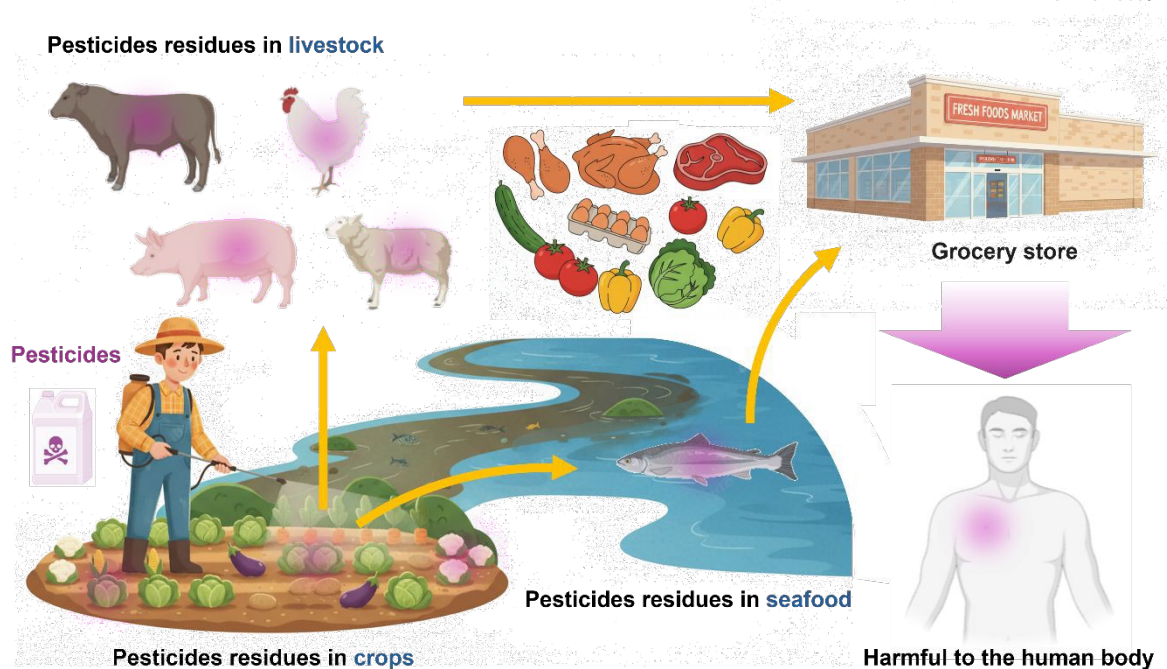


1 environmental impact analyses of nanozyme materials remain limited. Future research should
2 focus on selection of sustainable materials, scalable fabrication methods, and green synthesis
3 strategies, while also considering environmental safety, recyclability, and real-world
4 applicability to ensure that nanozyme-based pesticide sensing technologies contribute to
5 sustainable monitoring practices.^{34, 35}

7 **3. Toxic impact of pesticides on human and environmental health**

8 Pesticide residues are toxic chemicals that are often found in vegetables, fruits, cattle feed,
9 and agricultural food products, which occur in the form of metabolites, transformation products,
10 impurities, and reaction products. The effective and strategic use of pesticides requires proper
11 leaf coverage. However, farmers tend to apply excessive amounts of pesticides to ensure that
12 all crops are thoroughly covered.^{1, 4, 11, 12, 36} Therefore, the dietary intake of pesticide-
13 contaminated fruits, vegetables, edible plant leaves, and other food crops is currently among
14 the greatest threats to human health worldwide, and food poisoning incidents caused by the
15 accidental ingestion of chemical pesticide residues are not uncommon. Even low
16 concentrations of pesticide residues may affect not only food and water safety but may also
17 disrupt ecosystem dynamics. Chemical pesticide residues may affect human health through
18 direct exposure or indirect exposure via the consumption of contaminated food or water.⁶⁻¹²
19 Scheme 2 shows the routes of food contamination by pesticides.





1

2 **Scheme 2.** Systemic pathways of pesticide residues from environmental application to human
3 intake.

4

5 The key routes of pesticide exposure are ingestion, inhalation, and dermal penetration.

6 Chronic diseases have been reported in individuals that were exposed to pesticides. Direct or

7 indirect exposure to pesticide residues may cause nausea, headaches, skin irritation, birth

8 defects, developmental toxicity, dizziness, burning eyes, cancer, reproductive disorders,

9 chronic nephropathies, neurotoxicity, Parkinson's disease, central nervous system damage,

10 diabetes, genetic damage, amyotrophic lateral sclerosis, cardiovascular diseases, vomiting,

11 respiratory and digestive disturbances, and death. Organophosphorus pesticides (OPs) are

12 known to affect the reproductive system, particularly in males. For instance, parathion exposure

13 led to testicular abnormalities in male mice, in addition to chromosomal aberrations in the

14 peripheral lymphocytes of mice. Although pesticides are crucial for pest management and plant

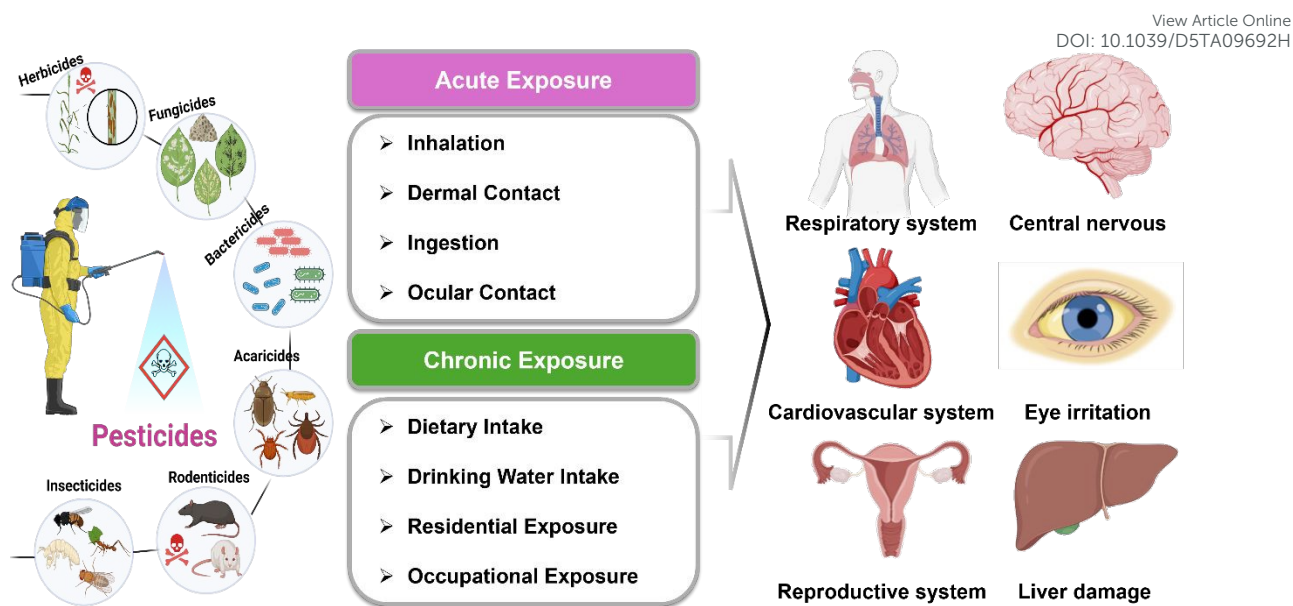


1 growth and productivity, the detection and regulation of pesticide residues in agricultural food
2 products and commodities should be strictly monitored and regulated.^{6-12, 19}

3
4 The use of chemical pesticides has considerably increased crop yields; however, their
5 utilization poses serious environmental risks. Pesticide residues have been recently identified
6 in a wide variety of environmental samples due to the rapid expansion of anthropogenic
7 activities. Massive amounts of chemical pesticides and other contaminants have also been
8 released into the atmosphere due to the recent industrialization of the agriculture sector.³⁷⁻³⁹
9 Additionally, pesticides are easily volatilized and transported, after which they are readily
10 assimilated into ecosystems and biota. The effects of elevated pesticide concentrations in the
11 environment may also be exacerbated by certain climate conditions, including temperature and
12 relative humidity. Pesticides may also percolate or leach into the soil, which in turn erodes the
13 soil and reduces its fertility and biodiversity. Further, these effects may also alter the physical
14 characteristics of water bodies, which ultimately reduces drinking water quality.⁴⁰ Honeybees
15 are generally considered outstanding sentinels for the evaluation of environmental pesticide
16 pollution. Chemical pesticides have been linked to increased honeybee mortalities, hive
17 abandonment, and decreased pollinating efficiency. In addition to honeybees, pesticide
18 residues affect a large variety of other non-target organisms such as birds and aquatic animals.^{1,}
19 ^{4, 11, 12, 19} Scheme 3 illustrates the adverse effects of pesticides on humans along with several
20 disorders.

21





Scheme 3. Adverse effects of pesticides on humans along with several disorders.

Another major cause of pesticide pollution is the improper, unsafe, and illegal disposal of chemical pesticides. Several quantitative and analytical techniques have been developed for the identification of pesticide residues in environmental samples. Importantly, these quantitative and analytical techniques can be utilized to detect residual pesticides in environmental samples with high sensitivity and selectivity. Some studies have utilized silver, gold, zinc, copper, and iron nanoparticles as effective colorimetric probes for detection of pesticide residues in environmental samples. Recently, the Quick, Easy, Cheap, Effective, Rugged, and Safe (QuEChERS) detection approach has also garnered increasing attention.^{41, 42}

Importantly, several methods have been developed based on this approach for the detection of pesticides in environmental and food samples. An efficient approach to diminish pesticide pollution and its associated adverse effects is to use safer, more reliable, natural, and non-chemical pest control agents. Furthermore, pollution control should also be addressed through the creation of environmental policies and the implementation and regular monitoring of agro-



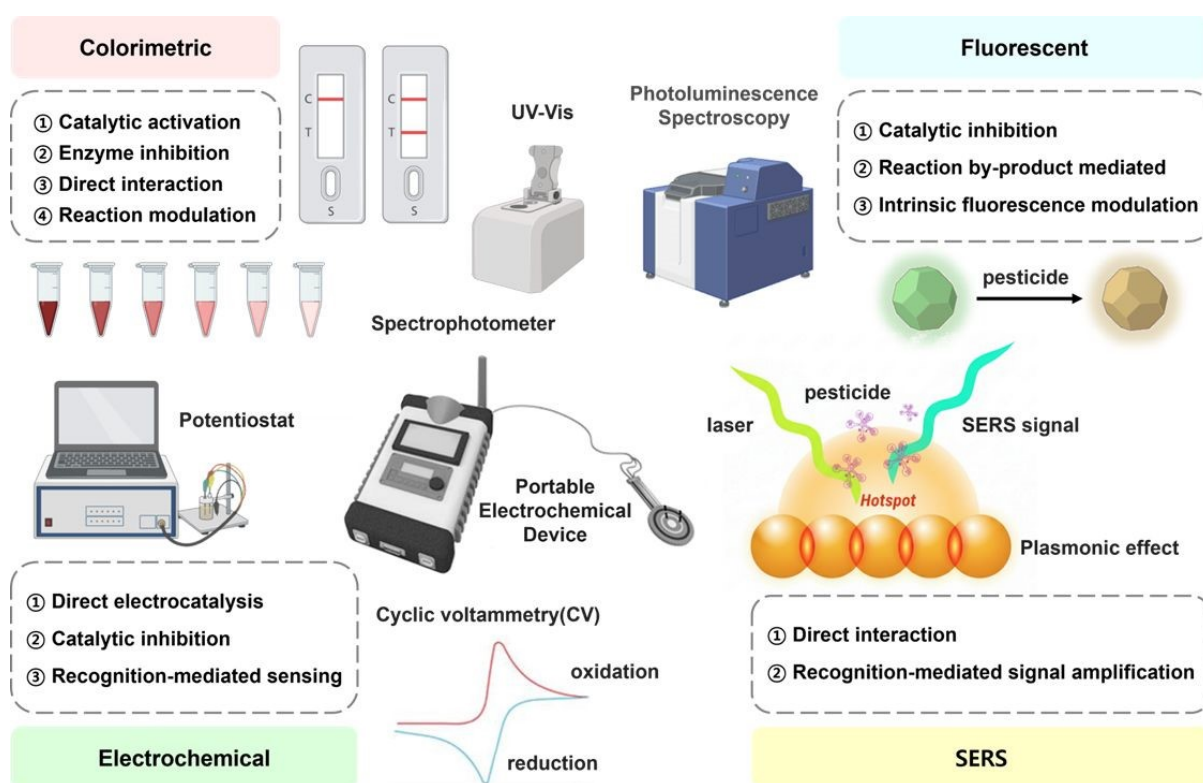
1 environmental indicators for pesticide risk assessment. In addition to reducing pesticide
2 pollution risks through natural and safer practices, these strategies may be later coupled with
3 crop rotation, land management, and field sizing to increase their efficacy.^{43, 44} Furthermore,
4 environmental, ecosystem, and biodiversity services should be clearly defined to empower
5 stakeholders, political authorities, industry professionals, conservationists, and farmers.
6 Therefore, global pesticide use should be strictly monitored to minimize their adverse effects
7 on human health and environmental safety.
8

9 **4. Potential of innovative nanozymes for the detection of pesticides**

10 Nanozymes highlighted the significance of colorimetric, electrochemical, SERS, and
11 fluorescence-based sensors for environmental monitoring applications. In this section, we
12 conducted an in-depth exploration of the innovative potential of nanozymes for the substantial
13 detection of pesticides. We analyzed the strategic advantages and operating principles of four
14 key detection platforms, i.e., colorimetric, fluorescence, electrochemical, and SERS based on
15 their signal transduction methods. In general, nanozyme-assisted pesticide detection relies on
16 four primary mechanisms, as shown in scheme 4. (i) catalytic activity modulation where
17 pesticides inhibit or enhance the enzyme-like activity of nanozymes; (ii) direct interaction
18 between pesticides and nanozyme surfaces that alters catalytic or optical properties; (iii)
19 recognition-element mediated sensing using antibodies, aptamers, or MIPs to provide
20 selectivity; and (iv) reaction by-product mediated signal generation. These mechanisms
21 emphasize various modes of signal transduction including colorimetric, electrochemical,
22 fluorescence, and SERS detection systems. Interaction mechanisms between the nanozymes
23 and pesticides were summarized and the latest research advances were elucidated. Further



1 demonstrated how nanzyme technology enables the development of ultra-sensitive and highly
 2 selective analytical methods. Through this multifaceted approach, we aim to provide a
 3 comprehensive outlook on the limitless potential of innovative nanozymes as next-generation,
 4 sensing technology for safeguarding future food security and environmental safety.



6 **Scheme 4.** Overview of detection mechanisms in nanzyme-based pesticide sensors.

9 **4.1. Nanozymes for the colorimetric detection of pesticides**

10 Pesticides are essential for enhancing agricultural productivity and play a vital role in
 11 modern agriculture. However, their residues in food and the environment are considered
 12 serious organic pollutants that pose a threat to human health and ecosystems⁴⁵. Therefore, the



1 development of technologies for rapid and convenient on-site screening of these substances in
 2 agricultural fields, during distribution, or at home is a key challenge for food safety and
 3 environmental protection. While traditional analytical methods offer high accuracy and
 4 sensitivity, their on-site application is impractical due to inherent limitations such as expensive
 5 equipment, complex sample pretreatment processes such as purification or concentration, long
 6 analysis times, and the need for skilled experts⁴⁶. One of the most promising alternatives to
 7 address these demands is the colorimetric detection method utilizing nanozymes. Colorimetric
 8 methods provide intuitive results through color changes, making them suitable for on-site use
 9 without expensive equipment or specialized training, but they have often been limited by low
 10 sensitivity and selectivity.⁴⁷ Most nanozyme-based colorimetric pesticide sensors operate
 11 through the catalytic activity modulation or through direct interaction with nanozyme surfaces,
 12 which influence the oxidation of chromogenic substrates. The advent of nanozymes has marked
 13 an innovative turning point, changing the paradigm of colorimetric detection.^{23, 46} Table 1
 14 shows the comparative analysis of nanozyme-based colorimetric sensors for pesticide detection.

15
 16 **Table 1. Comparative analysis of nanozyme-based colorimetric sensors for pesticide**
 17 **detection**

Nanozyme	Pesticides	Enzyme activity	LOD	Reaction time	Recovery rates	Ref.
NH ₂ -CuBDC	Chlorpyrifos	Peroxidase	1.57 ng/mL	-	103.25, 103.19%	48
Fe@PCN-224 NCs	Propiconazole	Peroxidase	2.74 ng/mL	10 min	94.6-109.2%	49
GSH-Fe	Thiram	Peroxidase	0.3 µg/mL	45 min	92.4-106.9%	50
Ce/ Zr-MOF	2,2-dichlorovinyl dimethyl phosphate	Peroxidase	0.32 ng/mL	30 min	95-107%	51
ASP-Cu, BPY-Cu, GMP-Cu	Sulfonylurea pesticides	Laccase, Peroxidase	0.03 µg/mL	5 min	-	52
V ₂ O ₅ -NC	Phorate, Chlorpyrifos, Methamidophos, Methyl parathion,	Catalase, Peroxidase	7.81, 42.07, 28.23,	20 min	88.1-110.5%	53



View Article Online
DOI: 10.1039/D5TA09692H

	Glyphosate		57.91, 10.14 ng/mL			
Cu-BDC-NO ₂	Glyphosate	Laccase	37 ng/mL	-	90.34-96.13%	54
PtPdNPs@g-C ₃ N ₄	Trichlorfon	Oxidase	83 pg/mL	20 min	95.78-104.34%	55
ASP-Cu	Sulfonylurea	Peroxidase	-	20 min	-	56
COF _{Fe} /CoP-Ph	Malathion	Oxidase, Catalase, Peroxidase	3.63 ng/mL	-	98.2-103.6%	57
CeO ₂ @NC	Carbosulfan	Oxidase	1.26 ng/mL	-	100.2-104.8%	58
ZnCo-ZIFs@MIL-101(Fe)	Glyphosate	Peroxidase	75 ng/mL	2 min	97.14-117.37%	59
Mn@NC	Phoxim	Peroxidase	1.27 ng/mL	20 min	95.00-107.93%	60
PDA/MnO ₂ NPs	Carbosulfan	Oxidase	0.63 ng/mL	1 min	95.80-103.19%	61
PtCu _{SA} @TriMOF	Carbosulfan	Peroxidase	1.60 ng/mL	50 min	85-121%	62
Porous nitrogen-doped carbon carrier (Pt-Np-C)	Chlorpyrifos, Glyphosate, Carbaryl, Pentachloronitro benzene, Metsulfuronmethyl	Peroxidase	8.76, 4.23, 5.03, 7.38, 9.53 µg/mL	-	-	63
Zr-based metal-organic framework	Paraoxon	Phosphatase	72.65 ng/mL	-	98.1-112.5%	64
PB@Fe-COF@Au	Chlorpyrifos	Peroxidase	0.61 ng/mL	-	94/3-110.2%	65
PtCu ₃ alloy nanocrystals	Trichlorfon	Peroxidase	0.5 ng/mL	5 min	-	66
Fe-N/C SAzyme	Malathion	Oxidase	79 pg/mL	10 min	98.6-105.4%	67
Oxidized MXene quantum dots@CuNi bimetal (MQDs@CuNi)	Glyphosate	Peroxidase	0.19 µg/mL		97.4-111.2%	68
CuO/Fe ₂ O ₃ nanozymes	Glufosinate	Peroxidase	5.07 pg/mL	66 min	91.48-106.02%	69
Fe-N/C SAzymes	Dimethoate	Oxidase	96 pg/mL	10 min	95.01-103.94%	70
GSH-Cu	Thiram	Laccase	2.00 ng/mL	-	87.1-106.1%	71
0.5 MQDs-NiCoP/NF nanozymes	Glyphosate	Peroxidase	0.18 µg/mL	-	97.4-111.2%	72
Cu-doped carbon nanozyme	Thiophanate-methyl	Oxidase	0.04 µg/mL	-	98.1-102.1%	73



Fe-Co MNPs	Phorate, Profenofos, Isocarbophos, Omethoate	Peroxidase	0.16, 0.16, 0.03, 1.6 ng/mL	30 min	89.19–108.35 %	74
N-CDs/FMOF-Zr	Glyphosate	Peroxidase	13.1, 1.5, 11.5 ng /mL	9 min	92.83–108.80%	75
FeCoCu flower-like structure (FeCoCu-FN)	Parathion methyl	Peroxidase	11 pg/mL	-	96.5 % to 105.0 %	76
Pt/Co ₃ O ₄	Thiram, Ziram	Peroxidase, Oxidase, Catalase, Superoxide dismutase	15.63 ng/mL, 1.03 µg/mL	-	95.33-101.60%	77

1

4.1.1. Strategic advantages of nanozyme-based colorimetric detection methods

The first strategic advantage of nanozyme-based colorimetric methods is inherent signal amplification for high sensitivity. As pesticides typically exist at very low concentrations, high sensitivity is essential for their detection. Nanozymes, like natural enzymes, act as catalysts, where a single particle can rapidly and repeatedly convert numerous chromogenic substrate molecules (e.g., 3,3',5,5'-tetramethylbenzidine, TMB).⁷⁸ This is a powerful mechanism that amplifies the minute initial signal caused by trace amounts of pesticides into a distinct color change visible to the naked eye. Because of catalytic amplification, nanozyme-based colorimetric assays afford lower limits of detection than their noncatalytic counterparts.⁷⁹ The second advantage is superior robustness and stability over natural enzymes. Traditional biosensors usually use the natural enzymes (for example AChE), which possess high specificity but suffer from inherent limitations of protein structure, making them susceptible to changes in external environmental conditions such as temperature, organic solvents, and pH.^{23, 24} This reduces the shelf life of sensor and inhibits its direct application to real-world samples such as soil extracts or fruit surface. Conversely, nanozymes derived from stable inorganic materials such as metal oxides, metals, or carbon exhibit chemical and physical adaptability, preserving their catalytic activity even in extreme pH or higher temperature environments. This



1 dramatically extends the sensor's shelf life and provides strong field applicability, allowing for
2 direct use on various real samples without the need for purification or complex pretreatment
3 steps. Finally, there is excellent cost-effectiveness and design flexibility. Natural enzymes
4 require complex biological expression and purification processes, leading to high production
5 costs. Nanozymes can be mass-produced at a low cost through simple chemical synthesis
6 methods, a condition highly favorable for developing low-cost, disposable strip-type diagnostic
7 kits.⁷⁸ Furthermore, by precisely controlling the size, composition, morphology, and surface
8 chemistry of the nanomaterials, it is possible to optimize their catalytic activity or easily
9 introduce molecular recognition elements, such as aptamers that specifically bind to certain
10 pesticide molecules, onto their surface.^{46, 80} This tailor-made design capability signifies the
11 limitless potential for developing high-performance sensors optimized for the target analyte of
12 interest and forms the basis for the various innovative nanozymes discussed in this review. The
13 subsequent sections will delve into the specific mechanisms through which these advantages
14 are realized, categorized by the nature of the nanozyme-pesticide interaction.

15

16 **4.1.2 Classification of nanozyme-based colorimetric sensors for pesticide detection**

17 The most intuitive method in nanozyme-based colorimetric sensing uses the principle
18 where the pesticide molecule itself interacts directly with nanozyme to alter its intrinsic
19 catalytic function. This process is substantially efficient as it eliminates the need for auxiliary
20 enzymes or intermediate steps, leading to faster and simpler analytical systems.⁸¹ The direct
21 interaction approach can be further classified into two primary modes: modulation of separate
22 chromogenic substrate's oxidation, and direct enzymatic conversion of pesticide itself into
23 signal molecule. Modulation of chromogenic substrate oxidation is the most widely explored

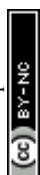


1 direct interaction pathway. Intrinsic enzyme-like activity (for example oxidase or peroxidase)
2 nanozyme's is utilized to catalyze the oxidation of chromogenic substrate like TMB.^{49, 74, 82} The
3 target pesticide molecule then interact directly with nanozyme, acting as either an enhancer or
4 an inhibitor of this catalytic activity, leading to a assessable change in intensity of color.
5 Indirect interaction mechanisms rely on a 'mediator' or 'byproduct', whose concentration is
6 modulated by the presence of the pesticide, to regulate the nanozyme's activity. This approach
7 can significantly expand the selectivity and design flexibility of the detection system by
8 incorporating auxiliary enzymes that react with a specific class of pesticides or by utilizing
9 characteristic chemical reactions of the pesticide.^{46, 83} This strategy allows the highly efficient
10 catalytic activity of nanozymes to be coupled with well-established biochemical recognition
11 events. The most representative and widely adopted example is using the principle of AChE
12 inhibition.^{48, 53, 57, 62, 66, 67, 70, 76} Organophosphorus and carbamate pesticides are potent inhibitors
13 that irreversibly bind to the active site of AChE, an essential enzyme for normal nerve signal
14 transmission. By coupling this principle with a nanozyme system, highly sensitive sensors can
15 be implemented. In this system, the key 'mediator' is thiocholine (TCh), which is produced
16 from the hydrolysis of the AChE substrate, and acetylthiocholine (ATCh). The concentration
17 of TCh mediators is inversely proportional to pesticide concentration, which acts as a switch
18 to control the nanozyme's activity.

19

20 **4.1.2.1 Detection mechanisms based on direct interaction**

21 Typically, colorimetric sensors based on the simple direct mechanism exhibit a 'turn-off'
22 response due to direct inhibition by the pesticide. In this mode, the pesticide binds to the
23 nanozyme's active sites or blocks substrate access, causing a decrease in the colorimetric signal.



1 In 2024, *Yan et al* designed bio-inspired glutathione-iron hybrid (GSH-Fe) nanozyme to detect
2 the fungicide thiram. Thiram molecules, containing sulfur atoms, exhibit strong affinity for the
3 iron centers of the nanozyme, leading to surface passivation and inhibition of its peroxidase-
4 like activity. This system achieved a low detection limit of 0.3 $\mu\text{g/mL}$ within a linear range of
5 0.3-10 $\mu\text{g/mL}$. Also, they achieved a recovery rate of 92.4% to 106.9%. To improve stability
6 and practicality, the nanozyme was immobilized in a hydrogel, and a simple smartphone with
7 ImageJ software was used for quantification, demonstrating its potential for on-site analysis of
8 fruit juice samples.⁵⁰ In 2025, *Zhang et al.* designed a ternary MOF, ZnCo-ZIFs@MIL-101(Fe),
9 and engineered the designed material with multiple metal sites to detect glyphosate. The
10 principle relies on the coordination of glyphosate's phosphonic acid group with these exposed
11 metal active sites, hindering TMB oxidation. This sensor demonstrated a wide linear range of
12 0-1 $\mu\text{g/mL}$ and an LOD of 75 ng/mL. Its utility for on-site applications was proven by
13 integrating it into a smartphone and paper-based microfluidic chips (μPAD) for analyzing
14 samples.⁵⁹ Advanced 2D materials have also shown great promise. MQDs@CuNi, a composite
15 of MXene quantum dots (MQDs) and a copper-nickel bimetal, and MQDs-NiCoP/NF, an
16 urchin-like nickel-cobalt phosphide structure modified with MQDs, both function by the
17 principle of glyphosate directly binding to their active sites and inhibiting their peroxidase-like
18 activity (Fig. 1a). These systems achieved remarkably low detection limits of 1.06 μM ,
19 respectively, showcasing their potential for highly sensitive trace analysis.⁷² In 2023, *Zhang et*
20 *al.*, used Cu-doped carbon nanozyme to detect the fungicide thiophanate-methyl. The
21 interaction between the pesticide's sulfur and nitrogen atoms and the copper active sites on the
22 nanozyme surface inhibits its oxidase-like activity, resulting in a 'turn-off' response with a low
23 LOD of 0.04 $\mu\text{g/mL}$.⁷³ *Li et al.* developed a bioinspired laccase-mimicking catalyst,
24 designated as GSH-Cu, by simulating the copper active sites and spatial amino acid



1 microenvironment of natural enzymes. This nanozyme, synthesized through the coordination
2 of Cu^{2+} with glutathione (GSH) peptides, exhibits catalytic functions that conform to the
3 Michaelis-Menten kinetics of natural laccase (Fig. 1b). The sensing mechanism is based on a
4 pesticide detection mechanism through a direct interaction method, where the catalytic activity
5 of the GSH-Cu complex is triggered for inhibition by thiram. Specifically, the coordination
6 between sulfur (S) and copper (Cu) atoms disrupts the surface structure of the catalyst and leads
7 to the formation of copper nanoparticles, a process characterized as catalytic inhibition (Fig.
8 1c). The developed sensor demonstrated outstanding performance with a laccase (LOD) of 2.00
9 ng/mL in solution, covering a linear range from 2.5 to 250 ng/mL. This system proved highly
10 effective for the sensitive detection of pesticides in real food samples, such as fruit juices and
11 environmental water. To determine sensing mechanism, absorption spectra were measured to
12 ascertain the coordination between copper ions and thiram (Fig. 1d).⁷¹

13

14

15



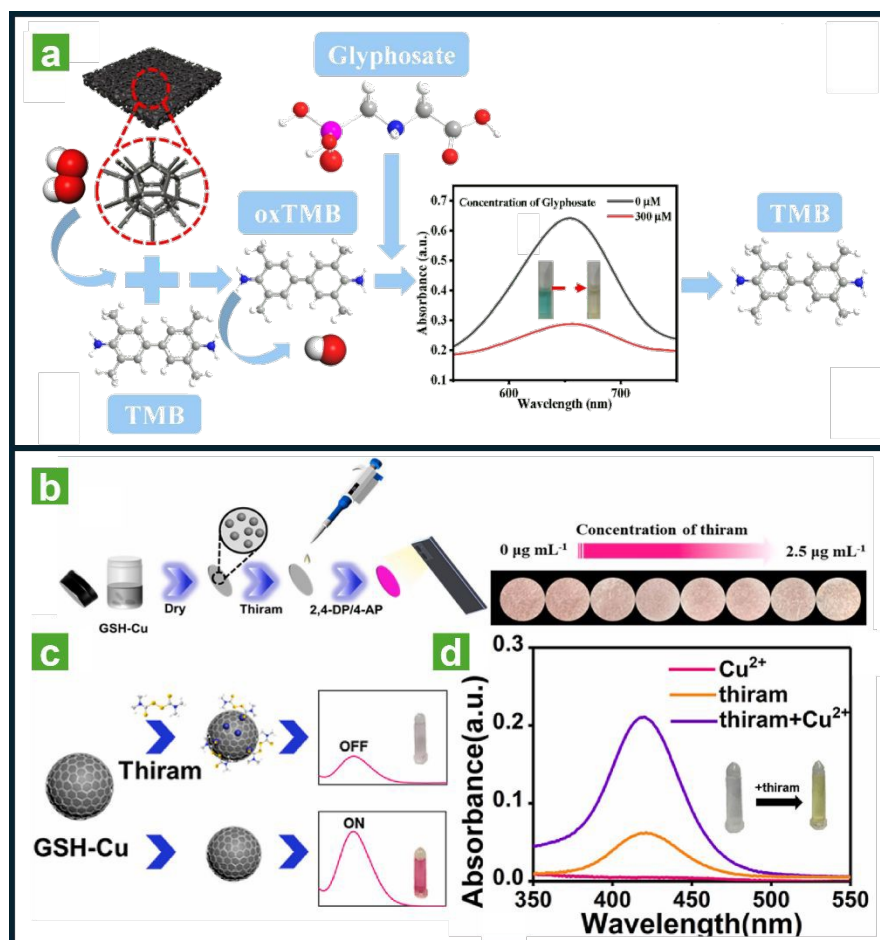
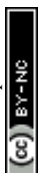


Fig. 1. Pesticide detection mechanism through direct interaction method. (a) Diagrammatic depiction of the colorimetric sensing of glyphosate facilitated by 0.5 MQDs-NiCoP/NF nanozymes. Reproduced from reference⁷² with permission from Elsevier, copyright 2025. (b) Conceptual design of a paper-based analytical platform for the monitoring of thiram residues. (c) Illustrative representation of the catalytic inhibition of GSH-Cu complexes triggered by thiram. (d) UV-vis electronic absorption profiles of Cu^{2+} , thiram, and the resulting Cu^{2+} -thiram coordination complex. Reproduced from reference⁷¹ with permission from Elsevier, copyright 2022.



1 Conversely, 'Turn-on' sensors based on direct enhancement are highly advantageous for
2 achieving superior sensitivity due to their inherently low background signal. In 2024, *Xu et al.*
3 developed an iron-based MOF, Fe@PCN-224 NCs, whose peroxidase-like activity is markedly
4 enhanced in the presence of the fungicide propiconazole. This is interpreted as propiconazole
5 inducing a conformational change in the MOF structure that improves TMB substrate
6 accessibility to the active iron-porphyrin sites. This 'turn-on' sensor achieved an impressive
7 LOD of 8 nM over a linear range of 0.03-90 μM in vegetable samples, demonstrating a rare
8 case of analyte-induced activation.⁴⁹ In 2025, *Liu et al.* developed a copper-based MOF, Cu-
9 BDC-NO₂, was rationally designed by tuning the ligand with an electron-withdrawing nitro
10 group, which enhanced its laccase-like activity. This activity was further promoted by
11 glyphosate, leading to a 'turn-on' response with an LOD of 37 ng/mL. This design ensures that
12 the nanozyme's activity is restored only in the presence of the target pesticide, which displaces
13 the aptamer from the nanozyme surface, thus securing overwhelming selectivity.⁵⁴ The most
14 representative phosphatase-like activity was exhibited by Zr-based MOFs. Nanozyme
15 developed by *Zhu et al.* in 2025, directly cleavage the phosphoester bond of the paraoxon
16 pesticide, converting the colorless paraoxon into the distinctly yellow-colored p-nitrophenol.
17 The Zr₆ clusters within the MOF structure act as potent Lewis acid sites, activating water
18 molecules for a nucleophilic attack on the phosphorus center of the pesticide, thereby cleaving
19 the ester bond. This constitutes a perfect 'turn-on' system where the presence of the pesticide
20 directly generates a color signal from 'none' to 'some', achieving a LOD of 0.195 μM . As this
21 method is not based on redox reactions, it can fundamentally exclude interference from
22 common redox-active substances in the sample, leading to very high reliability and analytical
23 accuracy. This method signifies a paradigm shift from the indirect signal generation to direct

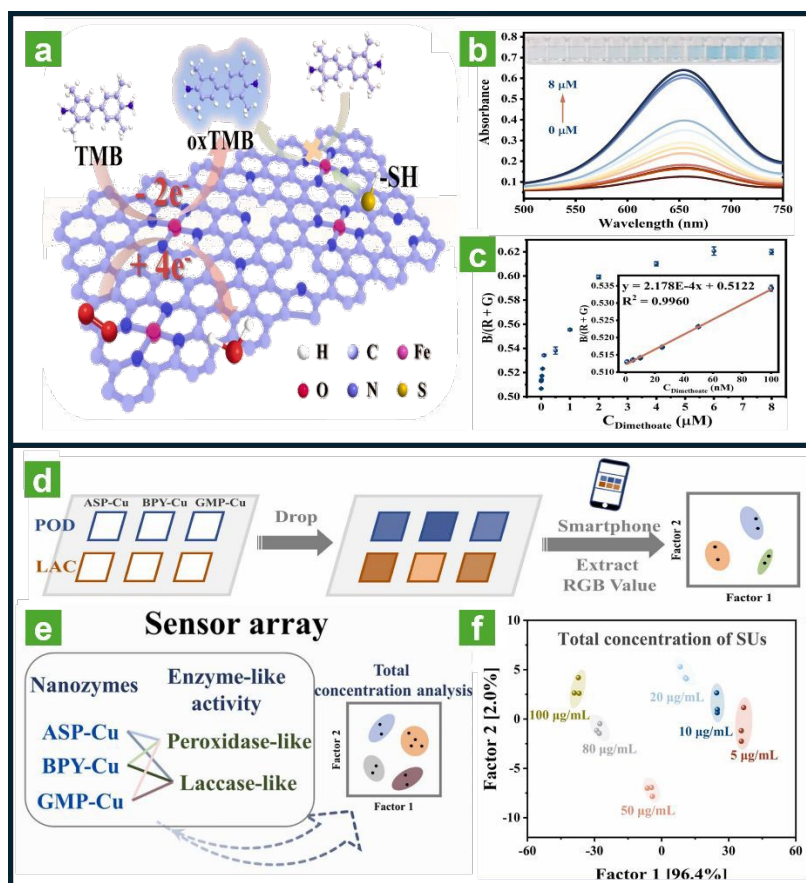


1 transformation of analyte, offering a highly robust and specific platform for the analysis
2 pesticide.⁶⁴

3 **4.1.2.2. Detection mechanisms based on indirect interaction**

4 In fact, indirect detection methods, inspired by the principle of AChE inhibition, are more
5 traditional and have been researched more extensively than direct detection methods. Wang *et*
6 *al.* (2024) applied molecular imprinting approach on Mn@NC nanozyme with phoxim as
7 template molecule. This method created molecularly imprinted sites on the surface of
8 nanozyme that precisely fit phoxim molecules, allowing it to detect selectively the phoxim
9 while excluding other pesticides with similar structures. Fabricated sensor recorded a wide
10 linear range of 5-1000 ng/mL and an LOD of 1.27 ng/mL in colorimetric approaches, proving
11 that molecular imprinting is an effective method for simultaneously enhancing selectivity and
12 sensitivity.⁶⁰



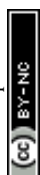


1
2 **Fig. 2.** Pesticide detection mechanism through indirect interaction method. (a) Conceptual
3 diagram illustrating the oxidase-mimicking behavior of Fe-N/C SAzymes and the inhibitory
4 effect of mercapto-containing molecules on their catalytic activity. (b) UV-vis absorbance
5 profiles of the colorimetric assay at various dimethoate concentrations (0 to 8 μM); the inset
6 provides the corresponding visual color transitions. (c) Analytical correlation between the
7 B/(R+G) color intensity ratio and dimethoate levels, with the inset displaying the established
8 linear regression model. Reproduced from reference⁷⁰ with permission from Elsevier,
9 copyright 2024. (d) Illustrative schematics of paper-based sensor arrays that leverage enhanced
10 multienzyme-like catalytic activities. (e) Development of a six-channel sensor array based on
11 the laccase- and peroxidase-mimicking activities of ASP-Cu, BPY-Cu, and GMP-Cu for the
12 selective detection of sulfonyleureas. (f) Linear discriminant analysis score plot for the



1 qualitative classification of sulfonylurea (SU) mixtures across a concentration range of 5 to
2 100 $\mu\text{g/mL}$. Reproduced from reference⁵² with permission from Elsevier, copyright 2024.

3
4 Zhang et al.⁷⁰ developed a sensor that indirectly detects pesticides using an Fe-N/C single-
5 atom nanozyme (SAzyme). The sensor's operating principle is based on an oxidase-like
6 reaction in which the Fe-N/C nanozyme converts colorless TMB into the blue-colored oxidized
7 TMB (oxTMB). In this process, an enzyme called acid phosphatase (ACP) is introduced, which
8 hydrolyzes L-ascorbic acid-2-phosphate (AAP) to produce ascorbic acid (AA), an inhibitor of
9 the color reaction (Fig. 2a). In the absence of pesticides, the ACP enzyme is active and produces
10 AA, which blocks the color reaction, causing the solution to remain colorless. Conversely,
11 when a pesticide is present, it inhibits the activity of the ACP enzyme, preventing the
12 production of AA. As a result, the Fe-N/C nanozyme is free to oxidize TMB, turning the
13 solution blue (Fig. 2b). Thus, the presence of the pesticide determines the expression of the
14 blue color, operating in a 'turn-on' fashion. Through this indirect detection method, the sensor
15 demonstrated excellent performance. Based on the pesticide dimethoate, it achieved a very low
16 LOD of 0.4177 nM (Fig. 2c). Additionally, in recovery experiments, which indicate accuracy
17 in real fruit and vegetable samples, it showed high reliability with results ranging from 95.01%
18 to 103.94%.⁷⁰ Additionally, Tian et al. reported a unique mechanism that amplifies a signal
19 through indirect interaction. In this study, three copper-based nanozymes ASP-Cu, BPY-Cu,
20 and GMP-Cu-were utilized to selectively detect sulfonylurea (SU) pesticides (Fig. 2d). The
21 core principle of this sensor is based on the phenomenon where sulfonylurea pesticides, instead
22 of directly inhibiting the enzyme-like activities (laccase-like and peroxidase-like) of the
23 nanozymes, actually enhance their activity. This is a 'turn-on' method where the pesticide



1 molecule indirectly promotes the nanozyme's catalytic reaction to generate a stronger
2 colorimetric signal, thereby detecting the pesticide's presence (Fig. 2e). This sensor achieved a
3 LOD of 0.03 $\mu\text{g/mL}$ for the total concentration of sulfonylurea pesticides and demonstrated
4 high accuracy and reliability with a recovery rate in real water samples ranging from 91.53%
5 to 103.5% (Fig. 2f).⁵²

6
7 The PtPdNPs@g-C₃N₄ nanozyme that developed by *Shen et al.* in 2023, has oxidase-like
8 activity, which is inhibited by the thiol group of the thiocholine produced in the absence of
9 pesticides. In contrast, when a pesticide is present, AChE is inhibited, no thiocholine is
10 produced, and consequently, the nanozyme's activity is 'turned on', resulting in a strong
11 colorimetric signal. This sensor showed an LOD of 83 pg/mL in colorimetric approaches,
12 demonstrating a sensitive indirect detection route.⁵⁵ Similarly, PtCu₃ alloy nanocrystals
13 developed as high-sensitivity 'turn-on' sensors for pesticides based on the AChE inhibition
14 principle. It achieved impressive LODs of 0.05 ng/mL to trichlorfon, and demonstrated the
15 feasibility of smartphone-based quantification for on-site analysis.⁶⁶ The triple composite
16 PB@Fe-COF@Au were also developed as 'turn on' sensor for trichlorfon detection. This
17 system is particularly noteworthy for its "triplet peroxidase-like activity" where the Prussian
18 blue, COF, and Au components all contribute to signal generation, creating a highly amplified
19 response. This multi-component design capitalizes on the high catalytic activity of PB and Au,
20 coupled with the large surface area and analyte-capturing ability of the COF, to create a highly
21 efficient cascade reaction.⁶⁵

22



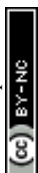
1 Regulation of nanozyme activity by byproducts generated from the chemical reaction of the
2 pesticide itself is also an effective strategy. This method can enhance selectivity by utilizing
3 chemical reactions unique to a specific pesticide, leading to AChE-free assays, which avoids
4 the use of unstable natural enzymes. In 2024, *Zhu et al.* developed CeO₂@NC nanozyme, also
5 utilizes this principle, promoting the hydrolysis of carbosulfan and using the generated SH
6 group in acidic condition to modulate its activity, achieving an LOD of 3.3 nM. This approach
7 is particularly valuable as it builds specificity into the system based on the unique chemistry of
8 the target analyte, rather than relying solely on a general enzymatic inhibition mechanism.⁵⁸
9 Sometimes, multiple steps of indirect interaction are combined to create more sophisticated
10 sensing cascades. In 2024, *Liu et al.* engineered copper-based MOF NH₂-CuBDC to detect the
11 insecticide chlorpyrifos. In this system, chlorpyrifos inhibits AChE activity, and the resulting
12 lack of thiocholine production alters its interaction with copper ions (Cu²⁺), which in turn
13 modulates the peroxidase-like activity of the nanozyme. Designed sensor achieved an LOD of
14 1.57 ng/mL in colorimetric method, showcasing how multiple equilibria can be coupled for
15 sensitive detection.⁴⁸ Another cascade system was built using the FeCoCu flower-like structure,
16 which could be switched between 'turn-off' and 'turn-on' modes. In one mode, AChE inhibition
17 prevented the production of the inhibitory thiocholine. In another mode, the product of AChE
18 (choline) was further converted by choline oxidase (ChOx) to produce H₂O₂, the substrate for
19 the nanozyme's peroxidase activity, leading to a 'turn-on' signal upon AChE inhibition. They
20 achieve LOD of 0.11 and 0.13 ng/mL and recovery rate 96.5-105.0% and 94.0-105.0% in
21 AChE method and ChOx single enzyme method. Thus, while indirect mechanisms based on
22 mediators or byproducts may require more complex system designs, they can be used as
23 potential strategy for enhancing sensor selectivity or diversifying signal transduction pathways
24 by incorporating specific chemical reaction routes into the design.⁷⁶



1

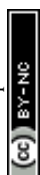
2 4.2. Nanozymes for the fluorescent detection of pesticides

3 Fluorescence detection is currently recognized as one of the most sensitive and selective
4 analytical techniques in the field of pesticide analysis. It plays a vital role in food safety and
5 environmental monitoring, particularly due to its high sensitivity, real-time monitoring
6 capabilities, and potential for visual discrimination. The emergence of nanozyme-based
7 fluorescent detection systems has brought innovative changes to this field. This is because they
8 effectively overcome the fundamental limitations of traditional natural enzyme-based systems
9 while further amplifying the inherent advantages of fluorescence detection. The integration of
10 nanozymes with fluorescent detection mechanisms is more than a simple combination of two
11 technologies; through synergistic effects, it is fundamentally changing the paradigm of
12 pesticide detection. These systems are widely applied for the detection of organophosphorus
13 pesticides (OPs), carbamate pesticides, glyphosate, neonicotinoid pesticides, and various other
14 pesticide compounds, providing tailored detection strategies suited to their respective chemical
15 properties and mechanisms of action.^{2, 84, 85} The core working principle of nanozyme-based
16 fluorescence detection is found on the complex and sophisticated interaction between target
17 pesticide and the nanozyme, which ultimately induces a quantitative change in the fluorescent
18 signal. This interaction is not a simple one-dimensional reaction but occurs through a multi-
19 step mechanism at the molecular level. Key mechanisms that work in combination include
20 enzyme activity modulation, the inner filter effect (IFE), Förster resonance energy transfer
21 (FRET), photoinduced electron transfer (PET), aggregation-induced emission (AIE) and
22 electron exchange (EE). Each of these mechanisms is selectively activated depending on the
23 molecular structure of a specific pesticide and the physicochemical properties of the nanozyme



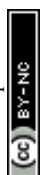
1 system. This enables the construction of detection platforms that can simultaneously achieve
2 extremely high specificity and sensitivity.^{86, 87}

3
4 The fluorescence detection mechanism based on enzyme activity inhibition and modulation
5 is one of the most extensively studied and practical approaches, demonstrating excellent
6 performance, particularly in the detection of organophosphorus and carbamate pesticides. The
7 core of this method is that pesticides specifically inhibit the activity of key biological enzymes,
8 such as AChE. This inhibition modulates the catalytic activity of a nanozyme through an
9 elaborately designed multi-step cascade reaction, ultimately inducing a quantitative change in
10 the fluorescent signal. For example, an organophosphorus pesticide forms a covalent bond with
11 the serine residue in the active site of AChE, irreversibly inhibiting the enzyme. This blocks
12 the hydrolysis of ACh. Consequently, the subsequent oxidation of choline to hydrogen
13 peroxide (H₂O₂) by choline oxidase (ChOx) is reduced. This ultimately leads to decreased
14 catalytic activity of a peroxidase-like nanozyme, which alters the oxidation state of a
15 fluorescent substrate and results in a measurable change in the fluorescence signal. The primary
16 advantage of such a cascade reaction system is its inherent signal amplification effect: a single
17 pesticide molecule can inhibit numerous enzyme molecules, and each enzyme can, in turn,
18 process a multitude of substrate molecules.^{88, 89} Fluorescence detection systems based on direct
19 interaction and molecular recognition operate on a mechanism where a pesticide interacts with
20 a nanozyme through direct chemical bonding or physical adsorption. This interaction alters the
21 nanozyme's structural and electronic properties, which in turn directly leads to a change in its
22 fluorescence characteristics. In this mechanism, specific chemical functional groups on the
23 pesticide molecule such as sulfur, phosphorus, and halogen atoms, or aromatic ring structures

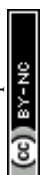


1 form various intermolecular interactions with the active sites or metal centers on the
2 nanozyme's surface, including hydrogen bonds, coordination bonds, π - π stacking, and van der
3 Waals forces. A particularly interesting example is when a sulfur-containing pesticide like
4 malathion forms a strong Ag-S coordination bond with a silver nanoparticle-based nanozyme.
5 This alters the nanozyme's surface electronic structure, which selectively inhibits its oxidase-
6 like activity and induces a change in the fluorescence signal associated with the oxidation of
7 TMB.⁹⁰ The greatest advantages of this direct interaction mechanism are that high specificity
8 can be achieved without using any biological enzymes and that it demonstrates stable
9 performance even in complex matrices.^{49, 91}

11 The FRET based fluorescence detection mechanism is a highly sophisticated and sensitive
12 method that utilizes non-radiative transfer of energy between a fluorescence acceptor and a
13 donor. In these systems, a nanozyme plays the role of either the donor or the acceptor, and the
14 presence of a target pesticide dynamically alters the FRET efficiency, which induces an
15 increase, decrease, or spectral shift in the fluorescence intensity. For FRET to occur effectively,
16 several conditions must be met, there must be significant spectral overlap between the donor's
17 emission spectrum and the acceptor's absorption spectrum, the distance between two molecules
18 must be within the 1-10 nm range, and the relative orientation between the transition dipole
19 moments of the donor and acceptor must be appropriate.^{92, 93} These conditions can be precisely
20 controlled in nanozyme-based systems in response to the presence of a pesticide. For instance,
21 nanozymes with fluorescent properties such as gold nanoparticles, graphene quantum dots
22 (GQDs), or MOFs doped with rare-earth elements can be utilized as key components in these
23 FRET systems.⁹⁴ The IFE is a phenomenon in which the observed fluorescence intensity



1 decreases because the emission or excitation light of a fluorophore is absorbed by an absorber
2 present in the solution. In nanozyme-based systems, this mechanism involves the products or
3 intermediates of a catalytic reaction acting as the absorber to modulate the fluorophore's signal.
4 A typical example is when a peroxidase-like nanozyme oxidizes a colorless substrate, such as
5 TMB, to produce a colored oxidation product (e.g., oxTMB). This product exhibits strong
6 absorption in a specific wavelength region, effectively quenching the signal of a fluorophore
7 that emits fluorescence in the same or a similar wavelength region. A unique advantage of IFE-
8 based detection systems is that, unlike FRET, physical proximity between the fluorophore and
9 the absorber is not essential. This provides greater choice in system design and enables various
10 combinations of nanozymes and fluorophores.⁹⁵⁻⁹⁷ The PET mechanism is a highly
11 sophisticated and sensitive detection method that utilizes the phenomenon of fluorescence
12 quenching or enhancement, which occurs when a fluorophore in a photoexcited state exchanges
13 electrons with an electron donor or acceptor. In the PET process, an electron from the excited
14 fluorophore is transferred to a nearby electron acceptor (quencher), or conversely, an electron
15 from a donor is transferred to the highest occupied molecular orbital (HOMO) of the
16 fluorophore, leading to the suppression of fluorescence emission.⁹⁸ In nanozyme-based systems,
17 the PET mechanism is widely observed, particularly in gold nanoclusters (AuNCs), carbon
18 quantum dots (CQDs), and various fluorescent MOFs, where the presence of a pesticide
19 modulates this electron transfer process to induce a change in the fluorescent signal. For
20 instance, in the BSA-stabilized gold nanoclusters (BSA-AuNCs), it has been shown that PET
21 quenchers such as ascorbic acid, mercury ions, and dopamine not only quench fluorescence but
22 can also simultaneously modify nanoclusters' intrinsic peroxidase-like activity. This attribute
23 can be utilized to develop dual-mode (colorimetric and fluorescent) sensing systems.⁹⁹ PET
24 quenchers do not inherently inhibit nanozyme activity. In certain circumstances, Cu^{2+} or



1 specific cyanine dyes, can augment the peroxidase-like activity. By effectively utilizing this
2 feature, it is possible to design highly selective and sensitive pesticide sensing systems.^{100, 101}
3 The electron exchange mechanism elucidates the change in fluorescence characteristics by
4 direct transfer of electrons between a target molecule and a nanozyme, a phenomenon closely
5 related to the surface quenching effect. This method involves defect sites or metal ions on the
6 nanozyme surface functions such as electron acceptors or donors, facilitating direct electron
7 transfer with a fluorophore, hence causing either amplification or quenching of fluorescence.
8 The electron exchange mechanism is highly sensitive to surface state and electronic structure
9 of nanozyme. Consequently, its direction and efficiency can vary substantially depending on
10 the nanozyme's size, composition, shape, and surface functionalization. The AIE process is an
11 innovative detection approach that employs a unique phenomenon. Unlike conventional
12 fluorophores that undergo fluorescence quenching in an aggregated state, AIE-active
13 compounds (AIEgens) demonstrate pronounced fluorescence upon aggregation. This
14 phenomenon is attributable to the limitation of intramolecular rotational effects. The
15 underlying principle is that when molecules aggregate, their internal rotations are physically
16 restricted. This restriction suppresses non-radiative energy dissipation pathways, which in turn
17 enhances fluorescence emission. In nanozyme-based AIE systems, the catalytic activity of a
18 nanozyme can control the aggregation state of AIE molecules. Conversely, a target pesticide
19 can alter the aggregation state of an AIE-active nanozyme (AIEzyme), thereby inducing a
20 change in the fluorescence signal. For instance, an AIEzyme with organophosphorus hydrolase
21 (OPH)-like activity can hydrolyze an organophosphorus pesticide. During this process, its own
22 fluorescence is quenched, enabling a self-reporting function. This allows for the specific and
23 sensitive detection of nerve agents and pesticides.¹⁰² Table 2 shows the comparative analysis
24 of nanozyme-based fluorescent sensors for pesticide detection.



1

2 **Table 2. Comparative analysis of nanozyme-based fluorescent sensors for pesticide**
3 **detection**

Nanozyme	Pesticides	Enzyme activity	LOD	Reaction time	Recovery rates	Ref.
66-IS-Zn	Profenofos, phoxim	phosphatase		30 min		103
Cu-BDC-NH ₂	Malathion, Dicofol, Fenvalerate, Pirimicarb, Dimethoate, Phosmet	Peroxidase		15 min		104
MB/COF@MnO ₂	Dichlorvos, Omethoate, Trichlorfon, Chlorpyrifos, Chlorfenvinphos	Oxidase	83 pg/mL	30 min	97.8 %-105 %	105
PtPdNPs@g-C ₃ N ₄	Dimethoate, Phosphine acetyloxalamine	oxidase	33 pg/mL	20 min	95.78–104.34 %	55
ASP-Cu	Metsulfuron, Rimsulfuron, Sulfosulfuron, Halosulfuron-methyl	Peroxidase		15 min	-	56
JLU-MOF22	Benfuracarb	Oxidase	76 ng/mL	1 min	98.8 to 102.55 %	106
Zr-TCPE	Paraoxon	Phosphatase	48.99 ng/mL	30 min	96.1–115.0%	64
Fe-N/C single-atom	Malathion	Oxidase	0.139 ng/mL	10 min	98.6%-105.4%	67
NH ₂ -CuBDC	Chlorpyrifos	Peroxidase	2.33 ng/mL	10 min	103.19%	48
NH ₂ -MIL-101(Fe)	Glyphosate	Peroxidase	3.16 ng/mL	15 min	92.4 %-115.6 %	107
Cu-MIL-53(Fe)-2	Malathion	Peroxidase	42.62 ng/mL	15 min	96.65 %-108.92 %	108
MQDs@CoO _x -300	Profenofos	Peroxidase	0.254 ng/mL	10 min	94.7 %-104 %	109



Cu-BH	Thiophanate methyl	oxidase	10.27 ng/mL	30 min	91.5 %- 93.5 %	110
AuNEs	Chlorpyrifos, Dimethoate, Methyl paraoxon	Peroxidase	7 ng/mL, 14 ng/mL, 16 ng/mL	25 min		111
MQDs@CuNi	Glyphosate	Peroxidase	0.12 pg/mL	10 min	97.4 %- 111.2 %	68
CTEzyme	Methidathion	Peroxidase	0.625 ng/mL	20 min	97.3%- 105.7 %	112
Zr-MOF	Paraoxon	Phosphatase	1.67 ng/m L	60 min	99.8 %- 100.6 %	113

1

2 4.2.1. Strategic advantages of nanozyme-based fluorescent detection methods

3 The strategic benefits of nanozyme-based fluorescence detection approaches stem from the
 4 synergistic integration of fluorescence technology and the distinctive characteristics of
 5 nanozymes, which are crucial for attaining significant enhancements in pesticide detection.
 6 These advantages extend beyond the general benefits of nanozymes in fluorescence detection;
 7 they encompass distinctive characteristics in the modulation, amplification, generation, and
 8 detection of the fluorescent signals, providing substantial performance improvements over
 9 conventional fluorescent or colorimetric techniques. A major strategic advantage is the
 10 capacity to attain exceptionally high sensitivity via catalytic signal amplification. Traditional
 11 fluorescence detection approaches typically depend on a 1:1 interaction between a target
 12 molecule and a fluorophore; however, in nanozyme-based systems, a single target molecule
 13 can influence the catalytic efficiency of a nanozyme to transform multiple fluorescent substrate
 14 molecules, thereby offering a natural signal amplification phenomenon. For instance, during
 15 the reaction in which a peroxidase-like nanozyme oxidizes a non-fluorescent substrate such as
 16 terephthalic acid (TA) to yield highly fluorescent 2-hydroxyterephthalic acid (2-HTA),



1 inhibition of the enzyme by one specific pesticide molecule results in the diminished
2 production of numerous fluorescent molecules, thereby attaining a significantly elevated signal
3 amplification ratio.¹¹⁴ This catalytic amplification significantly enhances the signal-to-noise
4 ratio, facilitating the precise detection of trace levels of pesticides in complex matrices.
5 Moreover, the catalytic efficacy of nanozymes facilitate enhanced signal amplification by fine-
6 tuning parameters such as temperature, pH, and reaction duration, a distinctive advantage
7 unattainable with conventional fluorescence techniques. A study involving Cu-BH nanozymes
8 indicated that molecules improved the catalytic activity of the nanozyme, intensifying the
9 fluorescent signal and attaining a remarkably low detection limit. This catalytic signal
10 amplification facilitates a linear enhancement in fluorescence intensity, markedly enhancing
11 the accuracy and precision of quantitative evaluation.¹¹⁰

12 Nanozyme-based fluorescence detection approaches offer distinct advantages for real-time
13 monitoring of pesticides. Because the fluorescent signal can be assessed instantly and the
14 nanozyme's catalytic reaction proceeds continuously, which enables real-time monitoring of
15 temporal variations in pesticide concentration. This is a significant advantage compared to
16 traditional detection methods that only provide static end-point analysis and can be utilized to
17 monitor pesticide degradation processes or their behavior in the environment. Furthermore, by
18 analyzing the nanozyme's catalytic reaction kinetics, information about the pesticide's
19 inhibition mechanism can be obtained, providing valuable data for distinguishing between
20 different types of pesticides or their mechanisms of action. The rapid response time of
21 fluorescence detection is a decisive advantage in urgent food safety inspections and on-site
22 monitoring, with most systems providing results within a few to ten minutes.¹¹⁵ In a glyphosate
23 detection system using Co_3O_4 nanozymes, for instance, detection is completed within 10



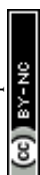
1 minutes, and the resulting color change remains stable for over 20 minutes, facilitating easy
2 visual discrimination. This rapid response and signal stability meet the requirements for point-
3 of-care testing and are also suitable for screening large volumes of samples.¹¹⁶

4 A unique advantage of nanozyme-based fluorescence detection is the ability to design a
5 selective fluorescence response for specific pesticides by engineering the nanozyme's structure
6 and surface functionalization. This is achieved through a ternary interaction system involving
7 the fluorophore, the nanozyme, and the target molecule, where high specificity can be
8 implemented by precisely controlling the properties of each component. For example, in
9 aptamer-functionalized gold nanoparticles, binding with a specific pesticide (acetamiprid)
10 selectively modulates the peroxidase-like activity of the nanoparticles, resulting in a
11 fluorescent response that distinguishes it from other structural analogs. Furthermore, by
12 controlling the pore structure or surface chemistry of the nanozyme, size selectivity or chemical
13 selectivity can be achieved, enabling the selective detection of a target pesticide within a
14 complex matrix.¹¹⁷

15

16 **4.2.2 Classification of nanozyme-based fluorescent sensors for pesticide detection**

17 The operating principles of nanozyme-based fluorescent pesticide detection systems can be
18 classified according to the fundamental interaction among the target analyte (pesticide), the
19 nanozyme catalyst, and the fluorescence signal-generating component. Nanozyme-based
20 fluorescence pesticide sensors primarily rely on catalytic activity modulation, reaction by-
21 product mediated fluorescence, or direct modulation of intrinsic fluorescence. This mechanistic
22 classification forms the basis for determining key performance indicators such as sensor
23 sensitivity, selectivity, and response speed, and it provides a theoretical foundation for

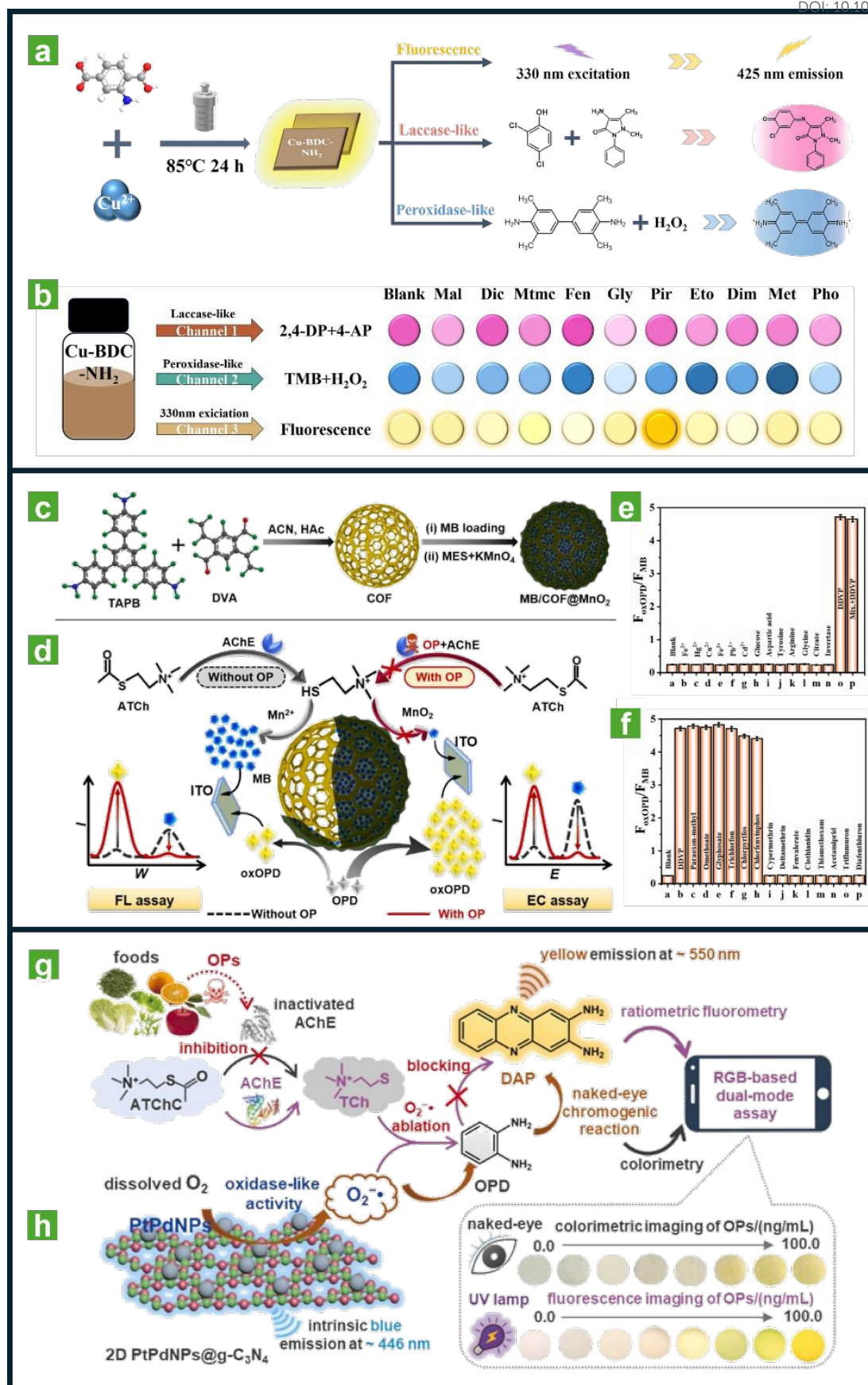


1 designing sensor platforms optimized for specific applications. To date, reported studies can
2 be broadly divided into three dominant strategies. The first is a classic approach that relies on
3 the indirect modulation of biological enzyme activity. In this method, the pesticide inhibits the
4 activity of a specific biological enzyme, and the nanozyme acts as a signal transducer to convert
5 this event into a fluorescent signal.^{55, 104, 105} The second is the direct degradation of pesticides
6 via the intrinsic catalytic activity of the nanozyme. This is a substantial approach, particularly
7 mimicking the function of a phosphatase to directly hydrolyze the phosphate ester bonds of
8 OPs.^{64, 108} Further, an in-depth discussion of each classification will be presented below.

10 **4.2.2.1. Enzyme activity inhibition by pesticides**

11 Indirect detection through the modulation of biological activity of enzymes is the most
12 thoroughly investigated and recognized method for detecting carbamate and organophosphorus
13 pesticides. This methodology fundamentally relies on the biochemical mechanism through
14 which these pesticides selectively impede the catalytic function of AChE, a crucial enzyme in
15 the central nervous system. To convert this change in AChE activity into a quantifiable
16 fluorescent signal, the sensor system employs a sophisticated enzymatic cascade reaction that
17 includes a nanozyme. This approach directly reflects the *in vivo* toxicity mechanism of the
18 pesticides, giving the detection results high biological significance. Furthermore, the multi-step
19 enzymatic and nanozyme catalytic reactions provide inherent signal amplification, making it
20 possible to achieve the high sensitivity required for trace analysis.^{3, 118}





1 **Fig. 3.** Pesticide detection through enzyme activity inhibition mechanism. (a) Fluorescent Cu-
2 BDC-NH₂ nanozyme demonstrating multienzyme-mimicking capabilities. (b) Implementation
3 of a multi-responsive sensor array using Cu-BDC-NH₂ for the classification of pesticides
4 across diverse concentration levels. The sensing platform is constructed by integrating laccase-
5 mimicking, peroxidase-mimicking, and fluorescence-based detection pathways. Reproduced
6 from reference¹⁰⁴ with permission from Elsevier, copyright 2024. (c) Schematic outlining the
7 fabrication process of the Multifunctional COF composite (MCM). (d) Conceptual illustration
8 of the dual-ratio enhanced FL/EC dual-mode sensor for OP analysis via target-regulated signal
9 modulation. (e) FL ratio response of the dual-modal sensing platform toward a range of
10 potential interferents (a–p: blank, Fe²⁺, Hg²⁺, Cu²⁺, Fe³⁺, Pb²⁺, Cd²⁺, glucose, aspartic acid,
11 tyrosine, arginine, glycine, citrate, invertase, dichlorvos (DDVP), and their mixture) (f)
12 Evaluation of FL ratios across various OPs and non-OP substances to demonstrate sensing
13 selectivity. Reproduced from reference¹⁰⁵ with permission from Elsevier, copyright [2024]. (g)
14 Design of a smartphone-integrated portable sensor based on 2D fluorescent oxidase-like
15 PtPdNPs@g-C₃N₄ nanozymes for dual-mode (colorimetric and fluorescence) imaging of
16 residual OPs. (h) Schematic illustration describing the oxidase-mimicking catalytic mechanism
17 of the 2D PtPdNPs@g-C₃N₄ nanozyme. Reproduced from reference⁵⁵ with permission from
18 Elsevier, copyright 2023.

19

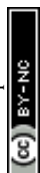
20 The most representative system is a cascade reaction involving AChE and a peroxidase-like
21 nanozyme. The operating principle of this system can be described in the following steps: (i)
22 In the absence of a pesticide, AChE hydrolyzes its substrate, acetylcholine or acetylthiocholine,
23 into choline or thiocholine. (ii) The generated choline is then oxidized by choline oxidase to



1 produce betaine and hydrogen peroxide. (iii) Finally, a nanozyme with peroxidase-like activity
2 uses H_2O_2 as an oxidant to convert a fluorogenic substrate (e.g., Amplex red and terephthalic
3 acid) into a product that emits strong fluorescence (e.g., Resorufin and 2-hydroxyterephthalic
4 acid).¹¹⁹ However, if the target pesticide is present in the system, the active site of AChE is
5 irreversibly or reversibly blocked. This reduces the production of choline, which in turn leads
6 to a sequential decrease in H_2O_2 generation and, ultimately, a reduction in the fluorescence
7 signal's intensity. Therefore, the degree of fluorescence decrease shows a direct correlation
8 with the concentration of the pesticide in the sample. In these signal transduction processes,
9 materials such as Fe_3O_4 , V_2O_5 , noble metal nanoparticles (Au NPs, Pt NPs), and various
10 carbon-based nanomaterials (graphene oxide, CQDs, etc.) have been successfully applied as
11 peroxidase-like nanozymes.^{67, 120} Additionally, in 2024, Song et al. presented an approach that
12 analyzes enzyme activity inhibition patterns by utilizing multiple signals from a single material.
13 In this study, a copper-based nanozyme (Cu-BDC-NH₂) with simultaneous laccase-like,
14 peroxidase-like, and fluorescent properties was used. The detection mechanism of this sensor
15 is based on the principle that different types of pesticides affect these three signals (two enzyme
16 activities, one fluorescence) to varying degrees, generating a unique 'inhibition pattern' or
17 'signal fingerprint' (Fig. 3a). Through this multi-signal analysis, Song et al. successfully
18 distinguished various types of pesticides down to a concentration of 1 $\mu\text{g/mL}$. Furthermore,
19 when applied to real fruit and vegetable samples, the sensor demonstrated high reliability by
20 identifying unknown samples with 100% accuracy (Fig. 3b).¹⁰⁴

21

22 A 2024 study by Wen et al. applied an enzyme activity inhibition mechanism by developing
23 a multifunctional covalent organic framework composite loaded with methylene blue and



1 coated with an MnO₂ nanozyme (Fig. 3c). Normally, thiocholine, produced when the enzyme
2 AChE breaks down its substrate (ATCh), decomposes the MnO₂ coating, releasing the
3 methylene blue signal molecule from inside. However, when OPs inhibit the activity of AChE,
4 thiocholine is not produced, so the MnO₂ coating is maintained, acting as gate that blocks the
5 release of MB (Fig. 3d). The dual-mode sensor demonstrated exceptional selectivity and robust
6 anti-interference capability for OPs detection. High concentrations of diverse inorganic ions
7 (such as Fe²⁺, Hg²⁺, and Cu²⁺) and organic substances produced negligible variations in FL and
8 electrochemical (EC) signals, whereas the presence of DDVP triggered significant responses
9 (Fig. 3e). Furthermore, the platform's versatility was validated by its ability to accurately
10 distinguish six typical OPs from representative non-OP pesticides. This high specificity is
11 fundamentally attributed to the target-regulated, high-affinity inhibition of AChE by
12 organophosphates (Fig. 3f). This sensor can detect dichlorvos based on fluorescence signal
13 change, and achieved very low LOD of 0.083 ng/mL. Additionally, it showed excellent
14 recovery rates in real vegetable and fruit samples, ranging from 95.2% to 104%.¹⁰⁵ Shen et al.
15 applied the principle of enzyme activity inhibition by developing platinum-palladium
16 nanoparticles combined with carbon nitride nanosheets (PtPdNPs@g-C₃N₄) (Fig. 3g). This
17 nanozyme acts like an oxidase, converting surrounding oxygen into reactive oxygen species
18 (O₂[·]). Normally, the enzyme AChE breaks down a substrate to produce TCh, which then
19 removes the reactive oxygen species, preventing a fluorescence color change. However, when
20 OPs inhibit the activity of AChE, thiocholine is not produced, allowing the reactive oxygen
21 species to freely oxidize a substrate (OPD), which generates distinct fluorescence signals (Fig.
22 3h). This sensor demonstrated LOD for trichlorfon of 0.033 ng/mL via fluorescence methods,
23 respectively, and recorded a high recovery rate of 95.78% to 104.34% in real food samples.⁵⁵

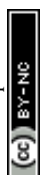


1 4.2.2.2. Byproduct mediated fluorescent detection

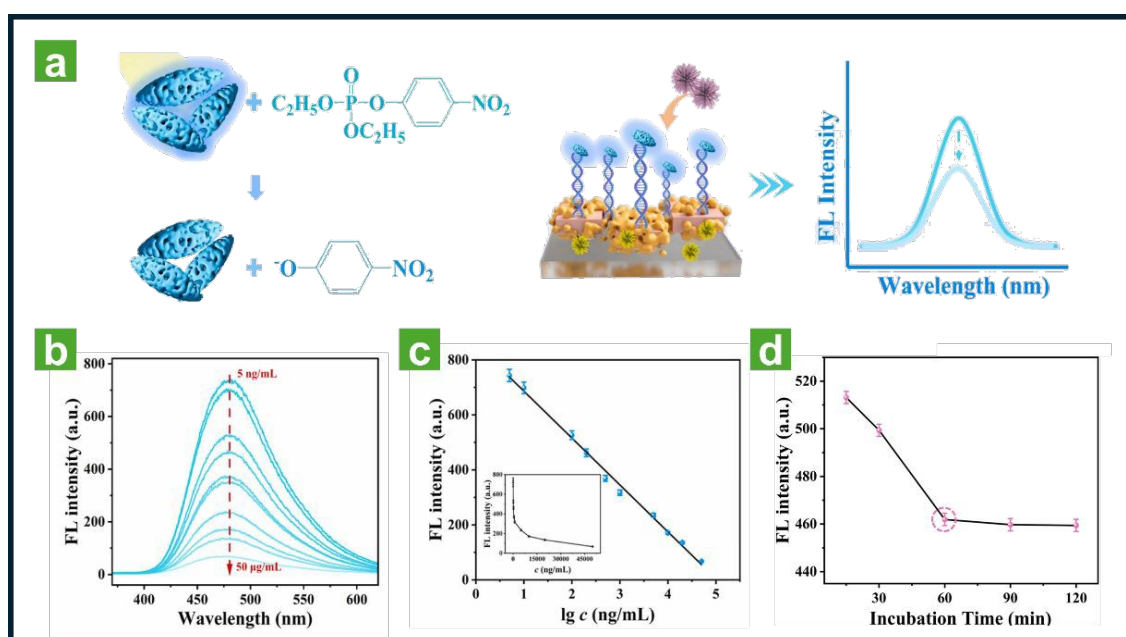
2 This approach utilizes the optical properties of the hydrolysis product. In this method, the
3 nanozyme functions as a biochemical converter, transforming a non-optical pesticide substrate
4 into a product with distinct optical characteristics. The most representative mechanism is
5 fluorescence quenching via the inner filter effect. For example, certain OPs, such as methyl
6 parathion, produce a chromophore called p-nitrophenol (PNP) upon hydrolysis. Under specific
7 pH conditions, PNP exhibits a distinct UV-visible absorption spectrum. If a separate
8 fluorophore (e.g., CQDs, organic dyes) whose excitation or emission wavelength overlaps with
9 PNP's absorption band is introduced into the system, the PNP generated by the nanozyme-
10 catalyzed reaction will absorb and extinguish the fluorophore's light. Consequently, a higher
11 initial pesticide concentration produces more PNP, leading to a greater degree of fluorescence
12 quenching. This relationship allows for the quantification of pesticides. A key advantage of
13 this method is that it does not require physical binding or complex energy transfer processes
14 between the nanozyme and the fluorophore, offering considerable versatility in system design.

15 121

16
17 In 2026, Zhang et al developed a nanozyme-based sensor utilizing a byproduct-
18 mediated fluorescent detection strategy for the sensitive monitoring of paraoxon. While the
19 platform was designed as a dual-mode system, its optical detection specifically relied on a
20 zirconium-based metal-organic framework (Zr-MOF), which served as a robust phosphatase-
21 like nanozyme with intrinsic aggregation-induced emission (AIE) properties. Focusing on the
22 fluorescence sensing mechanism, the approach utilizes a byproduct-mediated quenching
23 process. Upon target exposure, the biomimetic catalytic sites of the Zr-MOF hydrolyze



1 paraoxon to generate p-nitrophenol (p-NP). This generated byproduct acts as a strong quencher,
2 diminishing the intrinsic blue emission of the Zr-MOF via an Inner Filter Effect (IFE) (Fig.
3 4a). Based on this concentration-dependent fluorescence quenching, the platform yielded a
4 limit of detection (LOD) of 1.67 ng/mL across a broad linear range of 5 ng/mL to 50 μ g/mL
5 (Fig. 4b). By mapping these spectral changes, a corresponding calibration plot was established,
6 revealing a strong linear correlation ($R^2 = 0.995$) between the attenuated fluorescence intensity
7 and the logarithm of the paraoxon concentration (Fig. 4c). Furthermore, to evaluate its practical
8 applicability, the sensor was utilized to detect paraoxon in real-world samples. The
9 fluorescence mode provided highly reliable and accurate performance, showing recoveries
10 from 99.8% to 100.6% in tomato extracts and 99.5% to 100.4% in tap water matrices.
11 Additionally, optimization studies demonstrated clear fluorescence quenching after 60 minutes,
12 establishing this duration as the required incubation time for the sensor's optimal performance
13 (Fig. 4d).¹¹³



15



1 **Fig. 4.** Pesticide detection mechanism through byproduct mediated fluorescence detection. (a)
2 Diagrammatic of fluorescence sensing mechanism utilized byproduct-mediated quenching
3 process, (b) Fluorescence emission spectra obtained at varying paraoxon concentrations (from
4 5 ng/mL to 50 μ g/mL), (c) The corresponding calibration plot for the fluorescent assay, (d)
5 Optimization of incubation time for the fluorescence emission of the sensor. Reproduced from
6 reference¹¹³ with permission from Elsevier, copyright 2026.

8 **4.2.2.3. Modulation of intrinsic nanozyme fluorescence**

9 This approach is a self-reporting detection method that works by changing the nanozyme's
10 own fluorescence. The most ideal and streamlined system uses a self-reporting nanozyme,
11 which acts as both catalyst and fluorescent signal material at the same time. In this approach,
12 the nanozyme has intrinsic fluorescence, like protein-stabilized gold nanoclusters or certain
13 MOFs. When a pesticide molecule binds to the active site of one of these fluorescent
14 nanozymes and is hydrolyzed, it directly changes the nanozyme's electronic structure or surface
15 ligand environment. For instance, the binding of the substrate might activate a non-radiative
16 relaxation pathway, or the release of the product could alter the charge state of the fluorophore
17 core. This local environmental shift immediately leads to a decrease or increase in the
18 nanozyme's own fluorescence intensity, allowing for pesticide detection without any external
19 source. This self-reporting mechanism is a game-changer because the reaction and signal
20 generation are directly linked within a single entity, leading to fast reaction times and a very
21 simple system design.^{122, 123}

22



1 A key example of the direct pesticide quantification method is the 66-IS-Zn MOF nanozyme
2 developed by Huang *et al.* (2025). This innovative approach introduces biomimetic zinc active
3 sites into a UiO-66 framework to directly hydrolyze pesticides. This nanozyme directly cleaves
4 the P-O bond of pesticides, producing phosphate ions and alcohol derivatives, all without the
5 need for biological enzymes like AChE. During this process, a change in the ligand-to-metal
6 charge transfer within the MOF structure leads to a change in fluorescence intensity (Fig. 5a).
7 The system achieved a detection limit of 0.1-10 nM for methyl parathion, chlorpyrifos, and
8 diazinon, with a response time of less than 10 minutes. A 3-channel sensor array even makes
9 it possible to distinguish between multiple pesticides at the same time. Its practicality was
10 confirmed with a recovery rate of over 90% in real agricultural samples, and it showed
11 remarkable stability, maintaining over 90% of its activity after more than 100 reuses. This is a
12 perfect example of an all-in-one platform where the nanozyme both degrades the pesticide and
13 generates the detection signal, showcasing the core advantages of the direct degradation
14 method with simple, enzyme-free system and broad operating conditions (Fig. 5b).¹⁰³ Zhu et
15 al. (2025) proposed a method for detecting a specific OPs, paraoxon, utilizing a Zr-based MOF
16 termed an 'AIEzyme.' This AIEzyme hydrolyzes paraoxon to produce yellow p-nitrophenol.
17 This product induces a dual signal by simultaneously changing the solution's color (a
18 colorimetric 'On' signal) while quenching the AIEzyme's own strong blue fluorescence via an
19 inner filter effect (a fluorometric 'Off' signal) (Fig. 5c). The sensor exhibited excellent detection
20 limits of 0.178 μM (colorimetric) and 0.195 μM (fluorometric), demonstrating its reliability
21 with high recovery rates ranging from 96.1% to 115.0% in real samples (Fig. 5d, e). While this
22 study is very similar to the previously discussed research case, its focus is on the precise
23 quantitative analysis of a single pesticide, paraoxon, rather than on discriminating and
24 identifying multiple pesticides (Fig. 5f).⁶⁴



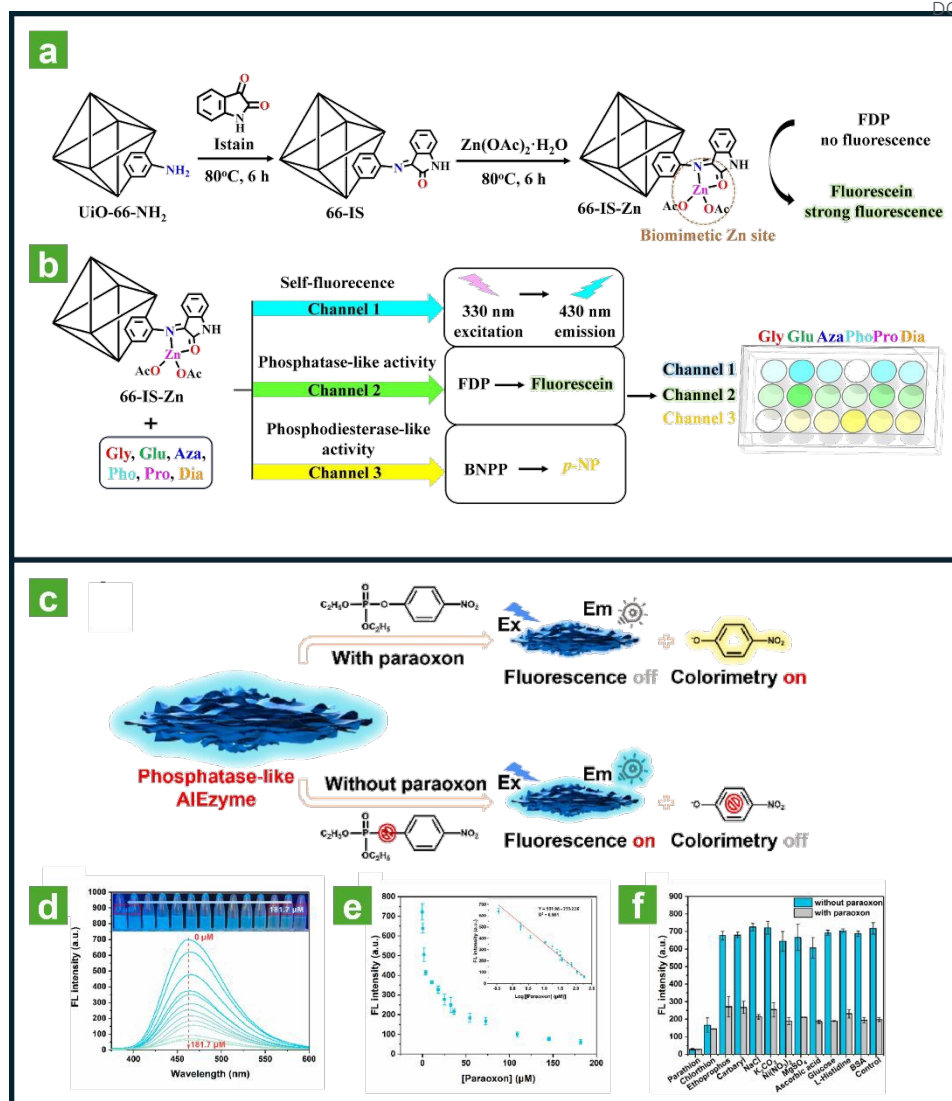


Fig. 5. Pesticide detection mechanism through modulation of intrinsic nanozyme fluorescence mechanism. (a) Diagram illustrating the synthesis of 66-IS-Zn via the covalent integration of biomimetic Zn centers into UiO-66-NH₂ and its catalytic role in FDP dephosphorylation. (b) Schematic representation of a 66-IS-Zn-based nanozyme sensor array designed for the differentiation of various OPs. Reproduced from reference¹⁰³ with permission from Elsevier, copyright 2025. (c) Detection mechanism of a dual-mode colorimetric and fluorometric platform for paraoxon sensing utilizing an AIEzyme. (d) Fluorescence emission profiles of the AIEzyme system in response to varying concentrations of paraoxon. (e) Calibration plot



1 showing the linear correlation between FL intensity and the logarithmic concentration of
2 paraoxon. (g) Selectivity evaluation of the proposed fluorometric method against diverse
3 interfering species. Reproduced from reference⁶⁴ with permission from Elsevier, copyright
4 2025.

6 **4.3. Nanozymes for the electrochemical detection of pesticides**

7 Electrochemical detection methods, owing to their outstanding advantages such as high
8 sensitivity, rapid response time, cost-effectiveness, and potential for miniaturization, have
9 established themselves as one of the most promising platforms for developing portable field-
10 oriented pesticide monitoring systems.¹²⁴ In particular, the technology of integrating
11 nanozymes onto electrode surfaces has opened new horizons in this field. By catalytically
12 amplifying the electrochemical signals generated from the target pesticide or its byproducts,
13 nanozymes dramatically enhance the sensor's analytical performance and play crucial role in
14 lowering the detection limit to ultra-trace levels. Traditionally, pesticide residue analysis has
15 relied on sophisticated instrumental techniques like GC-MS and LC-MS.¹²⁵ While these
16 methods boast high accuracy, capable of analysis at sub-ppb (parts per billion) levels and
17 simultaneous quantification of multiple components, they demand expensive equipment,
18 skilled analysts, and complex, time-consuming sample preparation procedures. This has posed
19 a critical limitation for on-site applications requiring immediate results, such as the rapid
20 inspection of agricultural products before shipment or extensive environmental pollution
21 monitoring. To bridge this gap, research on biosensors utilizing natural enzymes as
22 biorecognition elements began in the 1960s.¹²⁶ In particular, sensors using AChE were widely
23 employed for detecting organophosphate and carbamate pesticides¹²⁷; however, the inherent



1 instability of the enzyme itself was the greatest obstacle to commercialization. Natural enzymes
 2 are highly sensitive to changes in the external environment, such as temperature, pH, and ionic
 3 strength, causing their tertiary structure to denature easily and lose activity. Furthermore,
 4 persistent issues were raised regarding high production costs, stemming from the complexity
 5 of extraction and purification from biological systems, and the difficulty in achieving sensor
 6 reproducibility due to large batch-to-batch variations in activity.^{128, 129} It was at this
 7 convergence that 'nanozymes' artificial enzymes based on inorganic nanomaterials emerged as
 8 a powerful alternative, following the 2007 discovery that Fe₃O₄ nanoparticles exhibit
 9 peroxidase-like catalytic activity.¹³⁰ Nanozyme-based electrochemical sensors operate on the
 10 principle of converting a biochemical reaction into a measurable electrical signal, such as
 11 current, potential, or impedance. The nanozyme, immobilized on the working electrode,
 12 functions as the core recognition and signal amplification element.¹³¹ Its enzyme-like catalytic
 13 activity towards a specific substrate which is either consumed or generated in the presence of
 14 the target pesticide forms the basis of detection.¹³² This section begins by discussing the
 15 intrinsic advantages that nanozymes offer to electrochemical pesticide sensors. It then provides
 16 an in-depth analysis of the main detection mechanisms the enzyme inhibition-based and direct
 17 electrocatalytic methods and concludes with a thorough discussion on the latest design
 18 strategies to maximize sensor performance. Table 3 shows the comparative analysis of
 19 nanozyme-based electrochemical sensors for pesticide detection.

20
 21 **Table 3. Comparative analysis of nanozyme-based electrochemical sensors for pesticide**
 22 **detection**

Nanozyme	Pesticides	Enzyme activity	LOD	Reaction time	Recovery rates	Ref.
----------	------------	-----------------	-----	---------------	----------------	------



View Article Online
DOI: 10.1039/D5TA09692H

Ni-SPNC	Atrazine	Peroxidase	2.3 fg/mL	30 min	94.9%-98.5 %	133
Fe ₃ O ₄ /Cu-MOF	Thiram	Laccase	36.1 pg/mL	1 min	98.42%-105.44%	134
FePrNS, NiPcNS	Paraoxon	Peroxidase	1.35 μg/mL	15 min	98.4 %	135
Pd@Pt NP	Atrazine, Acetochlor	Peroxidase	1 ng/mL	2 min	97.0%-109%	136
MB/COF@MnO ₂ , MCM	Dichlorvos	Oxidase	26 pg/mL	30 min	97.8 %-105 %	105
Au@MnO _{2-x}	Ethion, Omethoate, Diazinon, Chlorpyrifos methyl, Trichlorfon	Oxidase	39 pg/mL	10 min	95%-105%	137
Pt/Co ₃ O ₄ nanoflowers	Thiram	Oxidase, Peroxidase, Catalase, Superoxide dismutase	15.63 ng/mL	15 min	95.33%-101.60%	77
NiCoFeS/rGO	Trichlorfon	Peroxidase	9.74fg/mL	20 min		138
MnNS	Paraoxon	Oxidase, Peroxidase	0.025 ng/mL	20 min		89
CeO ₂ nanozyme	Methyl-paraoxon	Phosphatase	14.83 ng/mL	60 min	116.84% ± 1.4 2%	139
NiO nanoplatelets	Parathion	Oxidase	6.99 ng/mL		103 ± 2.13%	140
SA-Fe-NZ	Ethyl parathion, Dichlorvos, Omethoate	Peroxidase	1.03 fg/mL	3 min	99.14 %-104.27 %	141
COF-OMe@Valine-CeO ₂	Methyl paraoxon	Phosphatase	2.72 ng/mL	5 min	97.46–111.10 %	142
ZrO ₂ @ZIF-90	Methyl parathion	Phosphohydrolyse	0.14 pg/mL	4 min	89.25%-102.78%	143
CuO _x /GDYO	Chlorpyrifos	Peroxidase	0.20 ng/mL	20 min	96.62%–103.25 %	144



Cobalt-Doped Ti ₃ C ₂ MXene	Paraoxon, Chlorpyrifos, Methidathion, Phoxim, Malathion, Profenofos, Carbaryl, Carbofuran	Peroxidase	0.02 ng/mL	30 min	97.4%-103.3%	145
Gold nanorods	Malathion, Methyl parathion, Chlorpyrifos, Parathion, Dichlorvos	Peroxidase	8.1 pg/mL	10 min	100%-110%	146
Au@Pt nanozyme	Parathion, Triazophos, Chlorpyrifos	Peroxidase	3.95 fg/mL	30 min	71.26%-117.47%	147

1

2

3 4.3.1. Strategic advantages of nanozyme-based electrochemical detection methods

4 The fusion of the unique catalytic properties of nanozymes with electrochemical
 5 transduction technology creates a significant synergistic effect in the field of pesticide detection.
 6 This goes beyond the mere sum of the two technologies, offering four key strategic advantages
 7 that can overcome the fundamental limitations of existing sensing paradigms. First is the
 8 intrinsic signal amplification and enhanced sensitivity. The sensitivity of an electrochemical
 9 sensor is ultimately determined by the magnitude of the Faradaic current generated at the
 10 electrode surface. Like natural enzymes, nanozymes follow Michaelis-Menten kinetics and can
 11 rapidly convert (turnover) numerous substrate molecules into products at a single catalytic site.
 12 This catalytic amplification mechanism enables a single molecule of the target analyte to
 13 trigger the generation or consumption of thousands, even tens of thousands, of
 14 electrochemically active molecules, thereby intrinsically amplifying the measurement
 15 signal.^{137, 148} For example, a nanozyme with peroxidase-like activity can generate a massive
 16 current by oxidizing or reducing numerous redox mediators (e.g., TMB, ABTS), a process
 17 mediated by a single molecule of hydrogen peroxide (H₂O₂).¹⁴⁹ While the turnover number of
 18 a natural enzyme like horseradish peroxidase (HRP) reaches several hundred per second, well-
 19 designed single-atom nanozymes or certain metal oxide nanozymes exhibiting comparable or



1 sometimes even superior catalytic efficiency. In particular, the K_m (Michaelis constant) value
2 of a nanozyme indicates its affinity for a substrate, and some nanozymes have demonstrated
3 lower K_m values than natural enzymes, proving they can operate efficiently even at lower
4 substrate concentrations.^{150, 151} Because of this inherent amplification capability, nanozyme-
5 based electrochemical sensors have the potential to detect ultra-trace levels of pesticides down
6 to the femtomolar (fM) or even attomolar (aM) range levels difficult to achieve with traditional
7 methods without the need for external, complex amplification systems (e.g., polymerase chain
8 reaction, rolling circle amplification). Second is the maximization of interfacial electron
9 transfer rates. An efficient electrochemical sensor relies on rapid and seamless electron transfer
10 between the catalyst's active site and the electrode surface.¹³⁴ In the case of natural enzymes,
11 the active site (e.g., a heme group) is often buried deep within a large protein structure
12 (typically several to tens of nanometers in size), making direct electron transfer (DET) with the
13 highly inefficient electrode. This is because the distance over which electrons can tunnel is
14 typically limited to 1-2 nm. Consequently, diffusible mediators such as ferrocene derivatives
15 or $[\text{Fe}(\text{CN})_6]^{3-/4-}$ are often required to facilitate electron transfer, which increases the sensor's
16 complexity and response time.^{28, 152} In contrast, nanozymes particularly nanozyme composites
17 integrated with 2D conductive nanomaterials like graphene, carbon nanotubes (CNTs), and
18 MXene create an ideal nano-environment at the electrode interface. These conductive
19 nanomaterials act as "electron wires," facilitating the rapid transfer of electrons generated at
20 the nanozyme's catalytic center to the electrode with minimal resistance.^{27, 28} This, in turn,
21 lowers the overpotential of the electrode reaction and reduces the charge transfer resistance
22 (R_{ct}), ultimately leading to higher sensitivity at lower potentials and faster response times in
23 the millisecond range.



1 Third is miniaturization, portability, and field applicability. Electrochemical analysis
2 systems, which intrinsically consist of a potentiostat and electrodes, have a simpler
3 configuration and are more easily miniaturized compared to other optical or mass spectrometry
4 instruments. The introduction of nanozymes further enhances this practicality. Natural
5 enzymes are typically irreversibly denatured at temperatures above 40-50°C and exhibit
6 optimal activity only within a narrow pH range. Therefore, enzyme-based sensors must be
7 operated under strict conditions and require frozen storage at temperatures below -20°C for
8 long-term preservation. In contrast, metal oxide, noble metal, and carbon-based nanozymes,
9 being intrinsically robust inorganic materials, maintain high stability at high temperatures
10 (above 80°C), across a wide pH range (2-11), and even in organic solvents like acetonitrile and
11 ethanol.¹⁵³ Thanks to this robustness, they can exhibit stable performance for several months
12 at room temperature without requiring refrigeration, making them highly suitable for the
13 development of field-oriented, disposable screen-printed electrode (SPE)-based sensors. By
14 combining the excellent stability of nanozymes with the simplicity of electrochemical detection,
15 it becomes possible to realize true point-of-care testing (POCT) devices that can be easily used
16 by non-experts in locations such as agricultural fields, food processing facilities, and
17 environmental water quality monitoring sites.¹⁵⁴⁻¹⁵⁶ Fourth is cost-effectiveness and ease of
18 mass production. The production of natural enzymes is expensive, requiring complex
19 biological expression (e.g., in *E. coli*, yeast) and multi-step purification processes (e.g.,
20 chromatography). Furthermore, large batch-to-batch variations in activity make it difficult to
21 ensure sensor reproducibility.^{128, 129, 135} In contrast, nanozymes can be produced in large
22 quantities and with high reproducibility from inexpensive precursors (e.g., FeCl₃, HAuCl₄, Ce
23 (NO₃)₃) through well-established chemical synthesis methods such as hydrothermal synthesis,
24 co-precipitation, and vapor deposition.^{146, 147} For instance, Fe₃O₄ nanozymes can be easily



1 synthesized on a gram-scale or larger through a simple method of reducing iron salts in an
2 aqueous solution, and their cost is merely a fraction as little as one-thousandth of that of
3 purified HRP enzyme.^{157, 158} This economy and productivity enable the mass production of
4 disposable sensor strips, which can lower the per-unit cost. This significantly lowers the cost
5 barrier for establishing extensive monitoring networks.¹⁵⁹ This serves as a particularly crucial
6 advantage in budget-constrained applications, such as food safety management in developing
7 countries or wide-area environmental surveillance.

9 **4.3.2. Classification of nanozyme-based electrochemical sensors for pesticide detection**

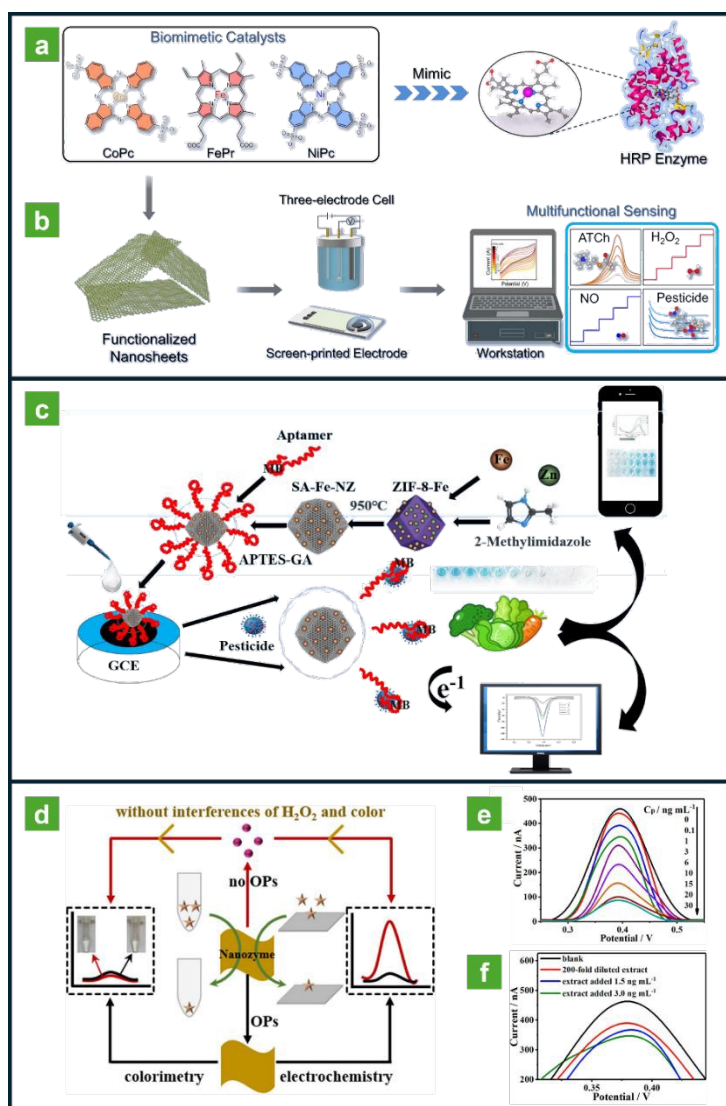
10 The advantages of nanozyme-based electrochemical sensors stem from the flexibility to
11 implement various detection mechanisms. This allows for the selection of an optimal sensor
12 design strategy tailored to the specific type of pesticide, the required sensitivity and selectivity,
13 and the application environment. Nanozyme-based electrochemical pesticide sensors primarily
14 operate through catalytic activity modulation, direct electrocatalytic oxidation or reduction
15 processes, or recognition-element mediated signal modulation, where interactions between
16 pesticides and the nanozyme surface influence the electrochemical response of redox probes or
17 electrode interfaces. In this section, nanozyme-based electrochemical pesticide sensors are
18 classified into three core detection mechanisms, (i) signal-suppression sensors via catalytic
19 activity inhibition, (ii) signal-generation sensors via direct electrocatalysis, and (iii) signal
20 modulation sensors based on specific molecular recognition. This section aims to provide an
21 in-depth analysis of the scientific principles, application examples, and technical limitations of
22 each approach.^{142, 160}

23



1 4.3.2.1. Signal-suppression sensors via catalytic activity inhibition

2 The signal-suppression approach is one of the most intuitive and widely studied methods for
 3 nanozyme-based sensors. The core principle of this mechanism involves the target analyte (the
 4 pesticide) inhibiting the activity of a catalyst—either a natural enzyme or a nanozyme within the
 5 system. This inhibition leads to a decrease in the electrochemical signal from its normal state,
 6 in proportion to the pesticide's concentration. This method is particularly effective for the
 7 detection of organophosphorus and carbamate pesticides, whose inhibitory effects on specific
 8 enzymes are well-established^{138, 161}.



9

57



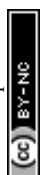
1 **Fig. 6.** Pesticide detection through “Signal-suppression” via catalytic activity inhibition
2 mechanism. (a) Schematic illustration of nanozymes functionalized with different biomimetic
3 catalysts (CoPc, FePr, NiPc). (b) Electrochemical detection of ATCh, H₂O₂, and nitric oxide
4 (NO) using functionalized nanozymes. Reproduced from reference¹³⁵ with permission from
5 Elsevier, copyright 2023. (c) Schematic illustration of single-atom Fe nanozyme based dual-
6 mode biosensor for multi-pesticide detection. Reproduced from reference¹⁴¹ with permission
7 from Elsevier, copyright 2024. (d) Schematic illustration of MnNS-based homogeneous
8 electrochemical sensor for OP detection. (e) DPV spectra of the TMB-MnNS-ATCh-AChE
9 platform in the presence of paraoxon at different concentrations. (f) DPV spectra of the TMB-
10 MnNS-ATCh-AChE platform in the extracting solution spiked with paraoxon. Reproduced
11 from reference¹⁶² with permission from American Chemical Society, copyright 2021.

12
13 As an example of a 'signal-suppression' sensor that uses a catalytic activity inhibition
14 mechanism, the paraoxon sensor developed by Niu et al. (2023) utilized a CoPcNS nanozyme
15 combined with the natural enzyme AChE to detect the paraoxon. The CoPcNS nanozyme is a
16 composite material created by combining sulfonated cobalt phthalocyanine (CoPc) with a base
17 of graphene and the conducting polymer PEDOT (Fig. 6a). The sensor operates based on a
18 cascade catalytic reaction between these two materials. First, the AChE enzyme hydrolyzes its
19 substrate, acetylthiocholine, to produce thiocholine. Subsequently, the CoPcNS nanozyme
20 efficiently oxidizes the generated thiocholine at the electrode, creating a strong baseline current
21 signal. However, if paraoxon is present in the sample, it inhibits the activity of the AChE
22 enzyme, which in turn prevents the production of thiocholine. As a result, the current signal
23 generated by the nanozyme proportionally decreases (Fig. 6b). Through this principle, the



1 developed portable sensor achieved a low limit of detection for paraoxon of 1.1 $\mu\text{g/L}$ and
2 demonstrated high reliability and applicability in complex environments by showing results in
3 real samples, such as tap water and cucumber, that were similar to those in standard
4 solutions.¹³⁵

5
6 In 2024, Wang et al introduced a dual-mode biosensor featuring a single-atom iron
7 nanozyme (SA-Fe-NZ), synthesized via high-temperature pyrolysis of Fe-doped ZIF-8, for the
8 multi-pesticide detection in vegetables (Fig. 6c). The platform utilizes the "toxic effect" of
9 target-aptamer complexes, which bind to the surface of SA-Fe-NZ and inhibit its peroxidase-
10 like activity, thereby preventing the catalytic oxidation of colorimetric substrates.
11 Simultaneously, the electrochemical mode records changes in DPV signals as methylene blue
12 (MB)-labeled aptamers undergo conformational changes upon binding to OPs, bringing the
13 signal molecules closer to the electrode surface for current amplification. This broad-spectrum
14 aptasensor achieved an exceptionally low LOD of 3.55 fM and a remarkably wide linear range
15 from 10^{-13} to 10^{-2} M, offering high accuracy through the mutual validation of optical and
16 electrochemical signals.¹⁴¹ Wu's research group detected pesticides using 2D manganese
17 dioxide nanosheets (MnNS) with dual oxidase- and peroxidase-like catalytic activities. This
18 study presented an innovative mechanism that completely excludes H_2O_2 and instead directly
19 utilizes dissolved oxygen as a coreactant. Fundamentally, the sensor relies on the AChE-
20 catalyzed hydrolysis of acetylthiocholine (ATCh) to generate thiocholine (TCh). The generated
21 TCh strongly inhibits the catalytic activity of the MnNS, thereby preventing the oxidation of
22 the electrochemical probe TMB and maintaining a high DPV signal (Fig. 6d). However, when
23 pesticides are present, AChE activity is inhibited, reducing TCh production. Consequently, the

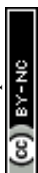
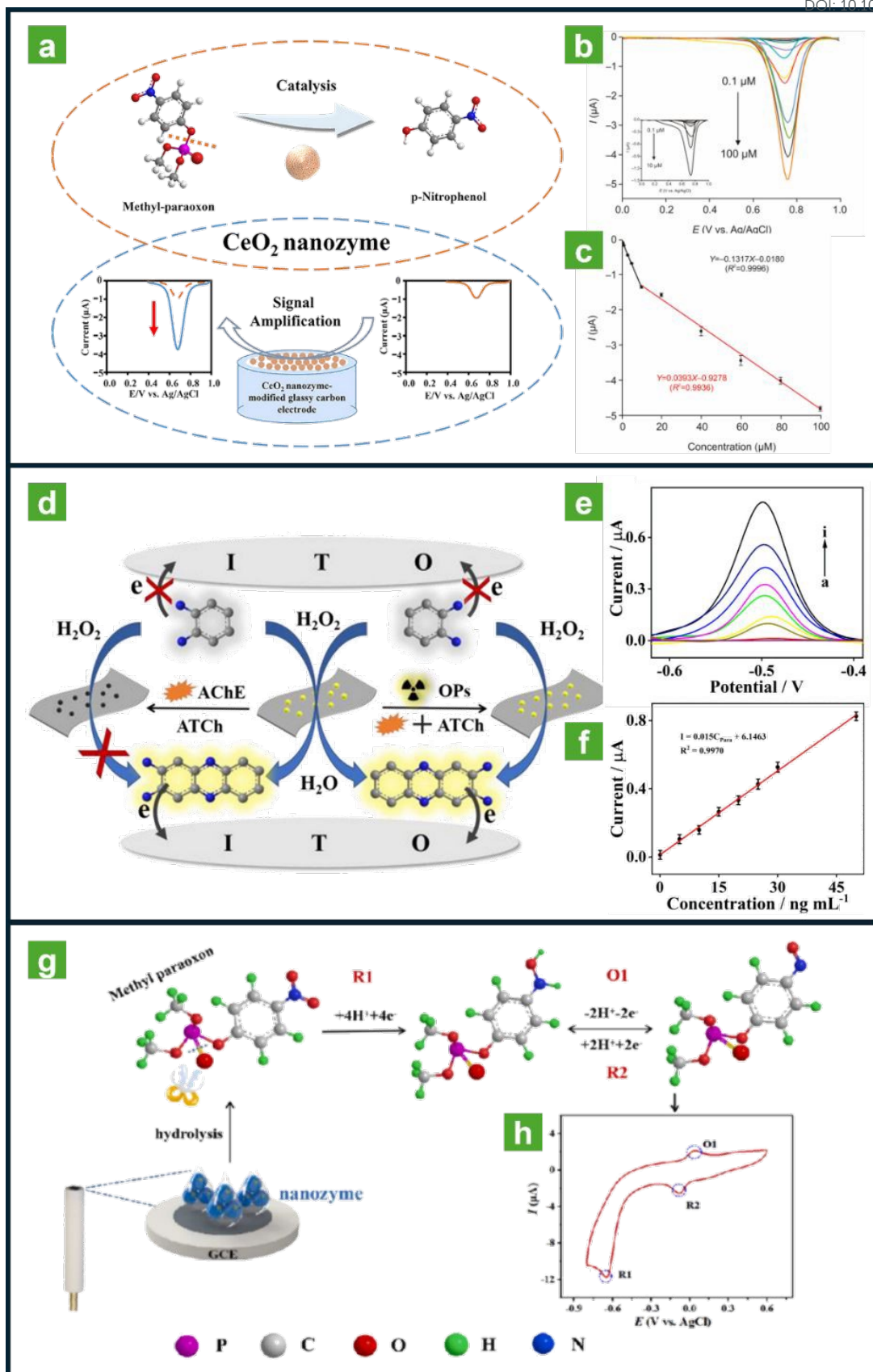


1 MnNS resumes the catalyzed oxidation of TMB, leading to a significantly declined DPV
2 current. At this point, nanozymes maximize the sensor's performance through two key roles
3 (Fig. 6e). Finally, when a pesticide inhibits AChE, no TCh is produced. Consequently, the
4 MnNS resumes the catalyzed oxidation of TMB, leading to a significantly declined DPV
5 current. This homogeneous electrochemical OP detection process based on the depressing
6 AChE activity enables the highly sensitive detection of the pesticide. The sensor demonstrated
7 high reliability, achieving a LOD of 0.025 ng/mL, excellent recoveries (99.72–102.41%), and
8 high precision with relative standard deviations under 2.52% (Fig. 6f).¹⁶²

10 4.3.2.2. Signal-generation sensors via direct electrocatalysis

11 As an alternative to inhibition-based methods, the 'signal-generation' approach is a versatile
12 strategy for detecting various types of pesticides to which the enzyme inhibition mechanism
13 cannot be applied. The core of this strategy involves using nanozymes as highly efficient
14 electrocatalysts to directly oxidize or reduce the pesticide molecules themselves or their
15 degradation byproducts, and then measuring the resulting Faradaic current. Consequently, the
16 magnitude of the electrochemical signal increases in direct proportion to the concentration of
17 the pesticide.



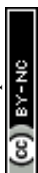


1 **Fig. 7.** Research trends in direct detection method. ‘Signal-generation’ sensors via direct
2 electrocatalysis mechanism. (a) Schematic illustration of the electrochemical sensing
3 mechanism for methyl-paraoxon detection utilized by CeO₂ nanozyme. (b) DPV responses of
4 the CeO₂ nanozyme-modified electrode interface across a gradient of MP concentrations (0.1–
5 100 μmol/L). (c) Quantitative calibration plot illustrating the linear dependence of the peak
6 current on the corresponding concentrations of MP.¹³⁹ (d) Schematic illustration of the
7 electrochemical sensing mechanism for paraoxon detection utilized by cobalt-doped Ti₃C₂
8 MXene nanozyme. (e) DPV responses of the common homogeneous electrochemical platform
9 across a gradient of paraoxon concentrations (0 to 50 ng/mL). (f) Linear calibration plot
10 correlating peak current intensity with the concentration of paraoxon. Error bars represent the
11 standard deviation derived from three independent replicates. Reproduced from reference¹⁴⁵
12 with permission from American Chemical Society, copyright 2022. (g) Schematic illustration
13 of the electrochemical sensing mechanism for MP detection utilized by COF-Ome@Valine-
14 CeO₂ nanozyme. (h) CV spectrum of COF-OME@Valine-CeO₂ in PBS (0.1 M pH 7.0) with
15 addition of MP. Reproduced from reference¹⁴² with permission from Elsevier, copyright 2024.

17 In 2021, the Sun research group reported a novel electrochemical method for detecting
18 pesticides by using cerium oxide (CeO₂) as a nanozyme with the dual functions of catalytic
19 reaction and signal amplification. This nanozyme possesses activity similar to
20 organophosphorus hydrolase, which promotes the decomposition of MP and generates p-
21 nitrophenol (Fig. 7a). Furthermore, by immobilizing the CeO₂ nanozyme on the electrode
22 surface, it serves to amplify the electrochemical response of the p-nitrophenol product (Fig.
23 7b). This study used bifunctional nanozyme for pesticide detection. Operating on this principle,



1 the sensor provided a LOD of 0.06 $\mu\text{mol/L}$ and successfully quantified methyl parathion in
2 herb samples (Semen nelumbinis, Adenophora stricta, and Coix lacryma-jobi) with recoveries
3 from $80.91\% \pm 4.86\%$ to $116.84\% \pm 1.42\%$ (mean \pm SD, $n=3$) across various concentrations
4 (Fig. 7c).¹³⁹ Organophosphate, phenolic, carbamate, and aniline-based pesticides possess
5 electrochemically oxidizable functional groups within their molecules, such as phenolic
6 hydroxyl groups, carbamate esters, or amine groups. In 2022, Yu's research group developed
7 cobalt-doped 2D Ti_3C_2 MXene nanosheets (CMNSs) as a nanozyme with excellent peroxidase-
8 like activity. This nanozyme can oxidize a specific substance, o-phenylenediamine (OPD),
9 converting it into oxidized OPD (OPDOX), which generates a strong electrical signal (Fig. 7d).
10 Interestingly, the nanozyme's activity is strongly inhibited by thiocholine, the product of the
11 AChE enzyme reaction. Therefore, when the target pesticide first inhibits the activity of AChE,
12 the production of thiocholine which would have interfered with the nanozyme is prevented.
13 This allows the nanozyme to become active, resulting in a strong signal. In fact, the study
14 observed that as the concentration of pesticides such as paraoxon, chlorpyrifos, and
15 methidathion increased, the oxidation peak current of OPDOX grew proportionally stronger
16 (Fig. 7e). In other words, this research successfully implemented a high-sensitivity sensor
17 based on a 'turn-on' mechanism, where the presence of the pesticide triggers the generation of
18 the signal molecule. The sensor exhibited both high sensitivity, with a LOD of 0.02 ng/mL,
19 and high accuracy, demonstrated by recovery rates of 97.4% to 103.3% (Fig. 7f).¹⁴⁵ In 2024,
20 Zhang et al. developed an ultrasensitive electrochemical sensor based on phosphatase-like
21 COF-OMe@Valine-CeO₂ nanozymes for the detection of organophosphorus pesticides,
22 specifically methyl paraoxon (MP). In this nanozyme, the Ce(IV)/Ce(III) species serve as the
23 active catalytic sites to polarize and hydrolyze the P=O bond in MP, leading to the formation
24 of electroactive p-nitrophenol (p-NP). Furthermore, the biomimetic substrate-trapping pockets

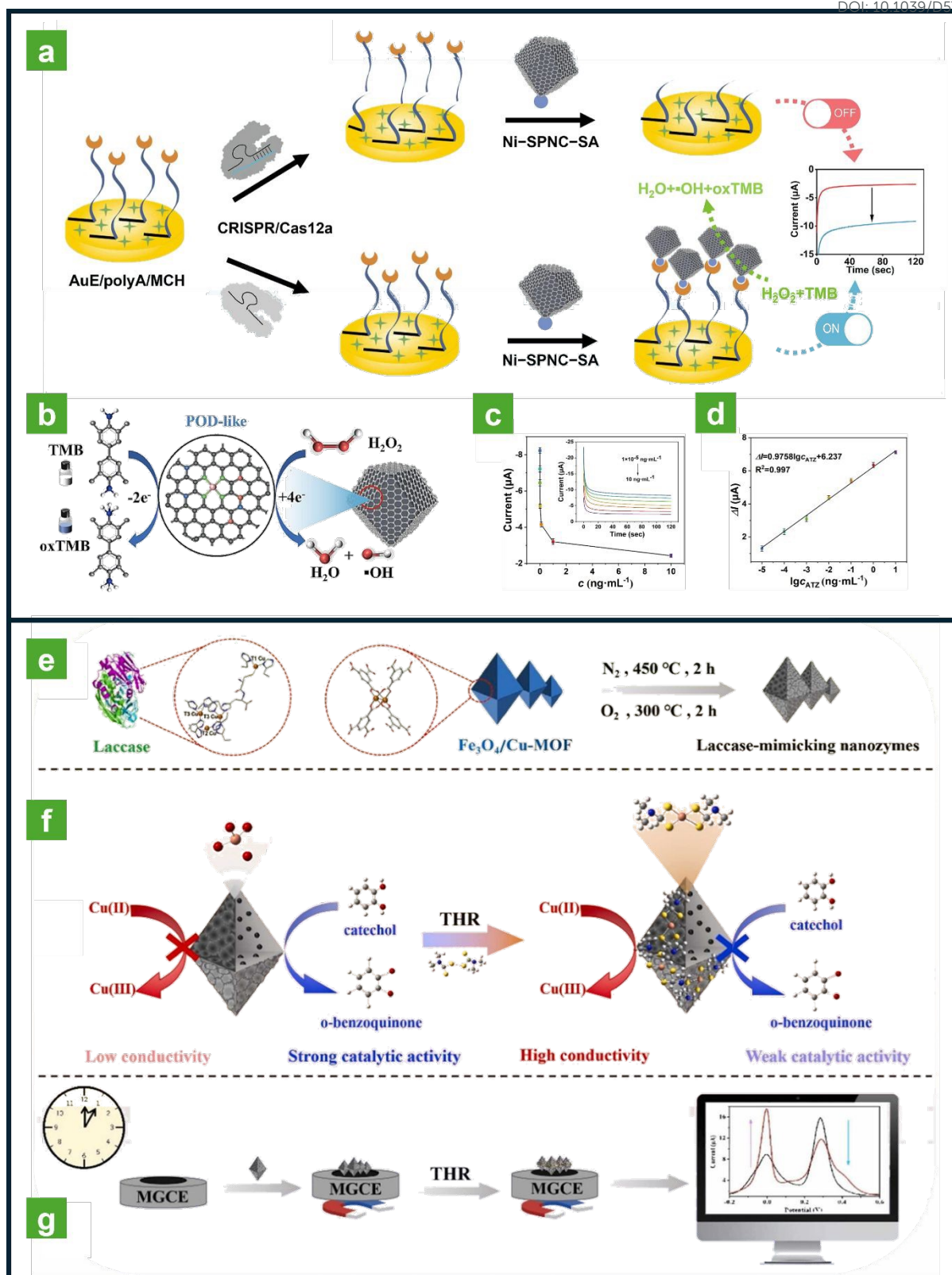


1 of the porous COF-OMe exhibit excellent adsorption performance to capture MP, which
2 synergistically promotes the hydrolysis process, while its covalent interface facilitates efficient
3 charge transfer for electrochemical signal amplification (Fig. 7g). Based on this catalytic
4 mechanism, the fabricated electrochemical sensor achieved a low limit of detection (LOD) of
5 0.011 $\mu\text{mol/L}$ for MP (Fig. 7h). The practicability of the sensor was successfully validated in
6 real samples (grapes and tap water), demonstrating satisfactory recoveries ranging from 97.46%
7 to 111.10%.¹⁴²

9 **4.3.2.3. Signal modulation sensors based on specific molecular recognition**

10 To overcome the limitations of the previous two mechanisms (e.g., reusability, selectivity),
11 sophisticated sensor designs that combine catalytic behavior of nanozymes with highly specific
12 molecular recognition elements, such as antibodies and aptamers, have recently been under
13 active investigation. This approach is based on the principle where the specific binding event
14 between a pesticide and a recognition element physically 'modulates' the nanozyme's activity
15 or the properties of the electrode interface to generate a signal. Aptamers are single-stranded
16 RNA or DNA sequences that bind to the specific molecules with high affinity. An aptamer
17 specific to a pesticide is immobilized on the nanozyme surface. In absence of pesticide, the
18 aptamer maintains a flexible structure, leaving the nanozyme's active site open or allowing a
19 redox probe to easily access the electrode. When the pesticide binds, the aptamer folds into a
20 three-dimensional structure (e.g., a hairpin, G-quadruplex), which can physically block the
21 nanozyme's active site to turn the signal off, or conversely, expose a previously blocked active
22 site to turn the electrochemical signal (Fig. 8a).¹³³





1

2 **Fig. 8.** Pesticide detection using signal modulation by specific molecular recognition

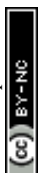
1 mechanism. (a) Schematic representation of the analytical workflow for atrazine (ATZ)
2 monitoring using the proposed biosensor. (b) Evaluation of the peroxidase-mimicking catalytic
3 performance of the synthesized SANs utilizing 20 mM TMB across various H₂O₂
4 concentrations. (c) Amperometric current-time (I - t) profiles of the electrochemical biosensor
5 in response to varying ATZ levels ranging from 1×10^{-5} to 10 ng/L. (d) Linear calibration
6 plot illustrating the relationship between the response current and the logarithmic concentration
7 of ATZ. Error bars indicate the standard deviation of triplicate measurements (n=3).
8 Reproduced from reference¹³³ with permission from Elsevier, copyright 2025 (e) Preparation
9 of the bioinspired laccase-mimicking nanozyme for thiram detection. (f) Schematic of the
10 ratiometric electrochemical sensing mechanism for thiram detection. (g) Label-free and rapid
11 detection of thiram utilizing the magnetic glassy carbon electrode platform. Reproduced from
12 reference¹³⁴ with permission from Elsevier, copyright 2024.

14 In the field of electrochemical biosensors, securing high selectivity for a target molecule is
15 a critical challenge. Recently, in 2025, Han's research group reported a noteworthy study that
16 solved this problem by utilizing an aptamer that specifically binds to ATZ, simultaneously
17 achieving ultra-high sensitivity through multi-stage amplification. The core working principle
18 of this sensor is as follows. In the presence of ATZ, the aptamer selectively binds with ATZ,
19 which in turn 'turns on' the activity of a DNAzyme motor. The activated DNAzyme motor
20 produces a large amount of trigger DNA, which then activates the CRISPR/Cas12a system to
21 induce a cascade of cleavage reactions. This demonstrates a sophisticated design where a single
22 selective recognition event by an aptamer triggers a massive signal amplification cascade. The
23 final signal of this complex amplification process is handled by a newly developed Ni-SPNC



1 single-atom nanozyme (Fig. 8b). This nanozyme, through co-doping with P and S, possesses
2 peroxidase-like activity superior to that of conventional materials. Finally, the amplification
3 reaction prevents the nanozyme from binding to the electrode surface, enabling quantitative
4 analysis through a mechanism where the signal decreases as the ATZ concentration increases.
5 The effectiveness of this simple mechanism is highlighted by the sensor's performance, which
6 includes a LOD of 2.3 fg/mL (Fig. 8c). Notably, a robust linear calibration plot was obtained
7 by plotting the response current against the logarithmic concentration of ATZ (Fig. 8d). It also
8 exhibited high accuracy, with recoveries of 94.9 - 98.5% and relative standard deviations of
9 1.18 - 5.26%.¹³³

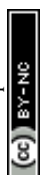
11 Building on the concept of signal modulation through molecular recognition,
12 bioinspired ratiometric sensors have been designed to achieve even higher accuracy. In 2024,
13 Geng et al. inspired by the multi-copper active centers ligated with amino acid residues in
14 natural laccase, a magnetic MOF (Fe₃O₄/Cu-MOF) was synthesized as a precursor for
15 electroactive nanozymes through a dual-step carbonization process (Fig. 8e). These nanozymes
16 utilize their abundant Cu(II) active sites to catalyze the oxidation of catechol into electroactive
17 o-quinone. When thiram is added, it forms a thiram-Cu(II) complex that specifically inhibits
18 the laccase-mimicking activity of the nanozyme, resulting in a significantly lower oxidation
19 current for catechol (Fig. 8f). Furthermore, the binding of thiram facilitates the internal electron
20 transfer of the nanozymes, generating a concomitantly higher signal for Cu(II). This
21 simultaneous decrease in the oxidation of catechol signal and increase in the Cu(II) signal
22 enables a sensitive ratiometric electrochemical response, which effectively cancels out
23 potential environmental interference. Finally, the intrinsic magnetic properties of the



1 nanocomposite allow for a label-less detection strategy using a magnetic glassy carbon
2 electrode, enabling the entire analytical process to be completed in less than 5 min (Fig. 8g).¹³⁴
3

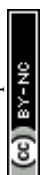
4 **4.4. Nanozymes for the SERS detection of pesticides**

5 SERS spectroscopy has firmly established itself as one of the most powerful and sensitive
6 molecular detection techniques in the field of analytical chemistry. This technique is based on
7 the phenomenon where the scattering signal of molecules adsorbed onto the surface of specific
8 metal nanostructures (primarily gold, silver, and copper) is dramatically amplified. SERS
9 provides information about the unique vibrational modes of a molecule, generating a highly
10 specific spectrum that acts like a "molecular fingerprint." The SERS technique can enhance
11 these signals by factors of 10^6 to 10^{14} , theoretically achieving detection sensitivity at the single-
12 molecule level. Due to this phenomenal sensitivity and high specificity, SERS has been
13 recognized for its potential in diverse fields such as chemistry, materials science, biology, and
14 environmental monitoring. It is considered an especially ideal tool for the analysis of pesticide
15 residues, which requires the rapid and accurate identification of trace amounts of hazardous
16 substances. Traditional SERS-based pesticide detection is broadly categorized into two
17 approaches.^{163, 164} The first is the "label-free" method, which directly measures the intrinsic
18 signal of the pesticide molecules themselves. The second is the "label-based" method, which
19 utilizes Raman reporter molecules that show distinct spectral changes upon interaction with a
20 specific pesticide.¹⁶⁵ However, these approaches have faced several inherent limitations. In the
21 case of the label-free method, there is a fundamental problem that most pesticide molecules
22 have a low affinity for the surface of metal nanoparticles and a small Raman scattering cross-
23 section, making it difficult to obtain a satisfactory SERS signal. While the label-based method



1 can improve sensitivity, it has disadvantages such as complex labeling processes, potential
2 steric hindrance, and stability issues with the reporter molecules, all of which complicate the
3 analytical procedure and increase costs.^{166, 167}

4
5 The introduction of nanozymes is emerging as an innovative alternative to overcome the
6 limitations of existing SERS techniques and to elevate analytical performance to a new level.
7 Nanozymes are functional nanomaterials that mimic catalytic behavior of natural enzymes,
8 possessing several advantages over them, such as superior stability, low production cost, and
9 the potential for easy mass production.^{168, 169} From an analytical chemistry perspective, the
10 most intriguing feature of nanozymes is that some can simultaneously serve as both a
11 plasmonic substrate for SERS signal enhancement and as an enzyme-like catalyst. For example,
12 gold nanoparticles (AuNPs) and silver nanoparticles (AgNPs) are not only the most widely
13 used plasmonic materials in SERS but also exhibit various catalytic functions, including
14 peroxidase-like activity that oxidizes specific substrates in presence of H₂O₂.^{170, 171} By
15 integrating the dual functions of 'catalyst' and 'signal amplifier' within a SERS platform,
16 nanozymes address the problems faced by conventional SERS sensors and enable new
17 analytical strategies. The product generated through the nanozyme's catalytic reaction can act
18 as a reporter molecule with a strong Raman signal. Since this product is generated in the
19 immediate vicinity of the nanozyme surface, it is positioned within the electromagnetic field
20 enhancement region where SERS signal amplification is maximized, thereby maximizing
21 signal enhancement efficiency. Pesticides can act by either inhibiting or enhancing this
22 catalytic activity of the nanozyme, which in turn leads to a variation in intensity of the finally
23 measured SERS signal.^{172, 173} Consequently, the introduction of nanozymes opens a new



avenue for detecting the presence of pesticide molecules indirectly, yet with very high sensitivity. This section aims to deeply discuss the latest research trends in the development of high-sensitivity, high-selectivity analytical platforms for pesticide detection by integrating SERS technology with the catalytic properties of nanozymes. Particularly, it will analyze the strategic advantages offered by nanozyme-based SERS detection methods and discuss the technological advancements in this field by classifying sensor systems according to their various operating principles. Table 4 shows the comparative analysis of nanozyme-based SERS sensors for pesticide detection.

Table 4. Comparative analysis of nanozyme-based SERS sensors for pesticide detection

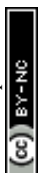
Nanozyme	Pesticides	Enzyme activity	LOD	Reaction time	Recovery rates	Ref.
FeMOF@OCTB	Isocarbophos	Oxidoreductase	2.89 pg/mL		97.7%–104%	174
Co-Fe PBA@Ag	Glyphosate	Peroxidase	0.482 ng/mL	10 min	97.64%–104.87%	175
Pt@BSA-hapten	Triazophos	Peroxidase	1 ng/mL	1 min	80.5%–109.8%	176
Cu-O-Mo nanozyme	Pirimicarb	Peroxidase	100 ng/mL	10 min	108.1%	164
Fe ₃ O ₄ @Ag NPs	Thiram	Peroxidase	0.827 ng/mL	5 min	81.6%–108.2%	163
Sea-urchin-like Au@Pt	Chlorpyrifos	Oxidase	0.742 ng/mL	30 min	100.45%–101.43%	177

4.4.1. Strategic advantages of nanozyme-based SERS detection methods

The fusion of nanozymes and SERS technology transcends a simple combination of two techniques, creating a powerful synergistic effect that maximizes analytical performance. In



1 the field of pesticide detection, this synergy offers unique strategic advantages that overcome
2 many limitations of conventional analytical methods. The core strengths of nanozyme-based
3 SERS sensors can be summarized as (1) the synergistic integration of catalytic activity and
4 signal amplification, (2) increased reproducibility of SERS signals and cost-effectiveness of
5 the sensor, (3) flexible analytical design and multiplex detection capabilities, and (4) enhanced
6 on-site applicability through simplified analytical procedures. Each of these benefits holds the
7 potential to change the paradigm of pesticide monitoring technology, and they will be discussed
8 in detail below. First, the synergistic integration of catalytic activity and signal amplification
9 is the most fundamental and powerful advantage of the nanozyme-based SERS platform.¹⁷⁸ In
10 traditional enzyme-linked SERS analysis, the enzyme that performs the catalytic reaction and
11 the plasmonic substrate that amplifies the SERS signal exist as separate components. In this
12 case, the Raman-active product generated by the enzymatic reaction must diffuse to the surface
13 of the SERS substrate for the signal to be amplified. This diffusion process not only limits the
14 reaction rate but also becomes a major cause of reduced signal amplification efficiency, as the
15 product may reach a location outside the substrate's electromagnetic field enhancement region.
16 However, the use of plasmonic nanozymes, such as AuNPs or AgNPs, fundamentally solves
17 this problem. These nanomaterials are themselves excellent SERS substrates while
18 simultaneously possessing enzyme-like activities, such as that of peroxidase. For instance,
19 consider a system using a AuNPs with peroxidase-like activity (AuNP-nanozyme). In this
20 system, the Raman-inactive substrate TMB is catalytically oxidized by the AuNP-nanozyme
21 and converted into a blue-colored oxidized TMB (oxTMB), which produces a strong Raman
22 signal. The key is that this conversion process occurs directly on the surface of the AuNP, the
23 very source of the SERS signal enhancement. This means the catalytic active site where the
24 Raman reporter is generated and the plasmonic surface where the SERS signal is amplified are

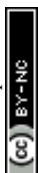


1 perfectly integrated within a single nanoparticle. This structure ensures that the generated
2 oxTMB molecules are immediately positioned in the region of the most intense
3 electromagnetic field enhancement, minimizing signal loss due to diffusion and maximizing
4 both the reaction rate and signal amplification efficiency. While this specific TMB-oxidation
5 system aptly illustrates the fundamental mechanism, its conceptual framework holds
6 significant promise for pesticide monitoring. Theoretically, if this integrated platform were
7 adapted so that a target pesticide selectively inhibits the catalytic activity of the AuNP-
8 nanozyme, a corresponding reduction in oxTMB production would be expected. This inhibition
9 could potentially translate into a quantifiable decrease in the SERS signal, offering an
10 exceptionally sensitive and innovative strategy for future pesticide detection.^{179, 180} As such,
11 this physical and functional integration of catalyst and signal amplifier serves as the core
12 mechanism for dramatically improving the sensitivity and responsiveness of the analytical
13 system. Second, nanozymes provide high signal reproducibility and cost-effectiveness, which
14 are essential for the practical application of SERS-based sensors. The intensity of a SERS
15 signal is extremely sensitive to the chemical state of the plasmonic substrate's surface and the
16 rate of the catalytic reaction. When using natural enzymes, even minor environmental changes,
17 such as in temperature or pH, can easily denature the enzyme, leading to a decrease or
18 irregularity in its catalytic activity. This, in turn, results in fluctuations in the generation rate of
19 the Raman analysis, which is a primary cause of severely compromised reproducibility in the
20 final measured SERS signals. In contrast, nanozymes, being robust inorganic nanomaterials,
21 maintain stable catalytic activity even under harsh conditions, thereby ensuring consistent and
22 reliable SERS signals in a variety of real sample environments. This stability not only allows
23 for the long-term storage of the sensors but also provides a foundation for the mass production
24 of uniform-quality SERS substrates with minimal batch-to-batch variation, thus enhancing the



1 reliability of analytical results. From a cost-effectiveness perspective, the low production cost
2 of nanozymes enables the development of affordable, disposable SERS sensors.¹⁸¹ In practical
3 field analysis, the use of disposable sensors is crucial to prevent cross-contamination between
4 samples. Substrates that rely on expensive natural enzymes or require complex fabrication
5 processes struggle to meet this demand. However, nanozymes are ideal for disposable
6 platforms due to their potential for low-cost mass production.¹⁸² This provides the economic
7 feasibility for SERS technology to expand beyond the confines of expensive laboratory
8 equipment into user-friendly, on-site diagnostic tools. Ultimately, the stability and economy of
9 nanozymes are key factors in achieving consistent SERS signals and practical analysis costs,
10 playing a decisive role in accelerating the technology's commercialization.

11
12 Third, the nanozyme-based SERS platform demonstrates outstanding potential in its
13 flexibility of analytical design and multiplexing capabilities. The catalytic and plasmonic
14 properties of a nanozyme can be optimized by precisely controlling the nanoparticle's size,
15 shape, composition, and surface chemistry.¹⁸³ For example, anisotropic nanostructures with
16 sharp vertices or edges, such as gold nanostars or silver nanocubes, can generate much stronger
17 electromagnetic fields compared to spherical nanoparticles, thereby maximizing the SERS
18 enhancement effect. At the same time, these structural variations also influence the nanozyme's
19 catalytic activity, making it possible to simultaneously tune both properties to fit specific
20 analytical objectives. Furthermore, the surface of a nanozyme can be easily functionalized with
21 various molecular recognition elements through methods like thiol (-SH) bonding or
22 electrostatic interactions. By immobilizing elements with high affinity and specificity for a
23 particular pesticide such as aptamers, antibodies, or molecularly imprinted polymers (MIPs)



1 onto the nanozyme-SERS substrate, the target pesticide can be selectively captured,
2 dramatically improving the selectivity of the analysis.^{184, 185} This design flexibility extends to
3 multiplex detection. A key feature of SERS spectra is that they exhibit very narrow peaks at
4 unique positions for each molecule. Therefore, by using several types of reporter molecules
5 that produce distinct Raman signals, it is possible to simultaneously quantify multiple types of
6 pesticides in a single measurement. For instance, one could design multiple nanozyme systems,
7 each regulated by a different pesticide and engineered to generate a unique Raman analysis.
8 By integrating these systems onto a single substrate, the concentrations of various pesticides
9 can be determined at once by analyzing the peak intensity of each analysis in the measured
10 SERS spectrum. This multiplexing capability saves time and money, reduces sample
11 consumption, and presents a highly effective solution for analyzing real agricultural and
12 environmental samples where multiple types of pesticides are often co-mingled.¹⁶³

14 Fourth, the integration of nanozymes and SERS significantly simplifies the analytical
15 procedure, greatly enhancing its applicability for rapid POCT. As mentioned earlier, because
16 the catalyst and the signal transducer are integrated into a single nanoparticle, the sensor
17 fabrication process is simplified, and the number of analytical steps is reduced. Many
18 nanozyme-based SERS analysis systems can be implemented in a "mix-and-read" format,
19 where the sample and reagents are mixed, allowed to react for a certain period, and then the
20 SERS signal is measured. This is a tremendous advantage compared to traditional methods like
21 chromatography-mass spectrometry (LC-MS/GC-MS), which require complex sample
22 pretreatment, separation, and purification steps.¹⁷⁷ The analysis time is reduced from hours to
23 minutes, and measurements can be performed without expensive, bulky equipment or highly



1 skilled analysts. This simplification has true potential when combined with a portable Raman
2 spectrometer. Recent technological advancements have led to the development of high-
3 performance, palm-sized portable Raman devices, paving the way for the direct use of
4 nanozyme-SERS platforms wherever they are needed at agricultural production sites,
5 distribution centers, food processing plants, and water quality management sites.¹⁸⁶ This
6 enables real-time, on-site screening for pesticide contamination, which maximizes the
7 efficiency of food safety management systems and contributes to proactively preventing threats
8 to public health through rapid responses to contamination incidents. As such, the procedural
9 simplicity and on-site applicability offered by nanozyme-based SERS technology will be a key
10 driving force in shifting the paradigm of pesticide safety management from centralized
11 laboratory analysis to decentralized, on-site diagnostics.

13 **4.4.2. Classification of nanozyme-based SERS sensors for pesticide detection**

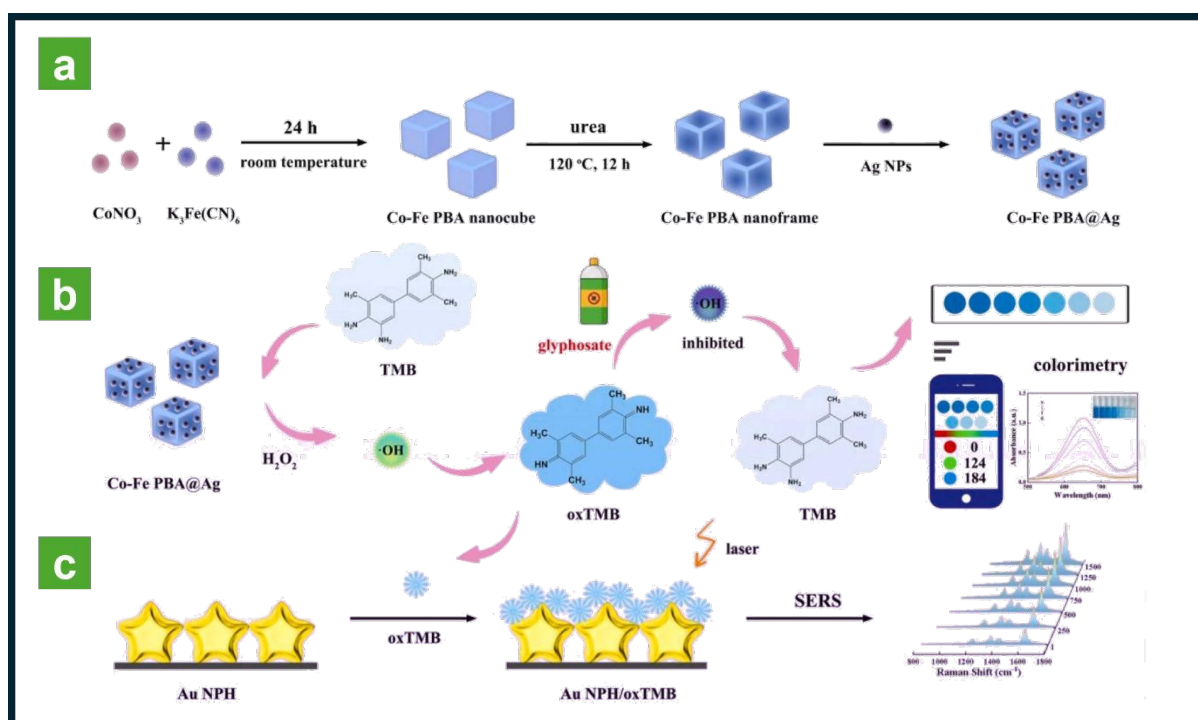
14 Although nanozyme-based SERS sensors for pesticide detection can be designed in a wide
15 variety of forms, their fundamental operating principles can be broadly classified into two key
16 mechanisms based on how the target pesticide affects the nanozyme's catalytic activity. The
17 first is the 'direct modulation of nanozyme activity' mechanism, in which the pesticide molecule
18 itself directly inhibits or enhances catalytic activity of nanozyme. The second is the 'indirect
19 modulation via recognition element' mechanism, where the pesticide first binds to a separate
20 recognition element, such as an aptamer or a MIP, and this binding event then indirectly
21 controls the active state of the nanozyme. This classification provides an important framework
22 for understanding the operating principles of these sensors.

23



1 4.4.2.1. Direct modulation of nanozyme activity

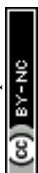
2 The first mechanism, 'direct modulation' is based on the principle that the pesticide molecule
3 alters the intrinsic catalytic efficiency of the nanozyme, either by binding directly to its active
4 site or by scavenging key intermediates like reactive oxygen species (ROS) that are generated
5 during the catalytic process. This approach is an intuitive strategy that utilizes the unique
6 chemical properties or reactivity of the pesticide as a core element for signal generation. This
7 mechanism can be further divided into two subtypes including inhibition and enhancement.



9
10 **Fig. 9.** Pesticide detection through direct modulation of nanozyme activity mechanism. (a)
11 Schematic for the synthesis of Co-Fe PBA@Ag for glyphosate sensing. (b) Sensing mechanism
12 of glyphosate based on Co-Fe PBA@Ag (c) SERS-based detection of glyphosate using Co-Fe
13 PBA@Ag. Reproduced from reference¹⁷⁵ with permission from Elsevier, copyright 2025.



1 Activity inhibition is the most common direct modulation method, implementing a 'turn-off'
2 sensor where the SERS signal decreases as the pesticide interferes with the nanozyme's
3 catalytic function. A prime example of direct inhibition was reported in 2025 by He's research
4 team for a glyphosate detection sensor. The Co-Fe PBA@Ag nanozyme was prepared through
5 a multi-stage strategy starting with the synthesis of Co-Fe PBA nanocubes. To increase the
6 specific surface area and active sites, these nanocubes were subjected to urea etching under
7 hydrothermal conditions at 100°C for 12 h, resulting in the formation of concave nanoframes
8 with a sunken surface. Finally, AgNPs were assembled onto the polyethylenimine (PEI)-
9 modified nanoframes to yield the bimetallic Co-Fe PBA@Ag composite, which exhibits
10 significantly enhanced peroxidase-like catalytic activity due to the synergistic effects between
11 the metals (Fig. 9a). The Co-Fe PBA@Ag nanozyme used in this sensor decomposes H₂O₂ to
12 generate hydroxyl radicals (•OH), a powerful oxidizing agent, which in turn oxidize TMB.
13 Glyphosate, rather than binding to the nanozyme itself, has the ability to effectively scavenge
14 the hydroxyl radicals, which are the key intermediates of the catalytic reaction. In other words,
15 glyphosate reacts with and deactivates the hydroxyl radicals before TMB can, thus hindering
16 TMB's oxidation (Fig. 9b).¹⁷⁵ For ultrasensitive quantification, gold polyhedra (Au NPH) were
17 employed as the SERS-enhancing substrate. The irregular morphology of the Au NPH
18 facilitates the generation of numerous SERS signal. This also induces a 'turn-off' response
19 where the SERS signal decreases as the glyphosate concentration increases (Fig. 9c). An
20 extremely low LOD of 2.85 nM was achieved using the nanozyme SERS sensor designed with
21 this mechanism. The sensor's reliability in complex matrices was confirmed with high recovery
22 rates for honey (97.64–104.87%), potato (97.87–104.86%), and tea (98.46–104.75%). As
23 shown, the direct inhibition method is highly effective as it directly leverages specific chemical
24 properties of pesticides, such as the affinity of certain functional groups (e.g., thiols, amines)

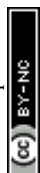


1 for the nanozyme's active site or the pesticide's radical-scavenging ability.

2

3 **4.4.2.2. Indirect modulation via recognition element**

4 The second mechanism, 'indirect modulation via a molecular recognition element' is a
5 strategy where the nanozyme's own catalytic activity is kept constant, while a third-party
6 molecular recognition element (e.g., aptamer, MIP) that specifically binds to the pesticide is
7 introduced to control the overall signal output. In this approach, the nanozyme focuses on its
8 role as a signal-generating reporter, and the sensor's selectivity depends entirely on the
9 performance of the molecular recognition element. The binding event between the pesticide
10 and the recognition element induces physical and structural changes in the system, and these
11 changes affect the nanozyme's catalytic reaction environment, ultimately leading to a change
12 in the final SERS signal. The aptasensor for isocarbophos (IPS) detection, developed by Li's
13 research team in 2022, is a typical example of this indirect modulation method. In this system,
14 a DNA aptamer that specifically binds to IPS is first physically adsorbed onto the surface of a
15 FeMOF@OCTB nanozyme. In this adsorbed state, the nanozyme's active sites are covered,
16 and the catalytic reaction is inhibited (Fig. 10a). However, if IPS is present in the sample, the
17 aptamer binds to IPS with a much stronger affinity than to the nanozyme surface, causing it to
18 detach. As the aptamer is released, the nanozyme's active sites are re-exposed, allowing the
19 nanozyme to recover its original catalytic activity. This restoration of catalytic performance is
20 manifested as a concentration-dependent colorimetric response, which can be quantitatively
21 monitored through UV-vis absorption spectra (Fig. 10b). Furthermore, the recovered activity
22 promotes the reduction of Ag⁺ ions into Ag nanoparticles. The resulting AgNPs act as a SERS
23 substrate, generating a strong signal. In other words, the pesticide acts as an inducer that

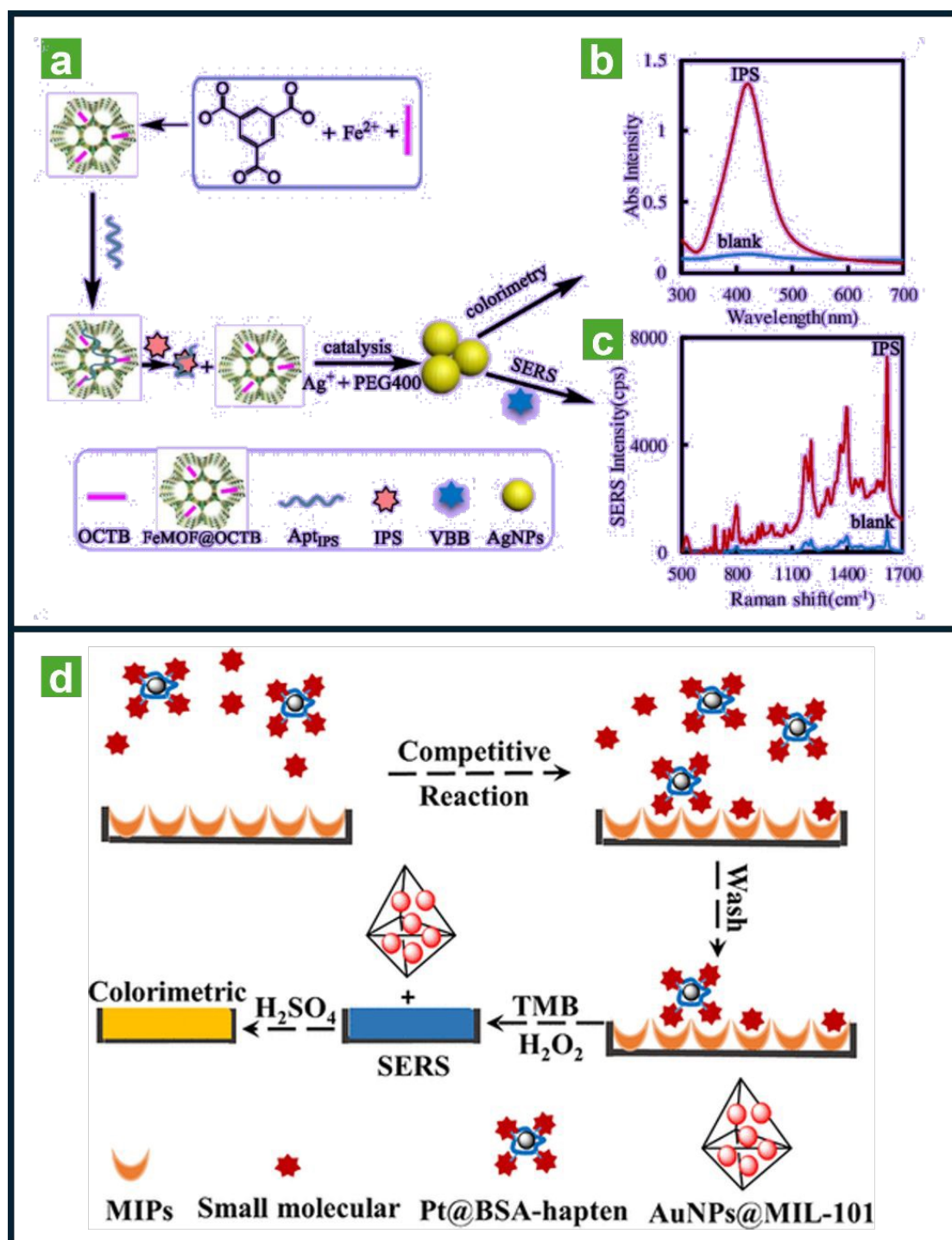


1 releases the aptamer that was inhibiting the catalytic activity, thereby restarting the suppressed
2 catalytic reaction and generating a 'turn-on' signal (Fig. 10c).¹⁷⁴

3
4 The sensor for triazophos detection, developed by Yan's research team in 2019, is another
5 example of indirect modulation that uses the principle of competitive immunoassay. In this
6 system, a MIP with recognition sites for triazophos is first immobilized on a 96-well plate.
7 Separately, a 'nanozyme-labeled competitor' (Pt@BSA-hapten) is prepared by conjugating a
8 nanozyme (PtNPs) to a structural analog (hapten) of triazophos. During analysis, the sample
9 containing triazophos and a fixed amount of the nanozyme-labeled competitor are added
10 simultaneously to the MIP-coated wells. At this point, the triazophos from the sample and the
11 nanozyme-labeled competitor compete for the limited number of MIP binding sites. If the
12 concentration of triazophos in the sample is high, it will occupy most of the MIP binding sites,
13 leaving the nanozyme-labeled competitor unable to bind and to be removed during the washing
14 step. Therefore, the final measured SERS signal is very weak. Conversely, if there is no or a
15 low concentration of triazophos in the sample, most of the nanozyme-labeled competitor will
16 bind to the MIP, generating a strong SERS signal. This results in a typical 'turn-off' competitive
17 analysis curve, where the signal decreases as the pesticide concentration increases (Fig. 10d).¹⁷⁶
18 Nanozyme-based SERS sensors can be clearly distinguished by whether the pesticide directly
19 influences the nanozyme's activity or indirectly affects it through a separate recognition
20 element. The direct modulation approach allows for the construction of relatively simple
21 systems by leveraging the chemical properties of the pesticide. In contrast, the indirect
22 modulation approach offers the advantage of securing excellent selectivity and versatility by
23 utilizing highly specific recognition elements such as aptamers or MIPs. An understanding of



- 1 these mechanisms will serve as a crucial theoretical foundation for designing and developing
- 2 high-performance SERS sensors optimized for specific pesticides and analytical environments
- 3 in the future.



4



1 **Fig. 10.** Pesticide detection using indirect modulation via recognition element mechanism. (a)
2 FeMOF@OCTB catalytic amplification mechanism for isocarbophos detection. (b) UV-vis
3 absorption spectra showcasing the concentration-dependent colorimetric response toward
4 isocarbophos. (c) SERS signal recovery resulting from the target-induced release of Fe-
5 MOF@OCTB from the aptamer-nanozyme complex. Reproduced from reference¹⁷⁴ with
6 permission from Springer Nature, copyright 2022. (d) Pt@BSA-Hapten catalytic amplification
7 mechanism for triazophos detection. Reproduced from reference¹⁷⁶ with permission from
8 American Chemical Society, copyright 2019.

9
10 A comparative analysis of the reported nanozyme-based pesticide sensors reveals
11 several common design principles that influence analytical behaviors of colorimetric,
12 fluorescence, electrochemical, and SERS sensing platforms. Structural engineering of
13 nanozymes such as heterostructure formation, surface functionalization, defect-rich
14 nanomaterials and increased catalytic activity, enhances active site exposure, and facilitates
15 transfer of electron processes, thereby improving response kinetics and sensitivity. In optical
16 sensing systems, including colorimetric and fluorescence methods, catalytic activity
17 modulation of nanozymes, particularly the inhibition or alteration of peroxidase-like activity
18 by pesticides, represents the most employed detection mechanism. Electrochemical sensors
19 frequently exploit nanozyme-assisted electrocatalysis or catalytic inhibition to modulate redox
20 signals at electrode interfaces, while SERS-based systems typically rely on nanozyme-assisted
21 catalytic generation of Raman-active species or plasmonic signal amplification mediated by
22 analyte surface interactions.

23



1 Despite the substantial progress reported in these sensing strategies, several trade-offs
2 remain between sensitivity, operational robustness, and material cost. Noble metal-based
3 nanozymes (e.g., gold, platinum, palladium) exhibit exceptional catalytic activity, yet their high
4 cost and limited availability raise concerns regarding material criticality and large-scale
5 deployment. Conversely, carbon-based nanozymes, transition metal oxides, and metal organic
6 framework-derived systems, provide more cost-effective, improved chemical stability, and
7 better scalability. Sometimes they exhibit comparatively lower catalytic efficiency. Another
8 important limitation observed across many studies is the limited validation of sensor
9 performance in complex real-world matrices. While numerous reports demonstrate excellent
10 analytical sensitivity under controlled laboratory conditions, systematic testing in real
11 agricultural products, food matrices, and environmental water samples is still relatively limited,
12 leaving potential interference effects insufficiently addressed. Hence, future development of
13 nanozyme-based pesticide sensors should focus on improving sensor robustness, enhancing
14 selectivity, and performing comprehensive real-sample validation to facilitate reliable
15 deployment in the practical monitoring applications.

17 Nanozymes have revolutionized pesticide analysis by providing robust alternatives to natural
18 enzymes, showcasing significant potential across colorimetric, fluorescent, electrochemical,
19 and SERS-based sensing platforms. An in-depth exploration of the discussed four key detection
20 modalities was provided, by systematically analyzing their strategic advantages, operating
21 principles, and interaction mechanisms. While nanozyme technology enables the development
22 of ultra-sensitive and highly selective analytical methods, the choice of a specific platform
23 depends heavily on specific analytical requirements, such as the desired sensitivity, sample



1 matrix complexity, and the availability of instrumentation at the point of need. Each modality
 2 offers unique benefits while facing distinct technical hurdles; for instance, colorimetric sensors
 3 excel in rapid, equipment-free screening, whereas SERS and electrochemical platforms are
 4 favored for ultra-trace quantification and high precision. Table 5 outlines best-use scenarios,
 5 key limitations, failure modes, and mitigation strategies for each modality. Through this
 6 multifaceted approach, we aim to provide a comprehensive outlook on the limitless potential
 7 of innovative nanozymes as next-generation, sensing technologies for safeguarding future food
 8 security and environmental safety.

9
 10 **Table 5. Decision matrix for selecting nanozyme-based sensing modalities for pesticide**
 11 **detection.**

Modality	Preferred Scenarios	Key Limitations	Potential Failure Modes	Mitigation Strategies
Colorimetric	Rapid on-site screening, naked-eye detection, low-cost disposable kits	Lower sensitivity compared to optical/electrical methods	High background color from real samples	Use of smartphone-based RGB analysis or ratiometric color shifts
Fluorescence	High-sensitivity requirements, real-time imaging	Susceptibility to photobleaching and background fluorescence	Signal quenching by non-target matrix components (Matrix effect)	Implementation of dual-emission ratiometric probes or AIE-active nanozymes
Electrochemical	High-sensitivity requirements, portability (POCT)	Electrode fouling due to bio-adsorption in complex samples	Loss of signal due to non-specific adsorption (Biofouling)	Surface modification with anti-fouling layers
SERS	Ultra-trace detection, molecular fingerprinting, multiplexing	Signal reproducibility issues and complex substrate fabrication	Inconsistent signals due to "hotspot" non-uniformity	Design of uniform 3D superstructures or use of internal standards

12

13 **Key design rules and practical recommendations for nanozyme-based pesticide sensors**



1 Based on collective analysis of recent studies, key design rules and practical
2 recommendations can be proposed for assistance in designing high-performance nanozyme-
3 based pesticide sensing systems across colorimetric, electrochemical, fluorescence, and SERS
4 platforms.^{187, 188} These include rational selection of catalytic substrates and signal reporters,
5 engineering nanozyme catalytic activity and nanostructure, enhancing selectivity through
6 surface functionalization, minimizing the matrix interference, signal amplification strategies,
7 improving stability and reproducibility, balancing sensitivity, cost, and scalability, and
8 validation in real-world matrices.^{14, 84, 189}

9

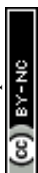
10 **5. Challenges and future perspectives**

11 **5.1. Colorimetric detection**

12 While nanozyme-based colorimetric sensors have made remarkable progress, several
13 significant challenges must be addressed to achieve widespread commercialization and on-site
14 application. First is the challenge of ensuring selectivity and reproducibility in complex real
15 samples. Real samples such as fruits, vegetables, and soil contain numerous organic substances,
16 salts, and proteins in addition to the target pesticide, which can cause a 'matrix effect' that
17 interferes with the nanozyme's activity.^{23, 46, 190} Although many studies discussed herein have
18 demonstrated feasibility by applying their sensors to real variant samples with excellent
19 recovery rates of over 90%, validation across a broader and more diverse range of matrices is
20 needed.^{48-51, 54, 57, 59, 61, 64, 65, 67, 68, 70, 73, 75-77} To overcome this, research into developing robust
21 nanozymes that are less affected by the sample matrix, or combining them with simple yet
22 effective pretreatment techniques, is essential. Second, a more fundamental understanding of
23 the catalytic mechanisms is still needed. Despite the reporting of numerous high-efficiency



1 nanozymes, identification of exact structure of active sites and the atomic-level elucidation of
2 how they interact with pesticide molecules to modulate activity are still in their early stages.¹⁹¹,
3 ¹⁹² For instance, the study on the Pt-Np-C nanozyme identified that a specific nitrogen
4 configuration (Pyridinic N) around the platinum active site plays a crucial role in pesticide
5 recognition, but in-depth studies at this level are still lacking.⁶³ Efforts to clarify reaction
6 mechanisms through computational science and in-situ characterization techniques will
7 provide the foundation for designing 'tailor-made nanozymes' in a more rational and
8 sophisticated manner, moving beyond empirical development. Future research on nanozyme-
9 based colorimetric sensors is expected to advance in several promising directions. The
10 development of intelligent and multifunctional systems will accelerate.²⁶ This includes the
11 creation of highly integrated materials where a single nanozyme can simultaneously detect
12 multiple pesticides or perform both degradation and detection of pesticides. Furthermore,
13 research on 'smart' nanozymes, such as the Pt/Co₃O₄ nanoflowers that engineered by *Sun et al.*
14 in 2022, whose function can be modulated by external stimuli like pH, will expand, allowing
15 for stimuli-responsive sensing platforms.⁷⁷ These advancements will be further amplified
16 through integration with portable smart devices. The use of smartphone cameras and dedicated
17 applications for quantitative analysis of colorimetric signals, already actively pursued in many
18 cases, could evolve into extensive data management systems by integrating with Internet of
19 Things (IoT) technology to transmit measurement data to the cloud in real-time and create
20 pesticide contamination maps.^{51, 66, 70} In conclusion, nanozyme-based colorimetric sensors,
21 through continuous and interdisciplinary research to overcome current technical challenges,
22 hold the potential to firmly establish themselves as a universal and reliable on-site analytical
23 technology for protecting the future of food and environmental safety, ultimately contributing
24 to public health and sustainable agriculture.



1 5.2. Fluorescence detection

2 While nanozyme-based fluorescence detection methods show outstanding potential in the
3 field of pesticide analysis, there are still technical challenges and limitations that must be
4 overcome for their transition to real-world applications and commercialization. At the same
5 time, this field is rapidly advancing, with new research directions and innovative approaches
6 continuously being proposed, so its future prospects can be considered very promising. One of
7 the most significant challenges is the complex matrix effect and the influence of interfering
8 substances. These substances can inhibit the catalytic activity of nanozymes or interfere with
9 the fluorescent signal. Therefore, ensuring selectivity and specificity is crucial. Structurally
10 similar pesticides or cross-reactive interfering substances can lead to the false positive or false
11 negative results. This is especially true in complex biological samples, where the concentration
12 of interfering substances (μM - mM) can be millions of times higher than the target analyte
13 concentration (fM - nM), requiring an extremely high selectivity on the order of 10^9 :1. Although
14 solutions utilizing aptamers or molecular imprinting technology are being proposed,
15 improvements in the efficiency of the aptamer screening process and precise control of
16 nanozyme-aptamer interactions are needed. Furthermore, direct detection methods using only
17 nanozymes are fundamentally limited in selectivity due to the lack of recognition units, making
18 the development of sensor arrays or pattern recognition-based approaches essential.^{193, 194}
19 Issues of reliability and reproducibility are key hurdles to commercialization. Single-mode
20 detection systems have inherent reliability limitations, making them vulnerable to false signals
21 or instrumental fluctuations. To address this, dual-mode (fluorescent-colorimetric) or multi-
22 mode detection systems are being developed, which can significantly enhance reliability
23 through intrinsic self-verification and self-correction functions. However, reproducibility



1 issues stemming from batch-to-batch variability in nanozyme synthesis, changes in activity
2 depending on storage conditions, and the influence of the measurement environment still need
3 to be resolved.^{195, 196} Despite the clear limitations of nanozyme-based fluorescence detection
4 technology, the potential for growth in this field is limitless, with several innovative research
5 directions indicating future breakthroughs. Particularly, designing sensing materials with high
6 selectivity will be a key focus. To solve the current problems of matrix effects and non-specific
7 interference, future nanozymes will be combined with advanced molecular recognition systems
8 such as molecular imprinting technology or aptamers. This is analogous to designing
9 "customized pockets" on the nanozyme surface to precisely capture specific pesticide
10 molecules. Furthermore, 'computer-aided nanozyme engineering' will be realized by utilizing
11 artificial intelligence and computational science to predict and design active sites that perfectly
12 match the 3D structure and charge distribution of a specific pesticide. This will grant the
13 ultimate selectivity, enabling a reaction only with the target molecule even in complex samples.

15 **5.3. Electrochemical detection**

16 Although nanozyme-based electrochemical pesticide sensors have shown remarkable
17 achievements, significant challenges remain in translating laboratory-level success into reliable,
18 commercially viable products for field use. Real samples, such as agricultural products, soil,
19 and river water, contain numerous interfering substances like proteins, fats, humic acids, and
20 inorganic salts. These substances can adsorb onto the electrode surface, inhibiting the
21 nanozyme's activity (biofouling) or causing non-specific signals, which severely compromises
22 the sensor's accuracy and reproducibility.¹⁹⁷⁻¹⁹⁹ Therefore, it is essential to develop effective
23 sample preparation techniques or to integrate anti-fouling surface modification technologies



1 (e.g., coating with polyethylene glycol or zwitterionic polymers).²⁰⁰⁻²⁰³ Despite the
2 experimentally proven high catalytic activity of many nanozymes, the exact structure of their
3 active sites and their catalytic reaction pathways often remain unclear. Clarifying the structure-
4 activity relationship by combining theoretical studies, such as density functional theory
5 calculations, with real-time spectroscopic analysis is a prerequisite for the rational design of
6 next-generation nanozymes with tailored activity and selectivity for specific pesticides.^{148, 204,}
7 ²⁰⁵ Standardization and quality control also make challenges. The shape, size, and surface
8 chemistry of nanozymes are highly sensitive to synthesis conditions, which leads to batch-to-
9 batch variations in catalytic activity. To ensure sensor reliability, the establishment of
10 standardized synthesis and quality control protocols for precisely controlling and analyzing
11 nanozyme properties is urgently needed.¹⁴⁰ Despite these challenges, the future is advancing
12 substantially. Artificial intelligence and machine learning can be utilized to analyze vast
13 experimental datasets to predict optimal nanozyme compositions and structures, and to screen
14 for new candidate materials. This will enable a shift from traditional 'trial-and-error' research
15 methods to efficient, data-driven nanozyme development.^{206, 207} Furthermore, integration with
16 microfluidics technology will enable the creation of 'Lab-on-a-Chip' systems that automate the
17 entire process from sample injection, mixing, and reaction to detection on a single chip. This
18 will minimize sample consumption, reduce analysis time, and decrease user error, thereby
19 maximizing the accuracy and convenience of on-site analysis.^{208, 209} Ultimately, multiplexed
20 analysis platforms will be developed by integrating multiple nanozyme sensors, each selective
21 for a different pesticide, onto a single array. This will allow for the simultaneous quantitative
22 analysis of various pesticides in a single measurement. When these technologies are combined
23 with wireless communication and the IoT, it will become possible to build distributed
24 environmental monitoring networks that can monitor pesticide contamination in real-time over

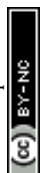


1 wide areas and transmit the data to a central server.²¹⁰

2

3 **5.4. SERS detection**

4 Combination of nanozymes and SERS technology has led to remarkable advancements in
5 the field of pesticide detection by providing excellent sensitivity and convenience, several
6 significant challenges still need to be addressed before this technology can move beyond
7 laboratory-level proof-of-concept to become a commercialized, on-site analytical tool. These
8 challenges include ensuring the reliability and reproducibility of quantitative analysis and
9 overcoming the matrix effects of complex real-world samples. However, these challenges also
10 serve as signposts for future research directions in this field. It is anticipated that future research
11 will move towards overcoming current limitations and maximizing the technology's practical
12 value through the rational design of nanozymes, the development of intelligent hybrid materials,
13 and the construction of multi-modal analysis systems. One of the most pressing challenges
14 currently facing by the nanozyme-based SERS sensors is ensuring the reliability and
15 reproducibility of quantitative analysis. The SERS signal is extremely sensitive to the distance
16 between and arrangement of nanoparticles, that is, the distribution of 'hotspots'.²¹¹ The simple
17 method of drying a nanozyme solution onto a substrate can lead to a non-uniform distribution
18 of nanoparticles due to the 'coffee-ring effect' that occurs as the solvent evaporates. This can
19 result in SERS signals that vary by more than an order of magnitude depending on the
20 measurement spot, which is a critical drawback that makes accurate quantitative analysis nearly
21 impossible.²¹² Furthermore, minor batch-to-batch variations in the size or shape of nanozymes
22 during synthesis lead to non-uniformity in their catalytic activity and plasmonic properties,
23 which is another factor that undermines the reproducibility of the sensor's performance.²¹³



1 Future research aimed at solving these challenges will unfold more sophisticated and intelligent
2 directions. First, the rational design of nanozymes will become a core research theme. Moving
3 beyond the conventional trial-and-error approach, research will actively utilize computer
4 simulations like DFT and machine learning to predict the correlation between the structure,
5 composition, and catalytic-plasmonic properties of nanomaterials. This will enable the design
6 of 'customized nanozymes' with active sites and surface characteristics optimized for specific
7 pesticides.^{214, 215} Particularly, nanozymes with active centers clearly defined at the atomic level,
8 such as platinum single-atom nanozymes, will provide crucial model systems for
9 fundamentally understanding and controlling catalytic mechanisms. Second, efforts to
10 overcome current limitations through the development of intelligent hybrid materials will
11 accelerate. For example, immobilizing nanozymes within porous materials like MOFs or COFs
12 can prevent nanozyme aggregation, thereby increasing stability and providing a uniform
13 reaction environment. Furthermore, by controlling the pore size or chemical properties of
14 MOFs, they can be designed to act as a 'molecular sieve,' selectively allowing only pesticide
15 molecules of a certain size to pass through. They can even be endowed with the ability to pre-
16 concentrate target pesticides by attaching molecular recognition elements like aptamers or
17 antibodies to the MOF surface.²¹⁶ Such hybrid systems can offer innovative solutions that
18 minimize matrix effects while simultaneously enhancing both selectivity and sensitivity.
19 Nanozyme-based SERS technology holds immense potential in the field of pesticide detection.
20 While current technical hurdles such as reproducibility and matrix effects are evident, these
21 limitations will gradually be overcome through future-oriented research, including rational
22 design via computational science, the development of intelligent hybrid materials, and
23 integration with smart devices. When these efforts come to realization, nanozyme-SERS
24 technology will establish itself as a powerful and reliable technology, keeping food and



1 environment safe.

2

3 **5.5. Concrete recommendations for standardization and practical application**

4 To bridge the gap between laboratory-level proof-of-concept and commercial on-site
5 applications, it is imperative to move beyond general challenges and establish strict, concrete
6 guidelines for the development of nanozyme-based pesticide sensors. Because the shape, size,
7 and surface chemistry of nanozymes are highly sensitive to synthesis conditions, leading to
8 batch-to-batch variations, future studies must adhere to minimum reporting standards.
9 Researchers should explicitly detail precursor concentrations, precise thermal protocols, and
10 kinetic parameters. Furthermore, to ensure commercial viability, it is highly recommended to
11 systematically evaluate and report the inter-batch relative standard deviation (RSD) for
12 catalytic activity. Ideally, this RSD should be maintained below 5–10%, which strongly aligns
13 with the universally accepted analytical guidelines and the high precision recently
14 demonstrated by state-of-the-art nanozyme sensors (e.g., RSDs of <2.52%, 1.18–5.26%, and
15 <6.17%).^{133, 162, 217} Achieving such low batch-to-batch variation across at least three
16 independently synthesized batches is essential for practical commercialization.

17

18 While nanozymes are fundamentally more robust than natural enzymes, empirical
19 long-term stability data is often missing in current literature. Concrete storage and shelf-life
20 testing must become standard practice. Proposed sensors should be subjected to systematic
21 aging tests to evaluating catalytic retention at both room temperature and 4 °C over a minimum
22 period of 3 to 6 months. This specific timeframe is not arbitrary. Instead, it represents the



1 minimum viable shelf-life required to accommodate the manufacturing-to-distribution supply
2 chain and covers a full seasonal cycle of agricultural pesticide monitoring. Furthermore, it
3 empirically validates the intrinsic advantage of inorganic nanozymes, which are known to
4 maintain stable catalytic performance for several months without rigorous refrigeration
5 requirements. Achieving an activity retention of >90% over this period should serve as a
6 baseline metric for practical, field-ready commercial sensors. In addition, biofouling and non-
7 specific adsorption on sensor surfaces drastically reduce reproducibility when analyzing
8 complex real-world samples.

9 Finally, matrix effects from real samples, such as agricultural runoffs or crude fruit
10 extracts, cannot be ignored during practical implementation. To mitigate these interferences,
11 simple external calibration is often insufficient. The routine adoption of standard addition
12 methods or ratiometric sensing approaches should be highly encouraged to self-calibrate and
13 eliminate background environmental fluctuations. Furthermore, any proposed sensor must
14 undergo rigorous validation across at least three structurally different sample matrices. The
15 analytical accuracy must be quantified by spiking standard pesticides and calculating recovery
16 rates, which should strictly fall within the universally acceptable analytical range of 80% to
17 120%, alongside intra-assay RSDs tightly controlled below 10%.

18 19 **6. Conclusions**

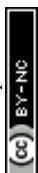
- 20 ➤ Chemical pesticides are essential for maintaining agricultural production and
21 guaranteeing global food security by efficiently managing pest infestations in crops
22 and storage facilities. Continual dependence on pesticides poses significant problems.
23 Insufficient safety protocols, including farmers' limited understanding of safety labels



1 and the absence of appropriate protective gear, intensify health and environmental
2 hazards. The detrimental consequences of pesticide application are profoundly
3 affecting human health both physically and neurologically while also hurting non-
4 target organisms, soil quality, groundwater, and local climatic conditions.
5 Acknowledging the essential role of pesticides in agricultural production, similar to
6 the function of medicine in disease management, underscores the significance of
7 proper application procedures. It is vital to implement targeted applications for the
8 successful management of pests while protecting food crops. Systematic surveillance
9 of pesticide residues prior to their transportation and consumption is essential to
10 mitigate risks, including the refusal of contaminated shipments in global trade.

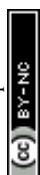
11
12 ➤ Advancements in nanotechnology have led to considerable interest in nanomaterials
13 owing to their remarkable physical, mechanical, optical, and chemical capabilities.
14 Nanozymes are a category of nanomaterials demonstrating enzyme-like functions.
15 Nanozymes have garnered significant attention since their inception in 2004 and have
16 emerged as a key research focus in the domain of artificial enzymes. Nanozymes
17 mitigate several drawbacks of natural enzymes, including complex manufacturing and
18 purification processes, inadequate stability, and reduced recycling effectiveness.
19 Various nanozyme-based techniques have been designed for identifying levels of OPs
20 in biosensor and food security applications. Nanozyme-based biosensors provide
21 considerable advantages in OPs detection, with their analytical performance primarily
22 demonstrated by operational simplicity, high sensitivity, excellent selectivity, and
23 reliability.

24



1 ➤ The distinctive properties of nanozymes, such as enzyme-like catalytic activity,
2 structural adaptability, operating durability, and versatility, facilitate their
3 incorporation into various detection methodologies, ranging from colorimetric and
4 fluorescent sensors that provide rapid visual or optical outputs, to electrochemical
5 platforms characterized by excellent sensitivity and real-time quantification, and
6 SERS systems offering highly sensitive molecular identification. Across these
7 detection methodologies, nanozymes will function through various mechanisms, such
8 as inhibition of intrinsic enzyme activity, direct catalytic interaction with the
9 molecules of pesticide, amplification of signals through molecular specific
10 recognition, and alteration of electron transfer pathways, enabling versatile design
11 strategies for the detection of various classes of pesticides. The growing demand for
12 improved detection systems has generated significant interest in innovative sensing
13 technologies. The latest advancements in materials chemistry and nanozymes
14 emphasize the importance of potential sensors.

15
16 ➤ In this review, we provided a critical overview on nanozymes for OPs detection,
17 encompassing their material compositions, strategies for activity enhancement, output
18 signals, and detection methodologies. Explicitly provided a detailed summary on the
19 progress and perspectives of nanozyme-based SERS sensors, electrochemical sensors,
20 fluorescence sensors, and colorimetric sensors for high-sensitivity sensing of
21 pesticides. By presenting the recent trends and advances of nanozymes for the
22 detection of pesticides, this review aims to provide a scientific reference for the
23 utilization of innovative nanozymes in pesticides detection, facilitating their practical
24 utilization in convenient and rapid analysis.



1 Acknowledgements

2 The authors acknowledge financial support from the Brain Pool Program (RS-2024-
3 00447409) and the Basic Science Research Program (RS-2026-25483852) through the
4 National Research Foundation of Korea (NRF), funded by the Ministry of Science, ICT, and
5 Future Planning. This research was supported by a grant (25192MFDS001) from Ministry of
6 Food and Drug Safety in 2025.



1 **References**

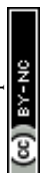
- 2 1. Z. Chen, R. Feng, Q. Zhou, X. Zhang, Y. Fan, D. Fang, R. Zheng, W. Zhang, Z. Lu, J.
3 Chen, Q.-W. Zhang, C. Jiang, P. Li, H. Yu and G. Li, Biomaterials and biosensing
4 technologies in the detection and removal of pesticide residues: Current trends
5 and future prospects, *Coordination Chemistry Reviews*, 2026, 547, 217110.
6 <https://doi.org/10.1016/j.ccr.2025.217110>.
- 7 2. S. Naveen Prasad, V. Bansal and R. Ramanathan, Detection of pesticides using
8 nanozymes: Trends, challenges and outlook, *TrAC Trends in Analytical Chemistry*,
9 2021, 144, 116429. <https://doi.org/10.1016/j.trac.2021.116429>.
- 10 3. F. Zhao, L. Wang, M. Li, M. Wang, G. Liu and J. Ping, Nanozyme-based biosensor
11 for organophosphorus pesticide monitoring: Functional design, biosensing
12 strategy, and detection application, *TrAC Trends in Analytical Chemistry*, 2023,
13 165, 117152. <https://doi.org/10.1016/j.trac.2023.117152>.
- 14 4. X. Li, M. Liu, W. Sun, X. Meng, K. Du, K. H. Row and Y. Chang, Visual Strategies of
15 Molecularly Imprinted Sensors for Rapid Warning of Pesticide Residues:
16 Smartphone-Based Visual Detection, *TrAC Trends in Analytical Chemistry*, 2025,
17 <https://doi.org/https://doi.org/10.1016/j.trac.2025.118472118472>.
18 <https://doi.org/10.1016/j.trac.2025.118472>.
- 19 5. R. Umapathi, B. Park, S. Sonwal, G. M. Rani, Y. Cho and Y. S. Huh, Advances in
20 optical-sensing strategies for the on-site detection of pesticides in agricultural
21 foods, *Trends in Food Science & Technology*, 2022, 119, 69–89.
22 <https://doi.org/10.1016/j.tifs.2021.11.018>.
- 23 6. R. Umapathi, S. Sonwal, M. J. Lee, G. Mohana Rani, E.-S. Lee, T.-J. Jeon, S.-M. Kang,
24 M.-H. Oh and Y. S. Huh, Colorimetric based on-site sensing strategies for the
25 rapid detection of pesticides in agricultural foods: New horizons, perspectives, and
26 challenges, *Coordination Chemistry Reviews*, 2021, 446, 214061.
27 <https://doi.org/10.1016/j.ccr.2021.214061>.
- 28 7. R. Umapathi, S. M. Ghoreishian, S. Sonwal, G. M. Rani and Y. S. Huh, Portable
29 electrochemical sensing methodologies for on-site detection of pesticide residues
30 in fruits and vegetables, *Coordination Chemistry Reviews*, 2022, 453, 214305.
31 <https://doi.org/10.1016/j.ccr.2021.214305>.
- 32 8. R. Umapathi, C. Venkateswara Raju, S. Majid Ghoreishian, G. Mohana Rani, K.
33 Kumar, M.-H. Oh, J. Pil Park and Y. Suk Huh, Recent advances in the use of
34 graphitic carbon nitride-based composites for the electrochemical detection of



- 1 hazardous contaminants, *Coordination Chemistry Reviews*, 2022, 470, 214708.
2 <https://doi.org/10.1016/j.ccr.2022.214708>.
- 3 9. R. Umapathi, G. M. Rani, E. Kim, S.-Y. Park, Y. Cho and Y. S. Huh, Sowing kernels
4 for food safety: Importance of rapid on-site detection of pesticide residues in
5 agricultural foods, *Food Frontiers*, 2022, 3, 666–676.
6 <https://doi.org/10.1002/fft2.166>.
- 7 10. R. Umapathi, S. M. Ghoreishian, G. M. Rani, Y. Cho and Y. S. Huh, Review—
8 Emerging Trends in the Development of Electrochemical Devices for the On-Site
9 Detection of Food Contaminants, *ECS Sensors Plus*, 2022, 1, 044601.
10 10.1149/2754-2726/ac9d4a.
- 11 11. T. G. Ambaye, A. Hassani, M. Vaccari, A. Franzetti, S. Prasad, F. Formicola, A.
12 Rosatelli, M. Z. u. Rehman, G. Mohanakrishna, S. V. Ganachari, T. M. Aminabhavi
13 and S. Rtimi, Emerging technologies for the removal of pesticides from
14 contaminated soils and their reuse in agriculture, *Chemosphere*, 2024, 362,
15 142433. <https://doi.org/10.1016/j.chemosphere.2024.142433>.
- 16 12. S. Tanveer, N. Ilyas, N. Akhtar, N. Akhtar, N. Bostan, Z. Hasnain, A. Niaz, G. Zengin,
17 A. Gafur and B. N. Fitriatin, Unlocking the interaction of organophosphorus
18 pesticide residues with ecosystem: Toxicity and bioremediation, *Environmental*
19 *Research*, 2024, 249, 118291. <https://doi.org/10.1016/j.envres.2024.118291>.
- 20 13. A. Jha, D. Pathania, Sonu, B. Damathia, P. Raizada, S. Rustagi, P. Singh, G. M. Rani
21 and V. Chaudhary, Panorama of biogenic nano-fertilizers: A road to sustainable
22 agriculture, *Environmental Research*, 2023, 235, 116456.
23 <https://doi.org/10.1016/j.envres.2023.116456>.
- 24 14. X. Lin, Y. Huang, J. Zou, G. Zhu, L. Pang and Y. Chen, Recent advances in
25 nanozymes for organophosphorus pesticide detection in food, *Chem Commun*
26 (Camb), 2025, 61, 14510–14530. 10.1039/d5cc03464g.
- 27 15. C. Venkateswara Raju, C. Hwan Cho, G. Mohana Rani, V. Manju, R. Umapathi, Y.
28 Suk Huh and J. Pil Park, Emerging insights into the use of carbon-based
29 nanomaterials for the electrochemical detection of heavy metal ions, *Coordination*
30 *Chemistry Reviews*, 2023, 476, 214920. <https://doi.org/10.1016/j.ccr.2022.214920>.
- 31 16. R. Umapathi, C. V. Raju, M. Safarkhani, J. Haribabu, H. U. Lee, G. M. Rani and Y. S.
32 Huh, Versatility of MXene based materials for the electrochemical detection of
33 phenolic contaminants, *Coordination Chemistry Reviews*, 2025, 525, 216305.
34 <https://doi.org/10.1016/j.ccr.2024.216305>.



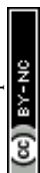
- 1 17. A. T. E. Vilian, R. Umapathi, S.-K. Hwang, Y. S. Huh and Y.-K. Han, Pd–Cu
2 nanospheres supported on Mo₂C for the electrochemical sensing of nitrites,
3 Journal of Hazardous Materials, 2021, 408, 124914.
4 <https://doi.org/10.1016/j.jhazmat.2020.124914>.
- 5 18. A. T. Ezhil Vilian, R. Umapathi, S.-K. Hwang, M. J. Lee, Y. S. Huh and Y.-K. Han,
6 Simple synthesis of a clew-like tungsten carbide nanocomposite decorated with
7 gold nanoparticles for the ultrasensitive detection of tert-butylhydroquinone,
8 Food Chemistry, 2021, 348, 128936.
9 <https://doi.org/10.1016/j.foodchem.2020.128936>.
- 10 19. B. Kariyanna, S. Senthil-Nathan, P. Vasantha-Srinivasan, B. V. Subba Reddy, A.
11 Krishnaiah, N. H. Meenakshi, Y. S. Han, S. Karthi, A. K. Chakravarthy and K. B. Park,
12 Comprehensive insights into pesticide residue dynamics: unraveling impact and
13 management, Chemical and Biological Technologies in Agriculture, 2024, 11, 182.
14 10.1186/s40538-024-00708-4.
- 15 20. R. Singh, A. Sharma, O. Bainik, B. Navyatha, C. Santhosh, E. Banoth, R. Balaji, N.
16 Chandrasekar, S. Kumar and H. K. Daima, Emerging trends, and prospects in
17 Nanozyme engineering to enhance dual-mode sensing applications, Coordination
18 Chemistry Reviews, 2025, 540, 216768. <https://doi.org/10.1016/j.ccr.2025.216768>.
- 19 21. A. Baranwal, R. Shukla and V. Bansal, Nanozyme-enhanced paper-based biosensor
20 technologies, TrAC Trends in Analytical Chemistry, 2024, 172, 117573.
21 <https://doi.org/10.1016/j.trac.2024.117573>.
- 22 22. R. Singh, A. Umapathi, G. Patel, C. Patra, U. Malik, S. K. Bhargava and H. K. Daima,
23 Nanozyme-based pollutant sensing and environmental treatment: Trends,
24 challenges, and perspectives, Science of The Total Environment, 2023, 854,
25 158771. <https://doi.org/10.1016/j.scitotenv.2022.158771>.
- 26 23. L. Yang, X. Xu, Y. Song, J. Huang and H. Xu, Research progress of nanozymes in
27 colorimetric biosensing: Classification, activity and application, Chemical
28 Engineering Journal, 2024, 487, 150612. <https://doi.org/10.1016/j.cej.2024.150612>.
- 29 24. R. Zhang, X. Yan and K. Fan, Nanozymes Inspired by Natural Enzymes, Accounts
30 of Materials Research, 2021, 2, 534–547. 10.1021/accountsmr.1c00074.
- 31 25. M. Liang and X. Yan, Nanozymes: From New Concepts, Mechanisms, and
32 Standards to Applications, Accounts of Chemical Research, 2019, 52, 2190–2200.
33 10.1021/acs.accounts.9b00140.
- 34 26. R. Zhang, X. Yan, L. Gao and K. Fan, Nanozymes expanding the boundaries of



- 1 biocatalysis, *Nature Communications*, 2025, 16, 6817. 10.1038/s41467-025-62063-
2 8.
- 3 27. I. Algov, A. Feiertag and L. Alfonta, Site-specifically wired and oriented glucose
4 dehydrogenase fused to a minimal cytochrome with high glucose sensing
5 sensitivity, *Biosens Bioelectron*, 2021, 180, 113117. 10.1016/j.bios.2021.113117.
- 6 28. K. Ito, J. Okuda-Shimazaki, K. Kojima, K. Mori, W. Tsugawa, R. Asano, K. Ikebukuro
7 and K. Sode, Strategic design and improvement of the internal electron transfer of
8 heme b domain-fused glucose dehydrogenase for use in direct electron transfer-
9 type glucose sensors, *Biosens Bioelectron*, 2021, 176, 112911.
10 10.1016/j.bios.2020.112911.
- 11 29. M. Zandieh and J. Liu, *Nanozymes: Definition, Activity, and Mechanisms*, *Advanced*
12 *Materials*, 2024, 36, 2211041. <https://doi.org/10.1002/adma.202211041>.
- 13 30. Y. Zhang, Y. Yang, Z. Yin, L. Huang and J. Wang, Nanozyme-based wearable
14 biosensors for application in healthcare, *iScience*, 2025, 28, 111763.
15 <https://doi.org/10.1016/j.isci.2025.111763>.
- 16 31. X. Li, J. Chen and S. Dong, *Nanozymes for Energy and Environmental*
17 *Sustainability*, *Advanced Science*, 2026, 13, e19402.
18 <https://doi.org/10.1002/advs.202519402>.
- 19 32. L. Singh, A. Sharma and J. Verma, Emerging green-synthesized nanozymes for
20 microbial applications: toward sustainable antimicrobial technologies, *World*
21 *Journal of Microbiology and Biotechnology*, 2025, 41, 444. 10.1007/s11274-025-
22 04655-6.
- 23 33. F. Li, C. Zhou, J. Zhao, K. Li, M. Yang, N. Cheng, L. Qiu, J. Zhou and L. Li,
24 Nanozyme taxonomy, mechanistic insights, and controllable synthesis strategies:
25 Advances in precision agricultural applications, *Chemical Engineering Journal*,
26 2025, 525, 169555. <https://doi.org/10.1016/j.cej.2025.169555>.
- 27 34. Y. Zhang, G. Tian, J. Lin, Y. Fu and Y. Li, Nanozymes to Enhance Plant
28 Photosynthesis: Mechanisms, Applications, and Future Prospects, *Journal of*
29 *Agricultural and Food Chemistry*, 2026, 74, 6593–6611. 10.1021/acs.jafc.5c12837.
- 30 35. H. Kim, D. Kim, J. Lee, G. M. Rani, Y. Park, S. Han, H. Jeong, K. Shin, T. J. Park, R.
31 Umapathi and Y. S. Huh, Beyond general catalysis: overcoming the specificity
32 limitations of nanozyme for organic contaminants detection, *Coordination*
33 *Chemistry Reviews*, 2026, 559, 217700. <https://doi.org/10.1016/j.ccr.2026.217700>.
- 34 36. B. R. Putra, E. Nurwidayanti, S. Fadilah, M. Khalil, E. Rustami and W. T. Wahyuni,



- 1 MWCNT-OH/graphene composite sensor for nonenzymatic detection of
2 paraoxon-ethyl in agricultural samples, *Carbon Letters*, 2025, 35, 1291–1310.
3 10.1007/s42823-025-00868-9.
- 4 37. Y. Jia, L. Kang, Y. Wu, C. Zhou, D. Li, J. Li and C. Pan, Review on Pesticide Abiotic
5 Stress over Crop Health and Intervention by Various Biostimulants, *Journal of*
6 *Agricultural and Food Chemistry*, 2023, 71, 13595–13611. 10.1021/acs.jafc.3c04013.
- 7 38. V. M. Pathak, V. K. Verma, B. S. Rawat, B. Kaur, N. Babu, A. Sharma, S. Dewali, M.
8 Yadav, R. Kumari, S. Singh, A. Mohapatra, V. Pandey, N. Rana and J. M. Cunill,
9 Current status of pesticide effects on environment, human health and it's eco-
10 friendly management as bioremediation: A comprehensive review, *Frontiers in*
11 *Microbiology*, 2022, Volume 13 - 2022, 10.3389/fmicb.2022.962619.
- 12 39. R. Bhoi, C. Sahu and A. Pradhan, A global review on health risks of pesticide
13 contamination in rice-based agricultural systems, *Discover Applied Sciences*, 2025,
14 7, 1316. 10.1007/s42452-025-07800-w.
- 15 40. W. Zhou, M. Li and V. Achal, A comprehensive review on environmental and
16 human health impacts of chemical pesticide usage, *Emerging Contaminants*, 2025,
17 11, 100410. <https://doi.org/10.1016/j.emcon.2024.100410>.
- 18 41. A. Santana-Mayor, R. Rodríguez-Ramos, A. V. Herrera-Herrera, B. Socas-Rodríguez
19 and M. A. Rodríguez-Delgado, Updated overview of QuEChERS applications in
20 food, environmental and biological analysis (2020–2023), *TrAC Trends in Analytical*
21 *Chemistry*, 2023, 169, 117375. <https://doi.org/10.1016/j.trac.2023.117375>.
- 22 42. L. Kim, D. Lee, H.-K. Cho and S.-D. Choi, Review of the QuEChERS method for the
23 analysis of organic pollutants: Persistent organic pollutants, polycyclic aromatic
24 hydrocarbons, and pharmaceuticals, *Trends in Environmental Analytical Chemistry*,
25 2019, 22, e00063. <https://doi.org/10.1016/j.teac.2019.e00063>.
- 26 43. S. Niell, V. Cesio, J. Hepperle, D. Doerk, L. Kirsch, D. Kolberg, E. Scherbaum, M.
27 Anastassiades and H. Heinzen, QuEChERS-Based Method for the Multiresidue
28 Analysis of Pesticides in Beeswax by LC-MS/MS and GC×GC-TOF, *Journal of*
29 *Agricultural and Food Chemistry*, 2014, 62, 3675–3683. 10.1021/jf405771t.
- 30 44. M. Li, C. Dai, F. Wang, Z. Kong, Y. He, Y. T. Huang and B. Fan, Chemometric-
31 assisted QuEChERS extraction method for post-harvest pesticide determination in
32 fruits and vegetables, *Scientific Reports*, 2017, 7, 42489. 10.1038/srep42489.
- 33 45. J.-H. Joo, S.-H. Kim, J. H. Kim, H.-J. Kang, J. H. Lee, H.-J. Jeon, Y. H. Jang, J.-H. Lee,
34 S.-Y. Lee, S.-J. Park and M.-K. Seo, Recent advances in activated carbon fibers for



- 1 pollutant removal, *Carbon Letters*, 2025, 35, 21–44. 10.1007/s42823-024-00803-4.
- 2 46. X. Lin, Y. Huang, J. Zou, G. Zhu, L. Pang and Y. Chen, Recent advances in
3 nanozymes for organophosphorus pesticide detection in food, *Chemical*
4 *Communications*, 2025, 61, 14510–14530. 10.1039/D5CC03464G.
- 5 47. A. M. Ashrafi, Z. Bytesnikova, J. Barek, L. Richtera and V. Adam, A critical
6 comparison of natural enzymes and nanozymes in biosensing and bioassays,
7 *Biosensors and Bioelectronics*, 2021, 192, 113494.
8 <https://doi.org/10.1016/j.bios.2021.113494>.
- 9 48. S. Liu, J. Zhou, X. Yuan, J. Xiong, M.-H. Zong, X. Wu and W.-Y. Lou, A dual-mode
10 sensing platform based on metal–organic framework for colorimetric and
11 ratiometric fluorescent detection of organophosphorus pesticide, *Food Chemistry*,
12 2024, 432, 137272. <https://doi.org/10.1016/j.foodchem.2023.137272>.
- 13 49. X. Xu, T. Sun, X. Zhou, Z. Liu and L. Zhang, Specific and enzyme-free monitoring
14 of propiconazole pesticide residues in vegetables with a portable nanozyme-
15 based paper sensor, *Food Chemistry*, 2025, 464, 141686.
16 <https://doi.org/10.1016/j.foodchem.2024.141686>.
- 17 50. X. Yan, R. Zou, Q. Lin, Y. Ma, A. Li, X. Sun, G. Lu and H. Li, Glutathione-iron hybrid
18 nanozyme-based colorimetric sensor for specific and stable detection of thiram
19 pesticide on fruit juices, *Food Chemistry*, 2024, 452, 139569.
20 <https://doi.org/10.1016/j.foodchem.2024.139569>.
- 21 51. C. Jin, S. Yang, J. Zheng, F. Chai and M. Tian, A smartphone-assisted portable on-
22 site detection system for organophosphorus pesticides in vegetables and fruits
23 based on all-in-one paper-based sensors: 2,2-Dichlorovinyl dimethyl phosphate as
24 a model, *Food Chemistry*, 2024, 459, 140369.
25 <https://doi.org/10.1016/j.foodchem.2024.140369>.
- 26 52. T. Tian, D. Song, L. Zhang, H. Huang and Y. Li, Facile and selective recognition of
27 sulfonylurea pesticides based on the multienzyme-like activities enhancement of
28 nanozymes combining sensor array, *Journal of Hazardous Materials*, 2024, 469,
29 133847. <https://doi.org/10.1016/j.jhazmat.2024.133847>.
- 30 53. Q. Ma, S. Shen, X. Hou, H. Gong, Y. Zhou, T. Liu and X. Wang, Multi-functional
31 colorimetric sensor arrays for accurate discrimination/quantitation of
32 organophosphorus pesticides based on enhanced bienzymatic activity
33 independent of routine peroxidase mimetics of V2O5-NC with C-O bridges-
34 connected wide interlayer-spacing, *Chemical Engineering Journal*, 2025, 509,



- 1 161222. <https://doi.org/10.1016/j.cej.2025.161222>.
- 2 54. Z. Liu, Y. Feng, Y. Cui, Y. Wang, Z. Fu, J. Zhao, H. Li and C. Sun, Tuning ligand
3 substituents of metal-organic frameworks enhances laccase-like activity for
4 pesticide detection, *Food Chemistry*, 2025, 488, 144892.
5 <https://doi.org/10.1016/j.foodchem.2025.144892>.
- 6 55. Y. Shen, X. Gao, H. Chen, Y. Wei, H. Yang and Y. Gu, Ultrathin C3N4 nanosheets-
7 based oxidase-like 2D fluorescence nanozyme for dual-mode detection of
8 organophosphorus pesticides, *Journal of Hazardous Materials*, 2023, 451, 131171.
9 <https://doi.org/10.1016/j.jhazmat.2023.131171>.
- 10 56. T. Tian, D. Song, L. Zhen, Z. Bi, L. Zhang, H. Huang and Y. Li, Colorimetric –
11 Fluorescence – Photothermal tri-mode sensor array combining the machine
12 learning method for the selective identification of sulfonylurea pesticides,
13 *Biosensors and Bioelectronics*, 2025, 277, 117286.
14 <https://doi.org/10.1016/j.bios.2025.117286>.
- 15 57. Y. Bai, W. Gao, J. Wei, B. Yu, L. Zhang, P. Zhu and J. Yu, Biomimetic cascade
16 intelligent paper chip sensor based on bimetallic porphyrin-based covalent
17 organic framework with triple-enzyme mimetic activities, *Chemical Engineering*
18 *Journal*, 2024, 490, 151628. <https://doi.org/10.1016/j.cej.2024.151628>.
- 19 58. D. Zhu, N. Li, M. Zhang, Y. Wang, F. Li and T. Hou, Hydrolysis enabled specific
20 colorimetric assay of carbosulfan with sensitivity manipulation via metal-doped or
21 metal-free carbon nanozyme, *Biosensors and Bioelectronics*, 2024, 243, 115786.
22 <https://doi.org/10.1016/j.bios.2023.115786>.
- 23 59. T. Zhang, M. Tang, S. Yang, H. Fa, Y. Wang, D. Huo, C. Hou and M. Yang,
24 Development of a novel ternary MOF nanozyme-based smartphone-integrated
25 colorimetric and microfluidic paper-based analytical device for trace glyphosate
26 detection, *Food Chemistry*, 2025, 464, 141780.
27 <https://doi.org/10.1016/j.foodchem.2024.141780>.
- 28 60. X. Wang, X. Zang, X. Wang, W. Zhang, Y. Fang and B. Cui, Molecularly imprinted
29 electrochemiluminescence-colorimetric dual-mode sensor based on Mn@NC
30 nanozyme amplification for the detection of phoxim, *Chemical Engineering*
31 *Journal*, 2024, 500, 156817. <https://doi.org/10.1016/j.cej.2024.156817>.
- 32 61. J. Guo, F. Zhao, Z. Yue and Z. Lei, Acetylcholinesterase-free colorimetric sensing
33 platform for carbosulfan detection based on hollow PDA/MnO₂ nanozyme, *Food*
34 *Chemistry*, 2025, 465, 142075. <https://doi.org/10.1016/j.foodchem.2024.142075>.



- 1 62. Y. Ren, L. Cao, H. Li, R. Jiao, Y. Zhan, X. Zhang, X. Yu, M. Li, W. Wu, Z. Liang, G. Li,
2 X. Xia, D. Zhang, N. Ling and Y. Ye, Trimetal–Organic Framework-Derived Diatomic
3 Nanozymes-Driven Cascaded Signal Sensitization Intelligent Flexible Sensors for In
4 Situ Detection of the Pesticide Residue, *ACS Nano*, 2025, 19, 23703–23718.
5 10.1021/acsnano.5c03245.
- 6 63. N. Guo, L. Jiao, L. Hu, D. Yan, C. Chen, P. Zong, X. Shi, Y. Zhai, Z. Zhu and X. Lu,
7 Deciphering Pyridinic N–Mediated Inhibition Mode in Platinum Nanozymes for
8 Pesticide Distinction, *Small*, 2025, 21, 2505731.
9 <https://doi.org/10.1002/sml.202505731>.
- 10 64. H. Zhu, B. Liu, J. Pan, L. Xu, J. Liu, P. Hu, D. Du, Y. Lin and X. Niu, Redox
11 interference-free bimodal paraoxon sensing enabled by an aggregation-induced
12 emission nanozyme catalytically hydrolyzing phosphoesters specifically, *Biosensors
13 and Bioelectronics*, 2025, 267, 116756. <https://doi.org/10.1016/j.bios.2024.116756>.
- 14 65. J. Li, M. Gao, X. Xia, Y. Cen, F. Wei, J. Yang, L. Wang, Q. Hu and G. Xu, Spherical
15 Hydrogel Sensor Based on PB@Fe-COF@Au Nanoparticles with Triplet
16 Peroxidase-like Activity and Multiple Capture Sites for Effective Detection of
17 Organophosphorus Pesticides, *ACS Applied Materials & Interfaces*, 2023, 15,
18 6473–6485. 10.1021/acsam.2c19921.
- 19 66. D. Li, J. Li, C. Wu, H. Liu, M. Zhao, H. Shi, Y. Zhang and T. Wang, Smartphone-
20 assisted colorimetric biosensor for the determination of organophosphorus
21 pesticides on the peel of fruits, *Food Chemistry*, 2024, 443, 138459.
22 <https://doi.org/10.1016/j.foodchem.2024.138459>.
- 23 67. J. Ge, L. Yang, Z. Li, Y. Wan, D. Mao, R. Deng, Q. Zhou, Y. Yang and W. Tan, A
24 colorimetric smartphone-based platform for pesticides detection using Fe-N/C
25 single-atom nanozyme as oxidase mimetics, *Journal of Hazardous Materials*, 2022,
26 436, 129199. <https://doi.org/10.1016/j.jhazmat.2022.129199>.
- 27 68. Y. Guo, X. Li, P. Shen, X. Li, Y. Cheng and K. Chu, Dendritic-like MXene quantum
28 dots@CuNi as an efficient peroxidase candidate for colorimetric determination of
29 glyphosate, *Journal of Colloid and Interface Science*, 2024, 661, 533–543.
30 <https://doi.org/10.1016/j.jcis.2024.01.177>.
- 31 69. M. Sun, L. Zhao, T. Liu, Z. Lu, G. Su, C. Wu, C. Song, R. Deng, M. He, H. Rao and Y.
32 Wang, Construction of CuO/Fe₂O₃ Nanozymes for Intelligent Detection of
33 Glufosinate and Chlortetracycline Hydrochloride, *ACS Applied Materials &
34 Interfaces*, 2023, 15, 54466–54477. 10.1021/acsam.3c12157.



- 1 70. Y. Zhang, J. Yang, W. Gao, S. G. Liu, Q. Zhao, Z. Fu and X. Shi, A smartphone-
2 integrated colorimetric sensor for sensitive detection of organophosphorus
3 pesticides based on large-scale synthesized Fe-N/C single-atom nanozymes,
4 *Sensors and Actuators B: Chemical*, 2024, 403, 135130.
5 <https://doi.org/10.1016/j.snb.2023.135130>.
- 6 71. A. Li, H. Li, Y. Ma, T. Wang, X. Liu, C. Wang, F. Liu, P. Sun, X. Yan and G. Lu,
7 Bioinspired laccase-mimicking catalyst for on-site monitoring of thiram in paper-
8 based colorimetric platform, *Biosensors and Bioelectronics*, 2022, 207, 114199.
9 <https://doi.org/10.1016/j.bios.2022.114199>.
- 10 72. X. Zhan, Z. Ding, F. Liu, K. Chu and Y. Guo, Synthesis of MXene quantum dot-
11 modified urchin-like NiCoP/NF nanostructures as high-performance peroxidase
12 mimics for sensitive colorimetric detection of glyphosate, *Food Chemistry*, 2025,
13 489, 144966. <https://doi.org/10.1016/j.foodchem.2025.144966>.
- 14 73. M. Zhang, Y. Wang, N. Li, D. Zhu and F. Li, Specific detection of fungicide
15 thiophanate-methyl: A smartphone colorimetric sensor based on target-regulated
16 oxidase-like activity of copper-doped carbon nanozyme, *Biosensors and*
17 *Bioelectronics*, 2023, 237, 115554. <https://doi.org/10.1016/j.bios.2023.115554>.
- 18 74. Z. Shen, D. Xu, G. Wang, L. Geng, R. Xu, G. Wang, Y. Guo and X. Sun, Novel
19 colorimetric aptasensor based on MOF-derived materials and its applications for
20 organophosphorus pesticides determination, *Journal of Hazardous Materials*,
21 2022, 440, 129707. <https://doi.org/10.1016/j.jhazmat.2022.129707>.
- 22 75. X. Luo, G. Huang, C. Bai, C. Wang, Y. Yu, Y. Tan, C. Tang, J. Kong, J. Huang and Z.
23 Li, A versatile platform for colorimetric, fluorescence and photothermal multi-
24 mode glyphosate sensing by carbon dots anchoring ferrocene metal-organic
25 framework nanosheet, *Journal of Hazardous Materials*, 2023, 443, 130277.
26 <https://doi.org/10.1016/j.jhazmat.2022.130277>.
- 27 76. Z. Yan, J. Zhong, Z. Liao, P. Xu and P. Qiu, Single-enzymatic (AChE/ChOx)
28 colorimetric detection of organophosphorus pesticides based on controllable
29 nanoparticles supported by FeCoCu flower-like structure with peroxidase-like
30 activity, *Sensors and Actuators B: Chemical*, 2025, 424, 136878.
31 <https://doi.org/10.1016/j.snb.2024.136878>.
- 32 77. M. Sun, S. Huang, G. Su, X. Wang, Z. Lu, Y. Wang, T. Liu, Y. Jiang, C. Song and H.
33 Rao, Synthesis of pH-switchable Pt/Co₃O₄ nanoflowers: Catalytic mechanism,
34 four-enzyme activity and smartphone biosensing applications, *Chemical*



- 1 Engineering Journal, 2022, 437, 134414. <https://doi.org/10.1016/j.ccej.2021.134414>.
- 2 78. H. Zhu, P. Liu, L. Xu, X. Li, P. Hu, B. Liu, J. Pan, F. Yang and X. Niu, Nanozyme-
3 Participated Biosensing of Pesticides and Cholinesterases: A Critical Review,
4 Biosensors (Basel), 2021, 11, 10.3390/bios11100382.
- 5 79. Y. Huang, J. Ren and X. Qu, Nanozymes: Classification, Catalytic Mechanisms,
6 Activity Regulation, and Applications, Chemical Reviews, 2019, 119, 4357–4412.
7 10.1021/acs.chemrev.8b00672.
- 8 80. R. Hou, N. Yin, Y. Wang, S. Song and H. Zhang, Nanozymes: recent advances for
9 sustainable agricultural development, Nanoscale Horizons, 2025,
10 <https://doi.org/10.1039/D5NH00281H>[10.1039/D5NH00281H](https://doi.org/10.1039/D5NH00281H).
- 11 81. A. Shamsabadi, T. Haghighi, S. Carvalho, L. C. Frenette and M. M. Stevens, The
12 Nanozyme Revolution: Enhancing the Performance of Medical Biosensing
13 Platforms, Advanced Materials, 2024, 36, 2300184.
14 <https://doi.org/10.1002/adma.202300184>.
- 15 82. S. Nayak, M. Aggarwal and P. Das, Bromine-doped carbon dot: concentration-
16 dependent multicolor emission, nanozyme activity, and visible-light-induced
17 photodynamic bacterial inactivation, Carbon Letters, 2025, 35, 593–605.
18 10.1007/s42823-024-00796-0.
- 19 83. Y. Ouyang, M. P. O'Hagan and I. Willner, Functional catalytic nanoparticles
20 (nanozymes) for sensing, Biosensors and Bioelectronics, 2022, 218, 114768.
21 <https://doi.org/10.1016/j.bios.2022.114768>.
- 22 84. E. M. Hamed, K. Kustomo, M. M. ElSaady, X. Wu, F. M. Fung and S. F. Y. Li,
23 Organic-Dominated Nanozymes for Pesticide Detection: Toward Sustainable
24 Agricultural Monitoring, Journal of Agricultural and Food Chemistry, 2025,
25 <https://doi.org/10.1021/acs.jafc.5c06721>[10.1021/acs.jafc.5c06721](https://doi.org/10.1021/acs.jafc.5c06721).
- 26 85. M. Levine, Fluorescence-Based Sensing of Pesticides Using Supramolecular
27 Chemistry, Frontiers in Chemistry, 2021, Volume 9 - 2021,
28 10.3389/fchem.2021.616815.
- 29 86. G. Kalaiarasi, A. Shanmughan, Y. Jegadeesan, M. K. Noushija, A. V. Krishna, H.
30 Gangadharan, D. Umadevi and S. Shanmugaraju, Recent progress in fluorescence-
31 based chemosensing of pesticides, Sensors & Diagnostics, 2025, 4, 460–488.
32 10.1039/d4sd00364k.
- 33 87. S. Paul, P. Daga and N. Dey, Exploring Various Photochemical Processes in Optical
34 Sensing of Pesticides by Luminescent Nanomaterials: A Concise Discussion on



- 1 Challenges and Recent Advancements, ACS Omega, 2023, 8, 44395–44423.
2 10.1021/acsomega.3c02753.
- 3 88. S. Zhang, Z. Wang, Y. Feng, C. Jiang, H. Li, Z. Yu, Y. Xiao, R. Hou, X. Wan and Y.
4 Liu, A novel fluorescent and photothermal probe based on nanozyme-mediated
5 cascade reaction for detecting organophosphorus pesticide residues, Talanta,
6 2024, 279, 126620. <https://doi.org/10.1016/j.talanta.2024.126620>.
- 7 89. J. Wu, Q. Yang, Q. Li, H. Li and F. Li, Two-Dimensional MnO₂ Nanozyme-Mediated
8 Homogeneous Electrochemical Detection of Organophosphate Pesticides without
9 the Interference of H₂O₂ and Color, Analytical Chemistry, 2021, 93, 4084–4091.
10 10.1021/acs.analchem.0c05257.
- 11 90. F. Faghiri, M. Hajjami and F. Ghorbani, Development of a sensing system based
12 on coupling magnetic solid phase extraction and colorimetric detection for
13 determination of organophosphorus pesticides in fruit extract and environmental
14 sample, Sensors and Actuators B: Chemical, 2021, 343, 130157.
15 <https://doi.org/10.1016/j.snb.2021.130157>.
- 16 91. D. Song, L. Lei, T. Tian, X. Yang, L. Wang, Y. Li and H. Huang, A novel strategy for
17 identification of pesticides in different categories by concentration-independent
18 model based on a nanozyme with multienzyme-like activities, Biosensors and
19 Bioelectronics, 2023, 237, 115458. <https://doi.org/10.1016/j.bios.2023.115458>.
- 20 92. H. Wu, J.-H. Li, W.-C. Yang, T. Wen, J. He, Y.-Y. Gao, G.-F. Hao and W.-C. Yang,
21 Nonmetal-doped quantum dot-based fluorescence sensing facilitates the
22 monitoring of environmental contaminants, Trends in Environmental Analytical
23 Chemistry, 2023, 40, e00218. <https://doi.org/10.1016/j.teac.2023.e00218>.
- 24 93. W. Hu, Q. Chen, H. Li, Q. Ouyang and J. Zhao, Fabricating a novel label-free
25 aptasensor for acetamiprid by fluorescence resonance energy transfer between
26 NH₂-NaYF₄: Yb, Ho@SiO₂ and Au nanoparticles, Biosensors and Bioelectronics,
27 2016, 80, 398–404. <https://doi.org/10.1016/j.bios.2016.02.001>.
- 28 94. Z.-D. Zhou, C.-Y. Wang, G.-S. Zhu, B. Du, B.-Y. Yu and C.-C. Wang, Water-stable
29 europium(III) and terbium(III)-metal organic frameworks as fluorescent sensors to
30 detect ions, antibiotics and pesticides in aqueous solutions, Journal of Molecular
31 Structure, 2022, 1251, 132009. <https://doi.org/10.1016/j.molstruc.2021.132009>.
- 32 95. H. Qiu, H. Yang, X. Gao, C. Nie, Y. Gu and Y. Shen, Inner filter effect-based
33 fluorescence assays toward environmental pesticides and antibiotics, Coordination
34 Chemistry Reviews, 2023, 493, 215305. <https://doi.org/10.1016/j.ccr.2023.215305>.



- 1 96. S. Li, J. Wu, S. Zhang, T. Jiao, J. Wei, X. Chen, Q. Chen and Q. Chen, Inner filter
2 effect-based upconversion nanosensor for rapid detection of thiram pesticides
3 using upconversion nanoparticles and dithizone–cadmium complexes, *Food*
4 *Chemistry*, 2024, 434, 137438. <https://doi.org/10.1016/j.foodchem.2023.137438>.
- 5 97. S. Tripathi and M. Chakravarty, Efficient dual-phase visual detection of pesticides
6 in real samples with electron-rich emitters carrying multiple twists, *Materials*
7 *Advances*, 2025, 6, 5928–5939. 10.1039/d5ma00630a.
- 8 98. D. Escudero, Revising Intramolecular Photoinduced Electron Transfer (PET) from
9 First-Principles, *Accounts of Chemical Research*, 2016, 49, 1816–1824.
10 10.1021/acs.accounts.6b00299.
- 11 99. Y.-S. Chen, Z.-W. Chen, Y.-W. Yuan, K.-C. Chen and C.-P. Liu, Fluorescence
12 Quenchers Manipulate the Peroxidase-like Activity of Gold-Based Nanomaterials,
13 *ACS Omega*, 2020, 5, 24487–24494. 10.1021/acsomega.0c02956.
- 14 100. Y. Chang, Z. Zhang, J. Hao, W. Yang and J. Tang, A simple label free colorimetric
15 method for glyphosate detection based on the inhibition of peroxidase-like
16 activity of Cu(II), *Sensors and Actuators B: Chemical*, 2016, 228, 410–415.
17 <https://doi.org/10.1016/j.snb.2016.01.048>.
- 18 101. M. Zhu, X. Zhu, M. Chen, X. Pang, Y. Hong and Y. Wang, A novel Cu²⁺-
19 coordinated fluorescent sensing system for specific detection of glyphosate and
20 its applications in environmental and biological systems, *Journal of Hazardous*
21 *Materials*, 2025, 488, 137424. <https://doi.org/10.1016/j.jhazmat.2025.137424>.
- 22 102. X. Li, Z. Wang, J. He, H. Al-Mashriqi, J. Chen and H. Qiu, Recent advances in
23 emerging nanozymes with aggregation-induced emission, *Chemical Science*, 2025,
24 16, 29–42. 10.1039/D4SC05709K.
- 25 103. Y. Huang, L. Gong, C. Xie, W. Qin, M. Wang, L. Hu and Z. Xia, Anchoring
26 biomimetic Zn site in metal–organic framework nanozyme to enhance
27 phosphatase-like catalytic activity for discrimination of organophosphorus
28 pesticides, *Chemical Engineering Journal*, 2025, 506, 160046.
29 <https://doi.org/10.1016/j.cej.2025.160046>.
- 30 104. D. Song, T. Tian, L. Wang, Y. Zou, L. Zhao, J. Xiao, H. Huang and Y. Li, Multi-signal
31 sensor array based on a fluorescent nanozyme for broad-spectrum screening of
32 pesticides, *Chemical Engineering Journal*, 2024, 482, 148784.
33 <https://doi.org/10.1016/j.cej.2024.148784>.
- 34 105. S.-H. Wen, H. Zhang, S. Yu, J. Ma, J.-J. Zhu and Y. Zhou, Nanozyme coating-gated



- 1 multifunctional COF composite based dual-ratio enhanced dual-mode sensor for
2 highly sensitive and reliable detection of organophosphorus pesticides in real
3 samples, *Journal of Hazardous Materials*, 2024, 480, 135791.
4 <https://doi.org/10.1016/j.jhazmat.2024.135791>.
- 5 106. J. Liu, X. Yu, Y. Zhou, L. Sun, Y. Liu and J. Li, Breaking the conventional: Ligand-
6 triggered Zn-MOF nanozyme with unusual oxidase activity for dual-channel
7 sensing of benfuracarb, *Biosensors and Bioelectronics*, 2025, 280, 117441.
8 <https://doi.org/10.1016/j.bios.2025.117441>.
- 9 107. C.-L. Yang, L.-H. Yu, Y.-H. Pang and X.-F. Shen, A ratiometric fluorescence sensor
10 for detection of organophosphorus pesticides based on enzyme-regulated
11 multifunctional Fe-based metal-organic framework, *Talanta*, 2024, 278, 126516.
12 <https://doi.org/10.1016/j.talanta.2024.126516>.
- 13 108. J.-J. Zhu, H.-Y. Niu, C. Yi, K.-X. Wang and C.-G. Niu, Dual-mode detection of
14 malathion under neutral conditions: Integrated application of specific nanozyme
15 and time-resolved europium probe, *Sensors and Actuators B: Chemical*, 2025, 435,
16 137650. <https://doi.org/10.1016/j.snb.2025.137650>.
- 17 109. Z. Ding, X. Zhan, F. Liu, K. Chu and Y. Guo, MXene quantum dots coupled with
18 amorphous CoOX: A high-performance peroxidase mimic for direct detection of
19 the pesticide profenofos, *Food Bioscience*, 2025, 68, 106658.
20 <https://doi.org/10.1016/j.fbio.2025.106658>.
- 21 110. S. Liu, H. Yu, S. Zhu and X.-E. Zhao, Copper-based fluorescent nanozyme used to
22 construct a ratiometric sensor for visual detection of thiophanate methyl, *Talanta*,
23 2025, 285, 127417. <https://doi.org/10.1016/j.talanta.2024.127417>.
- 24 111. J. Wang, Y. Huang, Z. Kuai, Y. Zhang, Q. Shen, P. Tian, W. Nong, W. Jiang, Y. He,
25 N. Ran, Y. Yin, T. Li and Q. Luo, Ultrasensitive and rapid detection of
26 organophosphates using a dual-signal naked-eye hydrogel sensor based on
27 acetylcholinesterase inhibition, *International Journal of Biological Macromolecules*,
28 2024, 283, 137778. <https://doi.org/10.1016/j.ijbiomac.2024.137778>.
- 29 112. Y. Zhao, H. Chen, C. Chang, L. Xu and L. Han, Clusterization-triggered emission
30 artificial enzyme (CTEzyme) with specific activity inhibition boosting ratiometric
31 fluorescence detection of pesticide, *Sensors and Actuators B: Chemical*, 2025, 432,
32 137491. <https://doi.org/10.1016/j.snb.2025.137491>.
- 33 113. Y. Zhang, L. Xue, F. Du, S. Yang, T. Liu, T. Wu and X. Jiang, Synergistic
34 ultrasensitive detection of paraoxon via a photoelectrochemical-fluorescent dual-



- 1 mode aptasensor with S-scheme Zn_{0.1}Cd_{0.9}S/Cu₂MoS₄ heterostructure and Zr-
2 MOF nanozyme, *Sensors and Actuators B: Chemical*, 2026, 449, 139164.
3 <https://doi.org/10.1016/j.snb.2025.139164>.
- 4 114. N. Bagheri, A. Khataee, J. Hassanzadeh and B. Habibi, Sensitive biosensing of
5 organophosphate pesticides using enzyme mimics of magnetic ZIF-8,
6 *Spectrochimica Acta Part A: Molecular and Biomolecular Spectroscopy*, 2019, 209,
7 118–125. <https://doi.org/10.1016/j.saa.2018.10.039>.
- 8 115. V. G. Panferov and J. Liu, Optical and Catalytic Properties of Nanozymes for
9 Colorimetric Biosensors: Advantages, Limitations, and Perspectives, *Advanced*
10 *Optical Materials*, 2024, 12, 2401318. <https://doi.org/10.1002/adom.202401318>.
- 11 116. D. Luo, X. Huang, B. Liu, W. Zou and Y. Wu, Facile Colorimetric Nanozyme Sheet
12 for the Rapid Detection of Glyphosate in Agricultural Products Based on Inhibiting
13 Peroxidase-Like Catalytic Activity of Porous Co₃O₄ Nanoplates, *Journal of*
14 *Agricultural and Food Chemistry*, 2021, 69, 3537–3547. 10.1021/acs.jafc.0c08208.
- 15 117. P. Weerathunge, R. Ramanathan, R. Shukla, T. K. Sharma and V. Bansal, Aptamer-
16 Controlled Reversible Inhibition of Gold Nanozyme Activity for Pesticide Sensing,
17 *Analytical Chemistry*, 2014, 86, 11937–11941. 10.1021/ac5028726.
- 18 118. Z. Wu, Z. Hao, Y. Chai, A. Li, C. Wang, X. Zhang, H. Chen and C. Lu, Near-infrared-
19 excitable acetylcholinesterase-activated fluorescent probe for sensitive and anti-
20 interference detection of pesticides in colored food, *Biosensors and Bioelectronics*,
21 2023, 233, 115341. <https://doi.org/10.1016/j.bios.2023.115341>.
- 22 119. D. Li, S. L. Garisto, P.-J. J. Huang, J. Yang, B. Liu and J. Liu, Fluorescent detection of
23 fluoride by CeO₂ nanozyme oxidation of Amplex red, *Inorganic Chemistry*
24 *Communications*, 2019, 106, 38–42. 10.1016/j.inoche.2019.05.028.
- 25 120. C. Shang, Q. Wang, H. Tan, S. Lu, S. Wang, Q. Zhang, L. Gu, J. Li, E. Wang and S.
26 Guo, Defective PtRuTe As Nanozyme with Selectively Enhanced Peroxidase-like
27 Activity, *JACS Au*, 2022, 2, 2453–2459. 10.1021/jacsau.2c00495.
- 28 121. J. Wei, L. Yang, M. Luo, Y. Wang and P. Li, Nanozyme-assisted technique for dual
29 mode detection of organophosphorus pesticide, *Ecotoxicology and Environmental*
30 *Safety*, 2019, 179, 17–23. <https://doi.org/10.1016/j.ecoenv.2019.04.041>.
- 31 122. D. Wei, M. Li, Y. Wang, N. Zhu, X. Hu, B. Zhao, Z. Zhang and D. Yin, Encapsulating
32 gold nanoclusters into metal–organic frameworks to boost luminescence for
33 sensitive detection of copper ions and organophosphorus pesticides, *Journal of*
34 *Hazardous Materials*, 2023, 441, 129890.



1 <https://doi.org/10.1016/j.jhazmat.2022.129890>.

- 2 123. Q. Fu, S. Tian, S. Wang, G. Sun, M. Wang and K. Fan, Advances in Metal–Organic
3 Framework-Based Nanozymes for Biosensing and Point-of-Care Testing
4 Applications, *ACS Applied Nano Materials*, 2025, 8, 17339–17355.
5 10.1021/acsnm.5c02319.
- 6 124. S. Alagarsamy, B. Ramachandran and Y.-C. Liao, Next-generation electrochemical-
7 optical sensors: Advanced and innovative strategies for pesticide detection,
8 *Chemical Engineering Journal*, 2025, 520, 10.1016/j.cej.2025.166021.
- 9 125. B. M. da Costa Filho, A. C. Duarte and T. A. P. Rocha-Santos, Environmental
10 monitoring approaches for the detection of organic contaminants in marine
11 environments: A critical review, *Trends in Environmental Analytical Chemistry*,
12 2022, 33, 10.1016/j.teac.2022.e00154.
- 13 126. L. C. Clark, Jr. and C. Lyons, Electrode systems for continuous monitoring in
14 cardiovascular surgery, *Ann N Y Acad Sci*, 1962, 102, 29–45. 10.1111/j.1749-
15 6632.1962.tb13623.x.
- 16 127. C. Gu, S. Ji, Z. Chen, W. Yang, Y. Deng, M. Zhao, W. Huang, W. Yang and W. Xu,
17 Enrichment-catalytic synergistically enhanced electrochemiluminescence sensors
18 based on IRMOF-3/CdTe for ultrasensitive detection of organophosphorus
19 pesticides, *Biosens Bioelectron*, 2025, 279, 117398. 10.1016/j.bios.2025.117398.
- 20 128. Q. Hou, M. Rooman and F. Pucci, Enzyme Stability-Activity Trade-Off: New Insights
21 from Protein Stability Weaknesses and Evolutionary Conservation, *J Chem Theory
22 Comput*, 2023, 19, 3664–3671. 10.1021/acs.jctc.3c00036.
- 23 129. A. Kunka, S. M. Marques, M. Havlasek, M. Vasina, N. Velatova, L. Cengelova, D.
24 Kovar, J. Damborsky, M. Marek, D. Bednar and Z. Prokop, Advancing Enzyme's
25 Stability and Catalytic Efficiency through Synergy of Force-Field Calculations,
26 Evolutionary Analysis, and Machine Learning, *ACS Catal*, 2023, 13, 12506–12518.
27 10.1021/acscatal.3c02575.
- 28 130. L. Gao, J. Zhuang, L. Nie, J. Zhang, Y. Zhang, N. Gu, T. Wang, J. Feng, D. Yang, S.
29 Perrett and X. Yan, Intrinsic peroxidase-like activity of ferromagnetic nanoparticles,
30 *Nature Nanotechnology*, 2007, 2, 577–583. 10.1038/nnano.2007.260.
- 31 131. X. Zhu, L. Lin, R. Wu, Y. Zhu, Y. Sheng, P. Nie, P. Liu, L. Xu and Y. Wen, Portable
32 wireless intelligent sensing of ultra-trace phytohormone alpha-naphthalene acetic
33 acid using self-assembled phosphorene/Ti(3)C(2)-MXene nanohybrid with high
34 ambient stability on laser induced porous graphene as nanozyme flexible



- 1 electrode, *Biosens Bioelectron*, 2021, 179, 113062. 10.1016/j.bios.2021.113062.
- 2 132. W. Wu and J. Li, Recent progress on nanozymes in electrochemical sensing,
3 *Journal of Electroanalytical Chemistry*, 2023, 936, 10.1016/j.jelechem.2023.117391.
- 4 133. W. Han, P. Wei, L. Xie, L. Zhu, B. He and X. Cao, Heteroatom modulation of nickel
5 single-atom nanozymes for enhanced interfacial catalytic activity in sensitive
6 electrochemical detection of herbicides, *Chemical Engineering Journal*, 2025, 505,
7 159557. <https://doi.org/10.1016/j.cej.2025.159557>.
- 8 134. L. Geng, X. Sun, L. Wang, F. Liu, S. Hu, S. Zhao and F. Ye, Analyte-induced laccase-
9 mimicking activity inhibition and conductivity enhancement of electroactive
10 nanozymes for ratiometric electrochemical detection of thiram, *J Hazard Mater*,
11 2024, 463, 132936. 10.1016/j.jhazmat.2023.132936.
- 12 135. K. Niu, J. Chen and X. Lu, Versatile biomimetic catalyst functionalized nanozymes
13 for electrochemical sensing, *Chemical Engineering Journal*, 2023, 475,
14 10.1016/j.cej.2023.146491.
- 15 136. X. Ruan, Y. Wang, E. Y. Kwon, L. Wang, N. Cheng, X. Niu, S. Ding, B. J. Van Wie, Y.
16 Lin and D. Du, Nanomaterial-enhanced 3D-printed sensor platform for
17 simultaneous detection of atrazine and acetochlor, *Biosensors and Bioelectronics*,
18 2021, 184, 113238. <https://doi.org/10.1016/j.bios.2021.113238>.
- 19 137. H. Zeng, H. Chen, B. Yang, J. Zeng, L. Meng, D. Shi, L. Chen and Y. Huang, Highly-
20 oxidizing Au@MnO(2-X) nanozymes mediated homogeneous electrochemical
21 detection of organophosphorus independent of dissolved oxygen, *J Hazard Mater*,
22 2023, 459, 132116. 10.1016/j.jhazmat.2023.132116.
- 23 138. R. Wang, B. Li, G. Li, Q. Shen and L. Zou, NiCoFeS/rGO nanozyme-mediated
24 multifunctional homogeneous sensing system for ultrasensitive electrochemical
25 assay of pesticides residues in fruits and vegetables, *Sensors and Actuators B:*
26 *Chemical*, 2025, 422, 10.1016/j.snb.2024.136664.
- 27 139. Y. Sun, J. Wei, J. Zou, Z. Cheng, Z. Huang, L. Gu, Z. Zhong, S. Li, Y. Wang and P. Li,
28 Electrochemical detection of methyl-paraoxon based on bifunctional cerium oxide
29 nanozyme with catalytic activity and signal amplification effect, *J Pharm Anal*,
30 2021, 11, 653–660. 10.1016/j.jpha.2020.09.002.
- 31 140. M. Khairy, H. A. Ayoub and C. E. Banks, Non-enzymatic electrochemical platform
32 for parathion pesticide sensing based on nanometer-sized nickel oxide modified
33 screen-printed electrodes, *Food Chem*, 2018, 255, 104–111.
34 10.1016/j.foodchem.2018.02.004.



- 1 141. G. Wang, J. Liu, H. Dong, L. Geng, J. Sun, J. Liu, J. Dong, Y. Guo and X. Sun, A
2 dual-mode biosensor featuring single-atom Fe nanozyme for multi-pesticide
3 detection in vegetables, *Food Chem*, 2024, 437, 137882.
4 10.1016/j.foodchem.2023.137882.
- 5 142. X. Zhang, N. Hao, S. Liu, K. Wei, C. Ma, J. Pan and S. Feng, Construction of
6 phosphatase-like COF-OMe@Valine-CeO₂ nanozymes for ultrasensitive
7 electrochemical detection of organophosphorus pesticides, *Sensors and Actuators*
8 *B: Chemical*, 2024, 417, 10.1016/j.snb.2024.136068.
- 9 143. X. Pang, G. I. N. Waterhouse, R. Wang, X. Qiao, Y. Sun and Z. Xu, Bifunctional
10 ZrO₂@ZIF-90 nanozyme with high phosphohydrolase activity for sensitive
11 electrochemical detection of methyl parathion, *Food Science and Human*
12 *Wellness*, 2025, 14, 9250095. 10.26599/FSHW.2024.9250095.
- 13 144. X. Zhang, C. Xue, H. Cao, Y. Wu, B. Yang, T. Zhou, W. Zhai and J. Deng, Ultra-small
14 CuOx/GDYO nanozyme with boosting peroxidase-like activity via electrochemical
15 strategy: Toward applicable colorimetric detection of organophosphate pesticides,
16 *Talanta*, 2024, 279, 126639. <https://doi.org/10.1016/j.talanta.2024.126639>.
- 17 145. L. Yu, J. Chang, X. Zhuang, H. Li, T. Hou and F. Li, Two-Dimensional Cobalt-Doped
18 Ti(3)C(2) MXene Nanozyme-Mediated Homogeneous Electrochemical Strategy for
19 Pesticides Assay Based on In Situ Generation of Electroactive Substances, *Anal*
20 *Chem*, 2022, 94, 3669–3676. 10.1021/acs.analchem.1c05300.
- 21 146. C. Praharaj, S. Singh, P. Tripathi and S. Nara, Investigating gold nanorod-mediated
22 hydrolysis of acetylthiocholine: a way for electrochemical detection of
23 organophosphate pesticides, *Environmental Science: Nano*, 2025, 12, 1558–1569.
24 10.1039/d4en00913d.
- 25 147. G. Chen, G. Liu, H. Jia, X. Cui, Y. Wang, D. Li, W. Zheng, Y. She, D. Xu, X. Huang, A.
26 M. Abd El-Aty, J. Sun, H. Liu, Y. Zou, J. Wang, M. Jin and B. D. Hammock, A
27 sensitive bio-barcode immunoassay based on bimetallic Au@Pt nanozyme for
28 detection of organophosphate pesticides in various agro-products, *Food Chem*,
29 2021, 362, 130118. 10.1016/j.foodchem.2021.130118.
- 30 148. M. Zandieh and J. Liu, Nanozyme Catalytic Turnover and Self-Limited Reactions,
31 *ACS Nano*, 2021, 15, 15645–15655. 10.1021/acsnano.1c07520.
- 32 149. Z. Yang, J. Guo, L. Wang, J. Zhang, L. Ding, H. Liu and X. Yu, Nanozyme-Enhanced
33 Electrochemical Biosensors: Mechanisms and Applications, *Small*, 2024, 20,
34 e2307815. 10.1002/sml.202307815.



- 1 150. Q. Liu, A. Zhang, R. Wang, Q. Zhang and D. Cui, A Review on Metal- and Metal
2 Oxide-Based Nanozymes: Properties, Mechanisms, and Applications, *Nanomicro*
3 *Lett*, 2021, 13, 154. 10.1007/s40820-021-00674-8.
- 4 151. Z. Guo, J. Hong, N. Song and M. Liang, Single-Atom Nanozymes: From Precisely
5 Engineering to Extensive Applications, *Accounts of Materials Research*, 2024, 5,
6 347–357. 10.1021/accountsmr.3c00250.
- 7 152. Q. Zhao, X. Zheng, L. Xing, Y. Tang, X. Zhou, L. Hu, W. Yao and Z. Yan, 2D
8 Co(3)O(4) stabilizing Rh nano composites developed for visual sensing bioactive
9 urea and toxic p-aminophenol in practice by synergetic-reinforcing oxidase
10 activity, *J Hazard Mater*, 2021, 409, 125019. 10.1016/j.jhazmat.2020.125019.
- 11 153. L. Fu, D. Yu, D. Zou, H. Qian and Y. Lin, Engineering the Stability of Nanozyme-
12 Catalyzed Product for Colorimetric Logic Gate Operations, *Molecules*, 2021, 26,
13 10.3390/molecules26216494.
- 14 154. S. Li, C. Pang, X. Ma, Y. Zhang, Z. Xu, J. Li, M. Zhang and M. Wang, Microfluidic
15 paper-based chip for parathion-methyl detection based on a double catalytic
16 amplification strategy, *Microchimica Acta*, 2021, 188, 438. 10.1007/s00604-021-
17 05084-6.
- 18 155. T. Sahare, N. Singh, B. N. Sahoo and A. Joshi, Smartphone-enhanced nanozyme
19 sensors: Colorimetric and fluorescence sensing techniques, *Biosensors and*
20 *Bioelectronics: X*, 2024, 21, 100544. <https://doi.org/10.1016/j.biosx.2024.100544>.
- 21 156. F. P. Gbonyea, J. Wu, M. Li, M. Liang, M. Zhang, X. Zhu, X. Li, S. He and P. Liu,
22 Smartphone-integrated Nanozyme approaches for rapid and on-site detection:
23 Empowering smart food safety, *Food Chemistry*, 2025, 486, 144678.
24 <https://doi.org/10.1016/j.foodchem.2025.144678>.
- 25 157. H. Dong, W. Du, J. Dong, R. Che, F. Kong, W. Cheng, M. Ma, N. Gu and Y. Zhang,
26 Depletable peroxidase-like activity of Fe(3)O(4) nanozymes accompanied with
27 separate migration of electrons and iron ions, *Nat Commun*, 2022, 13, 5365.
28 10.1038/s41467-022-33098-y.
- 29 158. K. Fan, H. Wang, J. Xi, Q. Liu, X. Meng, D. Duan, L. Gao and X. Yan, Optimization
30 of Fe(3)O(4) nanozyme activity via single amino acid modification mimicking an
31 enzyme active site, *Chem Commun (Camb)*, 2016, 53, 424–427.
32 10.1039/c6cc08542c.
- 33 159. H. Ouyang, Q. Lu, W. Wang, Y. Song, X. Tu, C. Zhu, J. N. Smith, D. Du, Z. Fu and Y.
34 Lin, Dual-Readout Immunochromatographic Assay by Utilizing MnO₂ Nanoflowers



- 1 as the Unique Colorimetric/Chemiluminescent Probe, *Analytical Chemistry*, 2018,
2 90, 5147–5152. 10.1021/acs.analchem.7b05247.
- 3 160. X. Zhang, C. Xue, H. Cao, Y. Wu, B. Yang, T. Zhou, W. Zhai and J. Deng, Ultra-small
4 CuO(x)/GDYO nanozyme with boosting peroxidase-like activity via electrochemical
5 strategy: Toward applicable colorimetric detection of organophosphate pesticides,
6 *Talanta*, 2024, 279, 126639. 10.1016/j.talanta.2024.126639.
- 7 161. S. H. Wen, H. Zhang, S. Yu, J. Ma, J. J. Zhu and Y. Zhou, Nanozyme coating-gated
8 multifunctional COF composite based dual-ratio enhanced dual-mode sensor for
9 highly sensitive and reliable detection of organophosphorus pesticides in real
10 samples, *J Hazard Mater*, 2024, 480, 135791. 10.1016/j.jhazmat.2024.135791.
- 11 162. J. Wu, Q. Yang, Q. Li, H. Li and F. Li, Two-Dimensional MnO(2) Nanozyme-
12 Mediated Homogeneous Electrochemical Detection of Organophosphate
13 Pesticides without the Interference of H(2)O(2) and Color, *Anal Chem*, 2021, 93,
14 4084–4091. 10.1021/acs.analchem.0c05257.
- 15 163. Y. Wang, Y. Liu, Z. Liu, X. Wang, M. Yu, G. Gao, Y. Zhao and J. Hong, A
16 colorimetric-SERS sensor based on MXene nanosheets loaded with Fe₃O₄@Ag
17 nanoparticles is used for sensitive and reliable detection of thiram, *Microchimica*
18 *Acta*, 2026, 193, 270. 10.1007/s00604-026-07988-7.
- 19 164. L. Shang, S. Zhou, X. Zhang, J. Chen, S. Wen, D. Liu, M. Mu, B. Zhao and W. Song,
20 Multiplexed discrimination and ultrasensitive determination of pesticides using a
21 Cu-O-Mo nanozyme tri-channel array driven by machine learning, *Biosensors and*
22 *Bioelectronics*, 2026, 297, 118374. <https://doi.org/10.1016/j.bios.2026.118374>.
- 23 165. Y. Chen, H. Liu, Y. Tian, Y. Du, Y. Ma, S. Zeng, C. Gu, T. Jiang and J. Zhou, In Situ
24 Recyclable Surface-Enhanced Raman Scattering-Based Detection of
25 Multicomponent Pesticide Residues on Fruits and Vegetables by the Flower-like
26 MoS₂@Ag Hybrid Substrate, *ACS Applied Materials & Interfaces*, 2020, 12, 14386–
27 14399. 10.1021/acsami.9b22725.
- 28 166. H. Li, S. A. Haruna, W. Sheng, Q. Bei, W. Ahmad, M. Zareef, Q. Chen and Z. Ding,
29 SERS-activated platforms for chemical contaminants in food: Probes, encoding
30 methods, and detection, *TrAC Trends in Analytical Chemistry*, 2023, 169, 117365.
31 <https://doi.org/10.1016/j.trac.2023.117365>.
- 32 167. A. Azzouz, V. Kumar, L. Hejji and K.-H. Kim, Advancements in nanomaterial-based
33 aptasensors for the detection of emerging organic pollutants in environmental
34 and biological samples, *Biotechnology Advances*, 2023, 66, 108156.



- 1 <https://doi.org/10.1016/j.biotechadv.2023.108156>.
- 2 168. T. Wu, Y. Mao, T. Wang, L. Ma, J. Cao, B. Xie, J. Wei and P. Li, Evolving trends in
3 nanozyme-based SERS systems for food contaminant monitoring: A review, *Food*
4 *Chemistry*, 2025, 486, 144621. <https://doi.org/10.1016/j.foodchem.2025.144621>.
- 5 169. X. Zhang, D. Wu, X. Zhou, Y. Yu, J. Liu, N. Hu, H. Wang, G. Li and Y. Wu, Recent
6 progress in the construction of nanozyme-based biosensors and their applications
7 to food safety assay, *TrAC Trends in Analytical Chemistry*, 2019, 121, 115668.
8 <https://doi.org/10.1016/j.trac.2019.115668>.
- 9 170. V. Kumar, D. Bano, D. K. Singh, S. Mohan, V. K. Singh and S. H. Hasan, Size-
10 Dependent Synthesis of Gold Nanoparticles and Their Peroxidase-Like Activity for
11 the Colorimetric Detection of Glutathione from Human Blood Serum, *ACS*
12 *Sustainable Chemistry & Engineering*, 2018, 6, 7662–7675.
13 10.1021/acssuschemeng.8b00503.
- 14 171. M. A. Abdel-Lateef, Utilization of the peroxidase-like activity of silver nanoparticles
15 nanozyme on O-phenylenediamine/H₂O₂ system for fluorescence detection of
16 mercury (II) ions, *Scientific Reports*, 2022, 12, 6953. 10.1038/s41598-022-10779-8.
- 17 172. X. Wang, X. Tang, C. Ji, L. Wu and Y. Zhu, Advances and Future Trends in
18 Nanozyme-Based SERS Sensors for Food Safety, *Environmental and Biomedical*
19 *Applications*, *Int J Mol Sci*, 2025, 26, 10.3390/ijms26020709.
- 20 173. Y. Yuan, J. Mi, Y. Cao, X. Hao, Y. Wu and J. Shi, A SERS sensing platform
21 leveraging Fe-ZIF nanozyme activity inhibition for ultrasensitive glyphosate
22 detection, *Opto-Electronic Engineering*, 2025, 52, 250115–250111–250115–250110.
23 10.12086/oe.2025.250115.
- 24 174. C. Li, F. Yu, J. Yang, H. Bai, X. Ma and Z. Jiang, SERS- and absorbance-based
25 catalytic assay for determination of isocarbophos using aptamer-modified FeMOF
26 nanozyme and in situ generated silver nanoparticles, *Microchimica Acta*, 2022,
27 190, 4. 10.1007/s00604-022-05549-2.
- 28 175. Y. He, R. Zhang, S. Xie, X. He, X. Yang, Y. Liu and M. Wang, A triple-mode strategy
29 for ultrasensitive and accurate point-of-care detection of glyphosate based on the
30 carved bimetallic prussian blue nanozyme and gold polyhedra, *Journal of Food*
31 *Composition and Analysis*, 2025, 144, 107661.
32 <https://doi.org/10.1016/j.jfca.2025.107661>.
- 33 176. M. Yan, G. Chen, Y. She, J. Ma, S. Hong, Y. Shao, A. M. Abd El-Aty, M. Wang, S.
34 Wang and J. Wang, Sensitive and Simple Competitive Biomimetic Nanozyme-



- 1 Linked Immunosorbent Assay for Colorimetric and Surface-Enhanced Raman
2 Scattering Sensing of Triazophos, *Journal of Agricultural and Food Chemistry*,
3 2019, 67, 9658–9666. [10.1021/acs.jafc.9b03401](https://doi.org/10.1021/acs.jafc.9b03401).
- 4 177. Y. Zhou, X. Liu, Y. Xu, Y. Jin, D. Wang and Y. Li, Sea-urchin-like Au@Pt nanozyme-
5 mediated sensing platform on single gold nanowire: Fabrication, cascade catalytic
6 amplification and highly sensitive dual-mode detection of organophosphorus
7 pesticides, *Sensors and Actuators B: Chemical*, 2026, 459, 139881.
8 <https://doi.org/10.1016/j.snb.2026.139881>.
- 9 178. Z. Wang, N. Liu, Y. Fan and A. Wu, Nanozyme-based biosensors for food
10 contaminants detection: advances, challenges, and prospects, *Talanta*, 2025, 295,
11 128290. <https://doi.org/10.1016/j.talanta.2025.128290>.
- 12 179. Y. Zhuang, H. Yin, Y. Huang, F. Jiang, L. Li, Z. Wu, Y. Yang, X. Cao and W. Wei,
13 Catalytic hairpin assembly-powered nanozyme-SERS dual-function sensing system
14 for ultrasensitive detection of gastric precancerous lesions, *Biosensors and*
15 *Bioelectronics*, 2025, 283, 117536. <https://doi.org/10.1016/j.bios.2025.117536>.
- 16 180. X. H. Pham, B. Seong, S. Bock, E. Hahm, K. H. Huynh, Y. H. Kim, W. Kim, J. Kim, D.
17 E. Kim and B. H. Jun, Nonenzymatic Hydrogen Peroxide Detection Using Surface-
18 Enhanced Raman Scattering of Gold-Silver Core-Shell-Assembled Silica
19 Nanostructures, *Nanomaterials (Basel)*, 2021, 11, 10.3390/nano11102748.
- 20 181. J. Jin, W. Song, J. Wang, L. Li, Y. Tian, S. Zhu, Y. Zhang, S. Xu, B. Yang and B. Zhao,
21 A highly sensitive SERS platform based on small-sized Ag/GQDs nanozyme for
22 intracellular analysis, *Chemical Engineering Journal*, 2022, 430, 132687.
23 <https://doi.org/10.1016/j.cej.2021.132687>.
- 24 182. F. Zhao, W. Wu, M. Zhao, S. Ding, Y. Lin, Q. Hu and L. Yu, Enzyme-like
25 nanomaterials-integrated microfluidic technology for bioanalysis, *TrAC Trends in*
26 *Analytical Chemistry*, 2023, 158, 116833.
27 <https://doi.org/10.1016/j.trac.2022.116833>.
- 28 183. J. Qin, N. Guo, J. Yang and J. Wei, Recent advances in metal oxide nanozyme-
29 based optical biosensors for food safety assays, *Food Chemistry*, 2024, 447,
30 139019. <https://doi.org/10.1016/j.foodchem.2024.139019>.
- 31 184. H. Li, Q. Lin, R. Zou, M. Zhang, X. Liu, H. Ye, C. Sun and X. Yan, A DNA tweezer-
32 actuated nanozyme-enzyme hybrid nanoreactor for pesticide detection,
33 *Biosensors and Bioelectronics*, 2025, 271, 117064.
34 <https://doi.org/10.1016/j.bios.2024.117064>.



- 1 185. S. Li, C. Pang, X. Ma, Y. Zhang, Z. Xu, J. Li, M. Zhang and M. Wang, Microfluidic
2 paper-based chip for parathion-methyl detection based on a double catalytic
3 amplification strategy, *Mikrochim Acta*, 2021, 188, 438. 10.1007/s00604-021-
4 05084-6.
- 5 186. H. Pan, W. Zhang, M. Jin, F. Zhang, X. Chen, X. Meng, H. Shao, L. Song, Z. Zhang
6 and C. Wang, Portable SERS device for rapid detection of indoxacarb and
7 chlorfenapyr in vegetable juice, *npj Science of Food*, 2025, 9, 137.
8 10.1038/s41538-025-00513-9.
- 9 187. N. Tabrizi, M. Ghafori-Gorab, A. Kashtiaray, M. Karimi, H. Aghamirza Moghim
10 Aliabadi, S. Adibzadeh and A. Maleki, Application of nanozyme-based biosensors
11 in pesticide residue detection: a review, *Materials Advances*, 2025, 6, 9272–9295.
12 10.1039/D5MA00800J.
- 13 188. B. Yin, Z. Jiang, R. Muhammad, J. Liu and J. Wang, Nanozyme-Powered
14 Multimodal Sensing for Pesticide Detection, *Foods*, 2025, 14, 1957.
- 15 189. E. M. Hamed, M. M. Elsaady and S. F. Y. Li, Single-Atom Nanozymes in Analytical
16 Chemistry: Opportunities and Challenges, *Analytical Chemistry*, 2025, 97, 26313–
17 26325. 10.1021/acs.analchem.5c05006.
- 18 190. E. L. S. Wong, K. Q. Vuong and E. Chow, Nanozymes for Environmental Pollutant
19 Monitoring and Remediation, *Sensors (Basel)*, 2021, 21, 10.3390/s21020408.
- 20 191. K. Wan, B. Jiang, T. Tan, H. Wang and M. Liang, Surface-Mediated Production of
21 Complexed •OH Radicals and Fe \square O Species as a Mechanism for Iron Oxide
22 Peroxidase-Like Nanozymes, *Small*, 2022, 18, 2204372.
23 <https://doi.org/10.1002/smll.202204372>.
- 24 192. J. Li, W. Liu, X. Wu and X. Gao, Mechanism of pH-switchable peroxidase and
25 catalase-like activities of gold, silver, platinum and palladium, *Biomaterials*, 2015,
26 48, 37–44. <https://doi.org/10.1016/j.biomaterials.2015.01.012>.
- 27 193. J.-F. Masson, Consideration of Sample Matrix Effects and “Biological” Noise in
28 Optimizing the Limit of Detection of Biosensors, *ACS Sensors*, 2020, 5, 3290–3292.
29 10.1021/acssensors.0c02254.
- 30 194. X. Yu, W. Zhuang, W. Bai, Q. Li, H. Tan, L. Zhang, J. Zhang, Y. Yao, S. Li, H. Bai, J.
31 Hu, X. Sun and W. Hu, Nanozyme-based biosensors with different combinations
32 of sensing modes for clinical biomarker detection: Biosensing strategies, practical
33 applications, and future perspectives, *Coordination Chemistry Reviews*, 2025, 542,
34 216850. <https://doi.org/10.1016/j.ccr.2025.216850>.



- 1 195. S. Wu, Z. Yu, J. Zhao, Y. Wang, J. Nan, L. Xu, X. Li, L. Yang and S. Dong,
2 Fluorescent Nanozyme Based on Bimetallic Fe/Co-Doped Carbon Dots for Dual-
3 Mode Detection of Ascorbic Acid and Hydrogen Peroxide, *Analytical Chemistry*,
4 2025, 97, 15320–15328. 10.1021/acs.analchem.5c02107.
- 5 196. L. Xi, Y. Chen, X. Zhang, M. Liu, J. Li, D. Xiao, P. Dramou and H. He, Less
6 interference fluorescence analytical strategy: Bridging substance-triggered
7 ratiometric sensor with convenient preparation and application, *Talanta*, 2024,
8 275, 126102. <https://doi.org/10.1016/j.talanta.2024.126102>.
- 9 197. X. Liao, X. Luo, Y. Li, Y. Zhou, Q. Liang, K. Feng, M. B. Camarada and J. Xiong, An
10 antifouling electrochemical sensor based on multiwalled carbon nanotubes
11 functionalized black phosphorus for highly sensitive detection of carbendazim and
12 corresponding response mechanisms analyses, *Microchemical Journal*, 2023, 190,
13 10.1016/j.microc.2023.108671.
- 14 198. S. Wang, P. Qi, S. Di, J. Wang, S. Wu, X. Wang, Z. Wang, Q. Wang, X. Wang, C.
15 Zhao and Q. Li, Significant role of supercritical fluid chromatography - mass
16 spectrometry in improving the matrix effect and analytical efficiency during multi-
17 pesticides residue analysis of complex chrysanthemum samples, *Anal Chim Acta*,
18 2019, 1074, 108–116. 10.1016/j.aca.2019.04.063.
- 19 199. Z. Kramplová, A. Ferancová, T. Maliar and A. Purdešová, Tuneable properties of
20 boron-doped diamond working electrodes and their advantages for the detection
21 of pesticides, *Journal of Electroanalytical Chemistry*, 2023, 949,
22 10.1016/j.jelechem.2023.117846.
- 23 200. Y. Li, R. Han, X. Yu, L. Jiang and X. Luo, A low-fouling electrochemical biosensor
24 based on multifunction branched peptides with antifouling, antibacterial and
25 recognizing sequences for protein detection in saliva, *Sensors and Actuators B:
26 Chemical*, 2024, 405, 10.1016/j.snb.2024.135322.
- 27 201. Y. Li, X. Pu, Y. Ding, L. Yi, Y. Yang, Y. Gu and S. Wang, An antifouling
28 electrochemical sensor based on a U-shaped four-in-one peptide and poly(3,4-
29 ethylenedioxythiophene) for vancomycin detection in fresh goat milk, *Food Chem*,
30 2025, 463, 141056. 10.1016/j.foodchem.2024.141056.
- 31 202. N. Hui, X. Sun, S. Niu and X. Luo, PEGylated Polyaniline Nanofibers: Antifouling
32 and Conducting Biomaterial for Electrochemical DNA Sensing, *ACS Appl Mater
33 Interfaces*, 2017, 9, 2914–2923. 10.1021/acsami.6b11682.
- 34 203. E. Jarosinska, Z. Zambrowska and E. Witkowska Nery, Methods of Protection of



- 1 Electrochemical Sensors against Biofouling in Cell Culture Applications, ACS
2 Omega, 2024, 9, 4572–4580. 10.1021/acsomega.3c07660.
- 3 204. X. Ma, H. Zheng, L. Fang, H. Zhou, W. Shi and W. Zhao, Anchoring atomically
4 distributed Mn with N, F Co-doped ultrathin carbon matrixes as multifunctional
5 nanozymes to boost electrochemical sensing performance, Sensors and Actuators
6 B: Chemical, 2025, 426, 10.1016/j.snb.2024.137075.
- 7 205. P. Wan, G. Chen, J. Fan, W. Tan, X. Li, L. Chen and K. Li, Modulating ion migration
8 realizes both enhanced and long-term-stable nanozyme activity for efficient
9 microplastic degradation, Chem Sci, 2025, 16, 15955–15963. 10.1039/d5sc04247j.
- 10 206. W. Pervaiz, M. H. Afzal, N. Feng, X. Peng and Y. Chen, Machine learning-enhanced
11 electrochemical sensors for food safety: Applications and perspectives, Trends in
12 Food Science & Technology, 2025, 156, 10.1016/j.tifs.2025.104872.
- 13 207. X. Wu, S. Shi, J. Jiang, D. Lin, J. Song, Z. Wang and W. Huang, Bionic Olfactory
14 Neuron with In-Sensor Reservoir Computing for Intelligent Gas Recognition,
15 Advanced Materials, 2025, 37, 2419159. <https://doi.org/10.1002/adma.202419159>.
- 16 208. Y. Chen, Y. Wu, W. Xu, Y. Tang, Y. Cai, X. Yu, J. Li, Y. Qiu, L. Hu, W. Gu and C. Zhu,
17 Nanozyme-Based Microfluidic Chip System for pH-Regulated Pretreatment and
18 Sensitive Sensing, Anal Chem, 2024,
19 <https://doi.org/10.1021/acs.analchem.4c02415>.
- 20 209. Q. Liu, H. Wei and Y. Du, Microfluidic bioanalysis based on nanozymes, TrAC
21 Trends in Analytical Chemistry, 2023, 158, 10.1016/j.trac.2022.116858.
- 22 210. M. Sun, S. Huang, G. Su, X. Wang, Z. Lu, Y. Wang, T. Liu, Y. Jiang, C. Song and H.
23 Rao, Synthesis of pH-switchable Pt/Co₃O₄ nanoflowers: Catalytic mechanism,
24 four-enzyme activity and smartphone biosensing applications, Chemical
25 Engineering Journal, 2022, 437, 10.1016/j.cej.2021.134414.
- 26 211. H. Li, P. Merkl, J. Sommertune, T. Thersleff and G. A. Sotiriou, SERS Hotspot
27 Engineering by Aerosol Self-Assembly of Plasmonic Ag Nanoaggregates with
28 Tunable Interparticle Distance, Advanced Science, 2022, 9, 2201133.
29 <https://doi.org/10.1002/adv.202201133>.
- 30 212. M. Yang, D. Chen, J. Hu, X. Zheng, Z.-J. Lin and H. Zhu, The application of coffee-
31 ring effect in analytical chemistry, TrAC Trends in Analytical Chemistry, 2022, 157,
32 116752. <https://doi.org/10.1016/j.trac.2022.116752>.
- 33 213. T. Bhardwaj and T. K. Sharma, A perspective on the selection and design of
34 nanozyme-based aptasensors for small molecules, Biosensors and Bioelectronics:



- 1 X, 2024, 21, 100533. <https://doi.org/10.1016/j.biosx.2024.100533>.
- 2 214. Q. Zhao, H. Wang, W. Jiang, H. Gao, S. Wen, X. Feng, Q. Wu, C. He, Y. Zhu, L. Hu,
3 B. Zhao and W. Song, SERS Resolving of the Significance of Acetate on the
4 Enhanced Catalytic Activity of Nanozymes, *Analytical Chemistry*, 2022, 94, 17930–
5 17938. 10.1021/acs.analchem.2c03992.
- 6 215. G. Wang, N. Feng, S. Zhao, L. Song, Y. Zhang, J. Tong, Y. Liu, X. Kang, T. Hu, I.
7 Ahmad Khan, K. Lu, H. Wu and J. Xie, Synthesis and DFT calculation of microbe-
8 supported Pd nanocomposites with oxidase-like activity for sensitive detection of
9 nitrite, *Food Chemistry*, 2024, 434, 137422.
10 <https://doi.org/10.1016/j.foodchem.2023.137422>.
- 11 216. Y. Wang, Y. Zhao, Q. Tan, G. Xiao, J. Baeyens and H. Su, Bioinspired Bi-amino Acid
12 Ce-MOFs Boosting Oxidase-like Activity: Dual-mode Aflatoxin Detection and
13 Antimicrobial Activity Platform, *Chemical Engineering Journal*, 2025, 512,
14 10.1016/j.cej.2025.161977.
- 15 217. Z. Lin, G. Sun, H. Liu, X. Zhang, Z. Bian and A. Liu, Mechanism of pesticide thiram
16 reversibly inhibiting of Pt single-atom peroxidase-mimicking nanozyme and its
17 application in colorimetric sensing thiram, *Talanta*, 2025, 294, 128201.
18 <https://doi.org/10.1016/j.talanta.2025.128201>.
- 19



Data Availability Statement

No primary research results, software or code have been included and no new data were generated or analysed as part of this review.

