MICROBIAL BIOPESTICIDES IN INDIA

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ISBN: 978-93-95319-88-1



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Foreword

In India, the crop losses due to pests and diseases are estimated to the tune of 30-40% under field condition and 9-10% post-harvest. The estimates suggest that pathways for introduction of invasive biotic stress have increased post globalization and free trade policy leading to losses associated with invasive insects, plants, and pathogens worth up to \$ 1.4 trillion annually. These biotic stresses are compounded by climate change influences. During Green Revolution and thereafter, the frequent and high application of chemicals and pesticides in agriculture, approximately 500 insect and related arthropod species are reportedly developed resistance against major groups of chemical pesticides, besides pesticide and chemicals loads in agriculture produces and outbreaks and resurgence of secondary pest etc.

The policy reorientation with strict regulations and ban on some of the hazardous chemical pesticides in the recent past and consumers awareness and preference for healthy agricultural products have put responsible focus on 'Greener Technologies' under environment friendly strategic framework. The 'National Policy for Farmers' (NPF), 2007 emphasised for increasing the organic agriculture which pushed the promotion of microbial biopesticides in India. Although the present policy ecosystem for greener technologies are supportive to the biopesticides, the factors such as high initial cost in identification and development of organisms, lack of subsidies to the small scale manufacturers, complex registration protocols, interest of multinational companies in new pesticide and associated business etc., are some hurdles in rapid growth of biopesticide industries and large scale utilization.

It is heartening to learn that currently 970 microbial biopesticide formulations from 15 microbial species are registered in India and 31 new microbial biopesticides developed by various ICAR institute against crop pests and pathogens are at various stages of registration and commercialization. I congratulate the authors for bringing out a compilation on 'Microbial Biopesticides in India' which shall be a useful reference book for policy makers, researchers, students and all other holders.

Dated: 6th September, 2022 Place: New Delhi (Himanshu Pathak)

Preface

In Indian agriculture, the widespread application of chemical pesticides to protect crops from biotic stresses has become a regular practice, with several unintended negative impacts on crops, people, animals, soil, water bodies, non-target creatures, and their surrounding environments. By 2050, the world's population is expected to increase to 9 billion people, necessitating a review and revision of current programmes for a 70% increase in food production. The government's strict regulations on chemical pesticides have created a favourable environment and provided a path for alternate, environmentally acceptable methods of managing biotic stress. Microbial biopesticide has been identified as an emerging tactic that is rapidly expanding in the context of plant protection in India. The appropriate modifications and simplifications to registration standards of biopesticides made by the Government, increase of the amount of land used for organic farming, subsidies to investors in the biopesticide industry, and other changes that attracted the attention of manufacturers. Additionally, stakeholders are concentrating on the advantages of biopesticides by raising knowledge of the value of high-quality goods that promote a healthy way of life and likelihood. Research and development efforts to improve the kill rate, shelf life, etc., of biopesticides are an added benefit for the steady growth of industry both internationally and in India. A steady rise in demand and consumption of biopesticides in India is a good sign for their ability to compete with or even surpass the market for chemical pesticides between 2040 and 2050.

A book titled "Microbial Biopesticides in India" consisting of 11 chapters with a focus on the need for biopesticide in plant protection, formulations, nanobiopesticide, genetic engineering, demand, consumption, and market, as well as government initiatives, awareness by growers, driving strategies and setback to enhance the biopesticide market in India was written by referring latest literature, dailies, review papers etc., in order to bring the Indian perspectives on microbial biopesticide to the most common platform for the benefit of readers, learned faculties and colleagues, corporate, stakeholders, students, youngsters etc. We, the writers, have been involved in the compilation of peer-reviewed data on the growth and development of biopesticide in India over the past two decades. Comparisons between India's policies and those of other industrialized nations' regulatory frameworks for biopesticide science include a number of concrete steps that the Indian Government has previously taken or plans to take. The coordinated work output by all authors in carefully gathering and compiling a variety of information from numerous sources is appreciated.

Editors

Contents

For	reword	vii
Pre	face	ix
Intr	roduction	xv
1.	Why We Need Biopesticides: Some Case Studies of Chemical Pesticides Introduction Case Studies on Acute Toxicity of Chemicals Pesticides	1
	Toxicities of Chemical Pesticides as Potential Endocrine Disruptors. Toxicities of Chemical Pesticides on Animals Effect of Neonicotinoid Insecticides to Bees Instances of Environmental Toxicities of Chemical Pesticides Why Biopetsicides? Conclusions References	9 9 11 12 13 14
2.	Insecticide Resistance and Case Histories Introduction Insecticide and Insecticide Resistance Importance of Insecticide Resistance and its Monitoring Mechanisms of Insecticide Resistance History of Insecticide Resistance Case Histories of Insecticide Resistance San Jose Scale Resistant to Lime Sulphur Paradigm Shift to Biopesticides	20 22 21 22 23 26 26
3.	Way Forward Conclusions References History and Development of Biological Control	28 29 29 31
	Introduction	34 36

4.	Biopesticides Classification and Their Formulations in Indi	a39
	Introduction	
	Common Method of Mass-production of Biopesticides	
	Biopesticide Formulations	
	Biopesticide Mass-production Units in India	
	Biopesticide Registrant in India	
	Conclusions	
	References	58
5.	Nano-Biopesticides for Management of Insect Pests of Croj	os61
	Introduction	61
	Nano-Biopesticides	
	Botanical Based Nano-Biopesticides	
	Nano-formulations of Semiochemicals	
	Nano-Biopesticides Derived from Plants	
	Nanoparticles as Nano-Biopesticides	
	Ag Nanoparticles (AgNPs)	
	Silica Nanoparticles (SiNPs)	
	Conclusions	
	References	
6	Impact of Biopesticides Application on Crop Quality and Environmental Quality Introduction	
0.	and Environmental Quality	91
	Introduction	01
	Merits of Biopesticides	
	Constraints in Exporting Agricultural Commodities	
	Impact of Biopesticides on Crop Quality Impact of Biopesticides on Environmental Quality	
	Case Studies	
	Entomopathogenic Nematodes (EPN) for Crop and Soil Health	
	Conclusions	
	References	
7.	Indian Biopesticide: Market Consumption, Growth	
	and Opportunities	
	Introduction	
	Biopesticide Market: Global and Indian Perspectives	
	India Biopesticide Market	
	Global Biopesticide Market Growth	
	India Biopesticide Market Growth	
	Conclusions	
	References	107

8.	Biopesticides: Research and Development Activities	109
	Introduction	109
	Entomopathogenic Bacteria	
	Insect Viruses	110
	Entomopathogenic Fungi	111
	Entomopathogenic Nematodes	112
	Protozoans	112
	Resistance to Microbes	113
	Genetic Improvement of Insect Pathogens	114
	Entomopathogenic Bacteria	114
	Insect Baculoviruses	
	Entomopathogenic Fungi	
	Entomopathogenic Nematode	118
	Conclusions	118
	References	118
9	Bionesticide Commercialization: World-wide Regulation	
	Biopesticide Commercialization: World-wide Regulation Policies for Registration and Use of Biopesticides	
	Introduction	125
	Regulation Policies of Biopesticide Registration	126
	China	126
	India	127
	Africa	127
	South Africa	128
	European Union	128
	USSR (formerly)	128
	United Kingdom	129
	United Kingdom	129
	Biopesticide Registration Protocol in India	130
	Policies on Biopesticide Use	
	Limitations in Regulations of Biopesticide Registration	
	Interventions in Regulations of Biopesticide	134
	Conclusions	135
	References	136
10	Promotion of Biopesticides in India: Role of Government	
10.	and Growers	141
	Introduction	
	Government Initiatives	
	Consumer Awareness on Biopesticides	
	Conclusions	
	References	

11.	Growth of Biopesticides: Driving Force and Set-Back.	145
	Introduction	
	Supporting Points for Biopesticide Growth	147
	Factors Restrain Biopesticide Market	149
	Conclusions	151
	References	



Introduction

The "Green Revolution" (GR) was brought about by the use of numerous inorganic outputs, such as pesticides, fertilisers, high input responsive cultivars, etc., in Indian agriculture, which greatly increased crop production and productivity. The stakeholders were motivated by the proportionate rise in yield indices to use numerous inorganic inputs carelessly, which had negative effects on soil quality, crop output quality, environmental pollution, human and animal health, etc. Inappropriately using synthetic chemical pesticides on a large scale to protect crops has led to a number of threats, such as insecticide residues in crop products, outbreaks, resurgence, the creation of secondary pests, insecticide resistance, and more. More than 500 insect and related arthropod species have been found to become resistant to significant classes of chemical insecticides.

The use of modern agricultural inputs needs to be reconsidered because the world's population is projected to increase to 9 billion people by 2050, which will result in an additional demand for food of about 70%. Despite this, global agriculture is still in the process of recovering from a number of negative effects. Wherever human action is used to correct the situation and return it to normal, climate change has been reported to amplify the negative effects. The current government's strict controls and rules on the registration, production, marketing, and subsequent field usage of chemical pesticides serve to lower demand and consumption. By raising knowledge among farmers, customers, the general public, etc. about the usage of high-quality goods free of pesticide residues, adulterations, etc., these groups' views were altered, and they were more interested in alternative tactics like "Green Technologies" for plant protection.

Many Government initiatives and programmes which include Sikkim Organic Mission (SOM), National Programme for Organic Production (NPOP), Organic Farming Policy (OFP), Strengthening and Modernizing Pest Management Approach in India (SMPMA), Capital Investment Subsidy Programme (CISP), National Action Plan on Climate Change (NAPCC), National Mission for Sustainable Agriculture (NMSA), "*Paramparagat Krishi Vikas Yojana*" (PKVY), Soil Health Management (SHM), Zero Budget Natural Farming (ZBNF) etc., are supportive for the scope of using 'Microbial Biopesticide' in India. Many small-scale industries were drawn to the biopesticide industry by the continued easing of regulation policies, but the initial investment costs for microbe identification, characterization, bioefficacy tests, toxicology tests, registration, commercialization, and other related costs are prohibitive. Based on their prior

experiences in international trade, multinational corporations (MNCs) are interested in innovative pesticide chemistries. In order to increase the demand and market for biopesticides, it is recommended that the government should provide adequate subsidies for early investment expenditures, set a fair pricing for biopesticides, and work with multinational corporations to support small-scale companies.

The number of bio-production units has currently increased to 361, of which 141 are in the private sector without GOI grant aids and 38 with GOI grant aids. Moreover, the Ministry of Agriculture and Farmers Welfare has assisted about 35 IPM centers to produce biopesticides since 2010. A total of 98 State Biocontrol Laboratories were established by the State Departments of Agriculture and Horticulture of Gujarat, Uttar Pradesh, Karnataka, Tamil Nadu, Andhra Pradesh and Kerala as well as the production of microbial pesticides by the Institutions of the Indian Council of Agricultural Research.

A total of 970 biopesticides registered in India by Central Insecticide Board and Registration Committee (CIB&RC) under the 1968 Insecticide Act which include microbial biopesticides of *Bacillus thuringiensis* var. kurstaki (42), var. israelensis (22), var. sphaericus (05), var. galleriae (01), *Pseudomonas fluorescence* (196), *Bacillus subtilis* (04), *Trichoderma viride* (289), *T. harzianum* (51), *Ampyliomyces quisqualis* (02), *Beauveria bassiana* (106), *Metarhizium anisopliae* (30), *Verticillium lecani* (93), *Verticillium chlamydosporium* (03), *Helicoverpa armigera* NPV (30) and *Spodoptera litura* NPV (03) and only 38 biopesticidal formulations. Fungal based- (Trichderma sp.) and Pseudomonas based- biopesticides are popular in India consumption-wise while *Bacillus thuringiensis* based formulations are widely used for plant protection of abroad agriculture.

In India, public sectors contribute 70% of the biopesticides production. Major companies are Biotech International Ltd., New Delhi, International Panaacea Ltd, New Delhi, Ajay Biotech (India) Ltd, Pune, Bharat Biocon Pvt. Ltd., Chhattisgarh, Microplex Biotech and Agrochem Pvt., Mumbai, Excel Crop Care Ltd., Mumbai, Govinda Agro Tech Ltd., Nagpur, Jai Biotech Industries, Satpur, Nasik, Ganesh Biocontrol System, Rajkot, Gujarat Chemicals and Fertilizers Trading Company, Baroda, Gujarat Eco Microbial Technologies Pvt. Ltd., Vadodara, Chaitra Agri-Organics, Mysore, Deep Farm Inputs (P) Ltd., Thiruvanandapuram, Kerala, Kan Biosys Pvt. Ltd., Pune, Indore Biotech Inputs and Research Pvt. Ltd., Indore, Romvijay Biotech Pvt. Ltd., Pondichery, Devi Biotech (P) Ltd., Madurai, Tamil Nadu, T. Stanes and Company Ltd., Coimbatore, Tamil Nadu, Harit Bio Control Lab., Yavatmal and Hindustan Bioenergy Ltd., Lucknow. Few Indian companies which work in biopesticde production in collaboration with foreign companies are Lupin Agro-chemicals, Mumbai, Sugar and distillery companies such as KCP Sugar and Industries Corporation Ltd., Andhra Pradesh, Rajshree Sugars and Chemicals Ltd., Tamil Nadu, New Swadeshi Sugar Mills, Bihar, and Bannari Amman Sugars Ltd., Tamil Nadu.

In India, the usage of biopesticides is growing at a faster pace than that of the chemical pesticides. According to the Directorate of Plant Protection, Quarantine and Storage, Ministry of Agriculture and Farmer Welfare, in the last 10 years, consumption of bio-pesticides increased by 23%, while that of chemical pesticides grew only by 2%. At 2020, the compound annual growth rate (CAGR) of global biopesticide market was approximately 3-5% of the total crop protection market while the market was anticipated to grow by 8.64 % at 2023; 9.7% at 2015-2023; 10.3% at 2014-2022; 15% at 2019-2024; 16% at 2020-2025.

Even though the biopesticide sector in India is growing rapidly, issues including slow kill times and short shelf lives are slowing it down. Around the world, researchers are creating recombinant organisms that contain spider, scorpion, and other venoms, adding and deleting genes of interest, and developing nanobiopesticides that have an efficacy that is comparable to chemical insecticides. For the control of important crop pests, a total of 31 fungal and bacterial based biopesticide formulations are under development and at various phases of commercialization. Between 2040 and 2050, the market for biopesticides is anticipated to surpass that for chemical pesticides, becoming one of the key elements of IPM all over the world.



1

Why We Need Biopesticides Some Case Studies of Chemical Pesticides

Abstract

Pesticides are used in most countries around the world to protect agricultural and horticultural crops against damage by pests and diseases. Injudicious use and unintentional poisoning of synthetic pesticides resulted deadly consequences. Exposure to chemical pesticides can have effects that are acute, chronic and long-term. Unregulated misuse of chemical pesticides lead to mobilization of toxic residues across the food chain, increasing bioaccumulation and environmental persistence. Non-target organisms, beneficial insects, land and aquatic animals are badly affected with the excessive use of chemical pesticides. Additionally, chemical pesticide poisoning poses a global concern due to unnatural death caused by mishandling of chemical pesticides. Biopesticide is one of the promising alternatives which can manage menace caused by pests in agriculture, persistency of pesticides, environmental pollutions, toxic and ill effects on nontarget species. The development of biopesticides stimulates modernization of agriculture and will, without a doubt, gradually replace chemical pesticides to a great extent.

Keywords: Biopesticides, Residue, Persistency, Environmental pollutions, Glyphosate

Introduction

The world population is expected to reach 9 billion by 2050. This population growth of 2 to 3 billion people over the next 30 years, combined with the changing diets, would result in a predicted increase in food demand of around 70% by 2050 (UNDESA 2009). To feed the burgeoning population, more food and livelihood opportunities from less per capita arable land and water

Aditi Kundu, Supradip Saha, Anirban Dutta, and Abhishek Mandal Division of Agricultural Chemicals, ICAR-IARI, New Delhi

required. Damage caused by insect and pest is one of the major limiting factors for agricultural food grain production. A major portion of expenditure on pesticides is for protecting the crop in the field (Kumar 2013).

Since the discovery of DDT, numerous pesticides (organochlorines, organophosphates, carbamates, pyrethroids, neonicotinoids, etc.) have been developed and used extensively worldwide with few guidelines or restrictions. Indeed, they help control agricultural pests (including diseases and weeds), plant disease vectors, human and livestock disease vectors and nuisance organisms, and organisms that harm other human activities and structures (gardens, recreational areas, etc.). However, many pesticides have been found to be harmful to the environment and human health. Some of them can persist in soils and aquatic sediments, bio-concentrate in the tissues of invertebrates and vertebrates, move up trophic chains, and affect top predators. They have caused adverse effects on soil health, water quality, produce quality and developed problems like insect resistance, genetic variation in plants, toxic residues food and feed. Moreover dependence on chemical pesticides and their indiscriminate use caused several detrimental effects on ecosystems.

Additionally, poisoning by agricultural pesticides is currently an important cause of human morbidity and mortality worldwide, increasing number of farm workers annually exposed to pesticides in developing countries (Jeyaratnam 1990). Developing countries use only 20% of the world's agrochemicals, yet they suffer 99% of deaths from pesticide poisoning (Jeyaratnam and Chia 1994). It has estimated that some form of poison directly or indirectly is responsible for more than one million illnesses worldwide annually. Acute pesticide toxicity is extremely common in developing countries of the Asia-Pacific region, particularly in settings of low education and poor regulatory frameworks. For, deliberate self-poisoning, a plausible range 233,997 to 325,907 with the estimated number of 258,234 probable deaths occurfrom pesticide self-poisoning worldwide each year (Gunnell et al. 2007). Fatality rates of 20% are common and the World Health Organization has estimated that more than 200,000 people die each from pesticide poisoning only (Singh and Unnikrishnan 2006).

Owing to huge adverse environmental impacts of synthetic chemicals, leading to resistance and resurgence of pests, forced to search alternate option for pest management. Further, the increasing public concerns and growing awareness about the potential adverse environmental effects as well as health hazards associated with the use of synthetic plant protection chemicals has prompted search for the technologies and products which are safer for the end users and the environment. Concerns of resistance development in pests and withdrawal of some of the products for either regulatory or commercial reasons, triggered to exploit naturally occurring pesticides. Biopesticides are environment friendly and safer than classical chemical pesticides. Hence, in the recent years, considerable attention has been paid towards exploitation of biopesticides in protection of food crops/commodities from pest infestations and the associated losses. They are more inclined to use eco-benign natural or herbal products in anticipation of any undesired side effects.Natural occurring phytochemicals have been an excellent option to replace toxic chemical pesticides. It has been speculated that botanical pesticides could reduce the pest resistance problem, thereby often subdue deleterious effects of hazardous chemicals. India has great diversity of flora and fauna. Treasure of bioactive phytochemicals from the diverse plant kingdom need to be exploited to develop newer bioactive molecules. Recent report published by WHO showed more than 21,000 plant species worldwide have tremendous potential for being used in medicinal and phytochemistry. It is estimated that more than 30% of the entire phyto-population possessed active constituents with complex biofunctional characteristics. Bioactive compounds derived from plants have proven to be valuable sources of bioactive secondary metabolites which can seldom be obtained from other sources (Kharshing 2012).

Case Studies on Acute Toxicity of Chemicals Pesticides

India is an agricultural country with a large rural population (60-80%), where pesticides are easily available and used extensively. Among different pesticides, organophosphates are most commonly used for self-poisoning, but being highly toxic, new compounds with high potency and lower toxicity are being developed continuously. The Poison Information Centre of the National Institute of Occupational Health, in Ahmedabad, reported that organophosphorus (OP) pesticides were responsible for the maximum number of poisonings (73%) among all agricultural chemicals (Dewan and Sayed 1998). Later another study was reported on season-long assessment of acute pesticide poisoning among cotton growing farmers across three villages in India. The study documented the serious consequences of pesticide use for the health of farmers, particularly women field helpers who were involved in mixing concentrated chemicals and refilling spraying tanks were as hazardous as direct pesticide application. Of 323 reported events, 83.6% were associated with signs and symptoms of mild to severe poisoning typical of poisoning by organophosphates (Mancini et al. 2005).

Acute intoxications after ingesting glyphosate was reported in the last four decades. Despite low potential toxicity of this herbicide, a number of fatalities and severe outcomes have been reported. Deaths following ingestion of 'Roundup' alone were due to a syndrome that involved hypotension, unresponsive to intravenous fluids or vasopressor drugs, and sometimes

pulmonary oedema, in the presence of normal central venous pressure (Talbot et al. 1991). Incidence on acute poisoning in case of suicidal or accidental cases after ingesting glyphosate and its lethal concentration in clinical samples have been published in literature (Hori et al. 2003). Glyphosate-poisoning is characterized by various symptoms such as gastrointestinal symptoms, altered consciousness, hypotension, respiratory distress, metabolic acidosis and renal failure (Tominack et al. 1991; Lee et al. 2000; Roberts et al. 2010). Glyphosate formulation contains surfactants that probably enhance its toxicity. The mortality rate due to glyphosate poisoning is reported at 3.2% in a study included 601 patients with glyphosate acute poisonings. Death was strongly associated with greater age, larger ingestions and high plasma glyphosate concentration $>734 \mu gmL^{-1}$. The most common symptoms were oropharyngeal ulceration, nausea and vomiting, mainly due to altered biological parameters of high lactate and acidosis. Respiratory distress, cardiac arrhythmia, hyperkaleamia, impaired renal function, hepatic toxicity and altered consciousness were the marked observations (Gress et al. 2015; Peillex and Pelletier 2020). Fatalities caused due to cardiovascular shock, cardio-respiratory arrest, haemodynamic disturbance, intravascular disseminated coagulation and multiple organ failure (Zouaoui et al. 2013).

Another case report in Thailand, where poisoning from glyphosate-surfactant herbicide has been displayed with rapid lethal intoxication. For a woman who ingested approximately 500 mL of concentrated Roundup formulation (41% glyphosate as the isopropylamine salt and 15% polyoxyethylene amine) showed glyphosate levels of 3.05 and 59.72 mg/mL in serum and gastric, respectively (Sribanditmongkol et al. 2012). During the re-approval process of glyphosate in Europe, it was mentioned that glyphosate-based products (GBF) were more toxic than glyphosate alone. This phenomenon was attributed to the surfactants and among them, polyethoxylatedtallowamine (POEA) has been suspected to significantly contribute to the toxicity of glyphosate products. In animal data acute oral toxicity of POEA has been suggested to be greater than glyphosate toxicity (Langrand et al. 2020).

Several episodes of mass poisoning by different pesticides have been reported to the Poison Information Centre (PIC) of the National Institute of Occupational Health (NIOH) in Ahmedabad, India, most notably endosulfan, phorate and ethion poisonings. It has been observed that OP poisoning from contaminated food ingestion is all too often treated empirically for food poisoning instead of specific treatment (Patel et al. 2012). A fatal accidental monocrotophos poisoning in adult female by dermal exposure while sleeping has been reported and elevated level of pesticide detected in post-mortem blood and skin by chromatography and spectroscopic techniques (Bodwal et al. 2019).

Endosulfan was one of the highly used organochlorine pesticides, and many poisoning cases have been reported from various regions of the world. In a case study, eighteen incidences of accidental endosulfan poisoning have been reported only from northern India between 1995 and 1997, which occurred after spraying of the pesticide. Analysis of various incriminating factors revealed that accidental overexposure was due to failure to adhere to the instructions for spray either due to ignorance or due to illiteracy (Chugh et al. 1998). Another case study from India revealed poisoning of endosulfan through consumption of endosulfan contaminated water by the entire age group (Srivastava et al. 2009). A survey for 11-year was carried out in various major cities including Ankara in Turkey, insecticides were found to be the most common cause (94%) of fatal pesticide poisoning, with organophosphates such as dichlorvos (25.7%) and organochlorines such as endosulfan (15.7%) being the most common types of pesticides involved (Kır et al. 2013). In Tehran, Iran, another case study showed high level of endosulfan poisoning and the most common culprit was organochlorines (57.1%) insecticide (Akhgari et al. 2018).

Another brief case study reported the inspection from January 2000 to December 2002, revealed 30 positive cases in 2000; 240 positive cases in 2001 and 38 positive cases in 2002. Organophosphorus insecticides were detected as the major component of most samples, representing 63% of the total positive cases and quinalphos is the most abundant pesticide, present in 32 of the 111 positive cases, followed by the herbicide paraquat (Teixeira et al. 2004). Unfortunately, poisoning and fatalities due to endosulfan, a halogenated carbohydrate derivative, have been widely reported in the Indian sub-continent. A fresh 23 cases of endosulfan poisoning have been reported describing symptoms were nausea and vomiting in 17 patients (73.9%), seizures in five patients (21.7%), and dizziness in one patient (4.3%) (Karatas et al. 2006). Two cases of unintentional exposure to endosulfan, one of which presented with neurological manifestations, liver toxicity, and required mechanical ventilation and emergent hemodialysis; the other had only neurological manifestations and liver toxicity, has been reported from a nine-year analysis study in Turkey (Yavuz et al. 2007).

Imidacloprid is a neonicotinoid insecticide developed for commercial use, belonging to the chloronicotinyl nitroguanidine chemical family. Imidacloprid has high potency against insects but with low mammalian toxicity and favorable persistence. On the basis of animal studies, it is classified as "moderately toxic" (class II by WHO and toxicity category II EPAV). Animal studies indicate relatively low toxicity to mammals because they have resistant nicotinic receptor subtypes compared to insects, as well as protection of the central nervous system by the blood brain barrier. Despite wide usage, human exposure experience resulting in toxicity is quite limited. Though imidacloprid

regarded as safe for human beings, toxicity can occur through inhalation exposure (Kumar et al. 2013).

Neuropsychiatric symptoms in imidacloprid poisoning have been reported in one case with inhalational exposure mainly due to central nicotinic stimulation (Huang et al. 2006). Cardiovascular manifestations like tachycardia, bradycardia, arrhythmia, and cardiac arrest were also described in different case reports (Wu et al. 2001). There is a paucity of information about human toxicity. Some reports also suggested that the other ingredients in the formulated product could be responsible for causing toxicity. In the same line of study imidacloprid formulation containing 9.7% active ingredient, <2 % surfactant, and the co-solvent, N-methyl pyrrolidone caused drowsiness, disorientation, dizziness, oral and gastroesophageal erosions, hemorrhagic gastritis, productive cough, fever, leukocytosis, and hyperglycemia. In fact, moderate to high dose imidacloprid in animals causes central nervous system activation similar to nicotine, including tremors, impaired pupillary function, and hypothermia, however, the causal role for the toxicity is still unclear (Wu et al. 2001). Similar observation was mentioned that moderate to relatively highdose imidacloprid in animals causes central nervous system activation similar to nicotine, including tremors, impaired papillary function, and hypothermia, it is more likely that the formulation ingredients caused most of the clinical symptoms including central nervous system depression and gastrointestinal irritation (Shadnia and Moghaddam 2008).

Two cases of acute poisoning with an insecticide formulation containing acetamiprid has been reported, exposed patients experienced severe nausea and vomiting, muscle weakness, hypothermia, convulsions, and clinical manifestations including tachycardia, hypotension, electrocardiogram changes, hypoxia, with the higher serum concentration of acetamiprid (Imamura et al. 2010). Similarly, another case study, Northeast China suggested toxic pesticides were responsible for comprehensive fatality (38.7%). Methomyl and fluoroacetamide were most commonly detected in the samples (Zhang et al. 2013).

In an autopsy case study of unnatural deaths in Northwest India, aluminium phosphide was found to be the most common suicidal poison, causing 68.4 % of total deaths due to poisoning between 1992 and 2000 (Chopra et al. 1986). Another case study revealed poisoning of 208 cases of death due to fatal poisoning of aluminium phosphide during the span of one year, January 2007 to December 2007. Studies decoded the mechanism of action of the fumigant in different animals described non-competitive binding of cytochrome oxidase by phosphine leading to valence change in the heme component of haemoglobin. However, other school of thoughts suggested inhibition of catalase, resulting to accumulation of hydrogen peroxide (Bogle et al. 2006).

Iable I: Case stud.	Image Image Image Image Voc.	Cal toxicolog	y or pesuci	alite. from	C. matoma	Doction do contract	Dafamana
1 Cal	resulting	country					Veletelle
2002-2007	Glyphosate-	Sri Lanka	601	27% were	Gastrointestinal	high plasma	Roberts et al.
	containing		patients	asymptomatic, 63.7%	symptoms,	glyphosate	2020
	herbicides			had minor poisoning,	respiratory distress,	concentrations on	
				and 5.5% had moderate	hypotension,	admission (>734	
				to severe poisoning. 19	altered level of	µg/mL)	
				deaths with a median	consciousness, and		
				time to death of 20 h)	oliguria		
2002 -2009	Glyphosate-	France	13 cases	1	Oropharyngeal	Blood glyphosate	Zouaoui et al.
	containing	<			ulceration, nausea	concentrations	2013
	herbicides				and vomiting,	had a mean	
					respiratory distress,	value of 61 mg/L	
		2			cardiac arrhythmia,	(range 0.6–150	
					hyper-kaleamia,	mg/L) and 4146	
		y			impaired renal	mg/L (range	
					function, hepatic	690–7480 mg/L)	
					toxicity and altered		
					consciousness		
2001 - 2007	Endosulfan	China	52 cases	30.7% of the 52 patients	Hypotension,	Amount ingested	Moon and
	poisoning			died, and 48 patients	rhabdomyolysis,	could be greater	Chun 2009
				experienced seizures.	hepatic toxicity, and	than 35 g, caused	
					hypotension.	mortality	
January to	Endosulfan	Singapore 23 cases	23 cases	Nausea and vomiting	Nausea, vomiting,	I	Karatas et al.
December 2005	poisoning			in 17 patients (73.9 %).	diarrhoea, stomach		(2006)
				seizures 21.7 %, and	ache, respiratory		
				dizziness 4.3 %	distress, seizures,		
					pulmonary oedema,		
					agitation, headaches,		
					dizziness etc.		

Year	Pesticide	Reported Patients		Fatality from acute	acute Symptoms	Pesticide content Reference	Reference
		country		poisoning			
January 2001 to	е	India	325	15% patients died			Singh and
May 2003	pesticides (65%)		patients				Unnikrishnan
9	and aluminium phosphide (15%)						2006
⁷ 2013 to June	July 2013 to June Organophosphate	India	133	36.1% were stable after	13.5% Patients		Banday et al.
2014	pesticides		patients	gastric lavage	showed episodic		2015
					convulsions,		
					derangement in renal function		
1997-2002	Monocrotophos	India	8040	Fatality 22.6%, 1819			Srinivas Rao
	and endosulfan		patients	patient died			et al. 2005
2019	Monocrotophos	India	1 patient Dead	Dead	Vomiting with		Sharma and
					breathlessness		Kumar, 2019
July, 2008	Organophosphate	China	335		Cerebral edema,	1	Zhang et al.
-December 2015	pesticides		cases		liver injury,		(2017)
					kidney injury,		
					myocardial injury,		
				(gastrointestinal		
					hemorrhage and		
					acute pancreatitis		
January 1985-	r	India	Sixteen		Profuse vomiting,		Chopra et al.
December 1985	phosphide		patients		pain in the upper		1986
				くとく	abdomen and shock		
					1		

Toxicities of Chemical Pesticides as Potential Endocrine Disruptors

Many chemicals that have been identified as potential endocrine disruptors, include pesticides. About 105 substances have been identified in this respect, of these, 46% are insecticides, 21% herbicides and 31% fungicides. Case studies indicated that thyroid hormone production can be inhibited by some ten pesticides (amitrole, cyhalothrin, fipronil, ioxynil, maneb, mancozeb, pentachloronitro-benzene, prodiamine, pyrimethanil, thiazopyr, ziram, zineb (Sugiyama et al. 2005; Leghait et al. 2009). Besides, effects linked to endocrine disruption have been largely noticed in invertebrates (Gooding et al. 2003), reptiles (Crain et al. 1997), fish (Purdom et al. 1994), birds (Vos et al. 2000) and mammals (Oskam et al. 2003). A case study on *Daphnia magna* has shown that endosulfan sulphate disrupts the ecdysteroidal system and juvenile hormone activity of crustaceans (Palma et al. 2009). Influence of linuron on reproductive hormone production has been reported in rats (Wilsor et al. 2009).

Epidemiological studies concluded that pesticide exposure affect spermatogenesis leading to poor semen quality and reduced male fertility, an increasing number of epidemiological studies linked environmental exposure to pesticides and hormone-dependent cancer risks. A case report on fat samples from women with breast cancer revealed elevated concentrations of PCBs, DDE, and DDT (Falck et al. 1992). Another epidemiological case studies performed in Spain between 1999 and 2009 shows that among 2,661 cases of breast cancer patients, 2,173 (81%) were associated with pesticide contamination (Parron et al. 2010).

Toxicities of Chemical Pesticides on Animals

Case report described the spraying of coca (*Erythroxylum coca*) with glyphosate (coca mixture, a combination of formulated glyphosate, Glyphos, and an adjuvant, Cosmo-Flux) in Colombia has raised concerns about possible impacts on amphibians. Mortality at 96 h in the control microcosms was between 0 and 16% and LC₅₀ values were between 8.9 and 10.9 kg glyphosate a.i./ha. Mortality >LC₅₀ was only observed in the tested species when the application rate was >2-fold the normal application rate (Bernal et al. 2009). Contrastingly, another report suggested no significant acute toxicity of the glyphosate end-use formulation Roundup Original[®] to four North American amphibian species (*Rana clamitans, R. pipiens, R. sylvatica,* and *Bufo americanus*) and the toxicity of glyphosate technical, the polyethoxylatedtallowamine surfactant (POEA) commonly used in glyphosate-based herbicides, and five newer glyphosate formulations to *R. clamitans*. For *R. clamitans*, acute toxicity

values in order of decreasing toxicity were POEA > Roundup Original > Roundup Transorb[®]>Glyfos AU[®]. However, relevant concentrations of POEA or glyphosate formulations containing POEA showed decreased snout-vent length at metamorphosis and increased time to metamorphosis, tail damage, and gonadal abnormalities. These effects may be caused, in some part, by disruption of hormone signaling, because thyroid hormone receptor β -m-RNA transcript levels were elevated (Howe et al. 2004).

The major mechanism of toxicity of OPs is the inhibition of Acetyl cholinesterase (AChE), resulting in a net accumulation of Acetylcholine (ACh) and increased stimulation of cholinergic receptors. In mammals, excessive stimulation of these cholinergic receptors in the central and peripheral nervous systems results muscarinic-receptor induced effects (excessive secretions, miosis, bradycardia) and nicotinic-receptor-induced effects (muscle tremors, convulsions, complete muscle paralysis). A case study in California, USA showed toxicity of phorate in a group of 300 Holstein cattle, a large number of cattle developed tremors, diarrhea, weakness, and paralysis. A total of 159, died within 24 h (Puschner et al. 2013).

An interesting study was carried out to assess the levels of atrazine, dimethoate, and dichlorodiphenyltrichloroethane on freshwater fish in Chiredzi, Zimbabwe revealed higher concentration of dichlorodiphenyltrichloroethane in water and fish muscle tissue at respective concentrations of 131.3 µg/l and 171.7 µg/kg, while concentrations of atrazine was 6.15 µg/l and 142.0 µg/kg in water and fish muscle tissue, respectively. The atrazine and DDT concentrations in water samples were above the limits permissible by the World Health Organization in drinking water. The pesticide in water were about three times higher than those in fish samples while significantly higher (p < 0.05) concentrations of atrazine (23-fold) were observed in fish samples compared to water. Levels of DDT and its metabolites in fish tissues were also higher than those in water samples (Basopo et al. 2020).

Another case study revealed the effect of glyphosate-based herbicide on aquatic organisms. Glyphosate has been widely used against terrestrial weeds, causes toxicity in plants include decreases in concentration of the aromatic amino acids, tryptophan, phenylalanine, and tyrosine, as well as decreased synthesis of protein, indole acetic acid and chlorophyll. Apart from their use in terrestrial environments, glyphosate-based formulas are also, however rarely, applied in order to control aquatic weeds, particularly invasive species e.g. common reed (*Phragmites australis*). Glyphosate was detected using gas chromatography mass spectrometry (GC-MS) in the water samples collected from the bathing area at a mean concentration of 0.09 mg dm⁻³. Significantly

lower numbers of Chironomidae (by 41%), Oligochaeta (by 43%), Vivipariae (by 75%), Hirudinae (by 75%), *Asellus aquaticus* (by 77%), *Gamma ruspulex* (by 38%) and *Dreissena polymorpha* (by 42%) were found at the glyphosate-treated site. The ranges of glyphosate concentration in the tissues of sampled macro-invertebrates and *P. australis* organs were 7.3-10.2 μ g kg⁻¹ and 16.2-24.7 μ g kg⁻¹, respectively (Rzymski et al. 2013).

Another instance reported five dead or debilitated bald eagles (Haliaeetus leucocephalus) and a red-tailed hawk (Buteo jamaicensis) from British Columbia (BC), Canada tested positive for residues of the organophosphorus insecticide, phorate (Elliott et al. 1997). Other cases of pesticide poisoning of wild birds diagnosed at the National Veterinary Research and Quarantine Service, Korea, where forty-one mortality events (759 birds) of 87 incidents (2,464 birds) were found affected by poisoning of six organophosphates or carbamates pesticides. Phosphamidon was most frequently identified as the cause of poisoning, accounting for 23 mortality events, besides, other pesticides identified as poisons for birds were monocrotophus, fenthion, parathion, EPN, and diazinon, carbofuran (Kwon et al. 2004). Later, serious threat of carbofuran has been reported in western Kenya, where, uncounted dead birds, Quelea species found in cereal fields. On investigation, it was observed that a large proportion of individuals of their populations were exposed to Furadan (Odino 2011). Similar cases were reported from January 2014 to October 2020, which confirmed pesticide poisoning substances in 503 samples of wildlife and domestic animals in Portugal. Toxicology results from domestic species (dog, cat, sheep, cows, and horses), wildlife species (red foxes, birds of prey, lynx, and wild boar), molluscicides, carbamates, rodenticides, strychnine and organophosphates (Grilo et al. 2020).

Effect of Neonicotinoid Insecticides to Bees

Pollinating insects, such as the honey bee, are mainly exposed to chemicals when visiting melliferous plants. Neonicotinoid insecticides were recently implicated by beekeepers who reported that hives placed near cropped plants, originated from seeds dressed with insecticide, showed high levels of damage due to a progressive decrease in the hive populations, until the complete loss of the colonies (Maus et al. 2003). The risk that systemic neonicotinoid insecticides induce for honey bees started in France with the use of Gaucho (active ingredient: imidacloprid) on sunflower (Maxima and van der Sluijs 2007). Generally, neonicotinoids are more toxic via oral route than contact mode. The difference between the oral and contact toxicity may be due to the weak hydrophobicity of the neonicotinoidsyielding a low penetration through the insect cuticle. Three species of bees, *Apis mellifera*, *Megachile rotundata*

and *Nomiam elanderi*, were found susceptible to imidacloprid (24-h LD_{50} 0.04 µg/bee) (Stark et al. 1995). Similar results were obtained for Admire and Provado that are two commercial formulations of imidacloprid (Devillers et al. 2003). The behavioral effects of neonicotinoid insecticides were largely investigated showed that foragers when collecting nectar and pollen were exposed to low doses of neonicotinoid insecticides during their foraging trips, which induced behavioral effects and subsequently no homing return to hive (Blacquiere et al. 2012).

A case study on the toxicity of dinotefuran, a neonicotinoid insecticide has been reported in Wilsonville, Oregon, USA which killed a large number of bumble bees. It was estimated that approximately 45,830 to 107,470 bumble bees from 289 to 596 colonies were killed with the chemical pesticide. Chromatographic analysis revealed concentration of dinotefuran in the samples flower was 7.4 ppm, which exceed 737% of the LC₅₀ of beneficial pollinator, honey bee. The dead bumble bees were tested for dinotefuran concentration which showed 0.92 ppm, far more than the maximum LC₅₀ (0.884 ppm) of *A. melifera* (Hatfield et al. 2021). Another study reported in Romania, where exposure of honey bees with neonicotinoids was estimated. In total, a set of fifty samples was collected from fields, located in different areas of intensive agriculture were analysed for five neonicotinoids which revealed 48% of the total samples contained one or more detected or quantified neonicotinoid residues (Cauia et al. 2020).

Instances of Environmental Toxicities of Chemical Pesticides

Despite the presence of rules and regulations, it has been observed that pesticides are not used in an appropriate manner. Much of the portion of chemical pesticides goes to wastage during their use. Pesticides are used in various types of pests control, remain a big source of air, water and soil pollution, which may negatively affect human health and the living organisms in the environment. Environmental impacts of pesticide use were commonly estimated through variables such as pounds of active ingredient applied or expenditure on pesticides. The disadvantage is that both these measures assume environmental damage is directly correlated with the quantity of pesticide used, regardless of the specific chemical formulation. The increased availability of low-dosage alternatives lend credence to the argument that weight and volume measures are not adequate proxies for assessing pesticide risk. Cornell University's Environmental Risk Analysis Program has identified eight of the indicators widely used worldwide: Environmental Potential Risk Indicator for Pesticides (EPRIP), Environmental Yardstick for Pesticides (EYP), Survey of National Pesticide Risk Indicators (SYNOPS), System for Predicting the Environmental Impact of Pesticides (SyPEP), Pesticide

Environmental Risk Indicator (PERI), Environmental Impact Quotient (EIQ), Chemical Hazard Evaluation for Management Strategies (CHEMS1), and Multi-Attribute Toxicity Factor (MATF). The first four indicators are referred to as predicted environmental concentration (PEC) indicators, and the later four constitute ranking indicators (Sande et al. 2011).

Negative effects of pesticides on the environment and the farmers awareness in Saudi Arabia has been described by Al-Zaidi et al. (2011). Another case study on assessment of hazards from methyl bromide and the proposed alternative fumigants to workers, consumers, beneficial arthropods, birds, fish, and bees in Florida, USA suggested the highest relative risks category under field workers and beneficial arthropods and fish and consumers the least risks (Sande et al. 2011). Similar kind of study was also conducted in Nepal to evaluate vegetable growers' knowledge on pesticide safety and pest management practices. Unfortunately, most of the farmer (>90%) did not know much about the harmful effects of pesticide residues nor practiced proper pesticide disposal methods (Rijal et al. 2018). A case study conducted the health risk associated with chemical pesticide contaminants in the drinking water sources of Dalian in China, revealed relatively higher concentration of atrazine and acetochlorat ng L-1 levels. Additionally, atrazine, acetochlor, hexachlorobenzene, p,p'-DDE, and p,p'-DDD were detected in the sediment/soil samples at ng g-1 levels. However, hexachlorobenzene and arsenic were identified as the main contributors to human carcinogenic risks, which were calculated at the high level of 10⁻⁴ (Dong et al. 2020).

Why Biopetsicides?

Biopesticide is gaining interest because of its advantages associated with the environmental safety, target-specificity, efficacy, biodegradability and suitability in the integrated pest management (IPM) programs. Thus, biopesticide is one of the promising alternatives to manage environmental pollutions. Though potential application of biopesticides in environmental safety is well known, it has gained interest in view of the growing demands for organic food. Although use of agrochemicals is indispensable to meet the ever growing demands of food, feed and fodder, opportunities do exist in selected crops and niche areas where biopesticides can be used as a component of IPM. The interest in biopesticides is based on the advantages associated with the products which are (i) inherently less harmful and environmentally safe, (ii) target-specific, (iii) often effective in very small quantity, (iv) naturally and quickly decomposable and, (v) usable as a component of IPM.

Biopesticides are very effective in the agricultural pest control without causing serious harm to ecological chain or worsening environmental pollution. The research and development of practical applications in the field of biopesticides greatly mitigate environmental pollution caused by chemical pesticide residues and promotes sustainable development of agriculture. Since the advent of biopesticides, a large number of products have been released, several of which have already played dominant roles in the market. The development of biopesticides stimulates modernization of agriculture and will, without doubt, gradually replace chemical pesticides. Many biopesticides are ideal substitutes for their traditional chemical counterparts in pollution-free agricultural production. Research in production, formulation and delivery may greatly assist in commercialization of biopesticides. More research is needed towards integrating biological agents into production system, improving capability of developing countries to manufacture and use biopesticides. At the same time, it is also required to encourage public funded programmes, commercial investors and pesticide companies to take up biopesticide enterprises.

Conclusions

Impact of synthetic pesticides, due in particular to an excessive use (including environmental pollution and implications to human health) have led to modifications in agricultural practices and various national and international regulations limiting their use. Further limitations and/or bans often encourage to find alternative solutions that are safer and non-toxic to the environment and humans. Most of the countries have amended their policies to minimize the use of chemical pesticides and promote the use of biopesticides. Policy measures need to be strengthened in order to reduce excessive use of chemical pesticides and to promote the use of biopesticides. Better understanding of the mode of action of biopesticides, their effects and regulatory issues that arise in their adoption may help further to raise their profile among the public, policymakers and hence enable them to realize their contributions to sustainability. The interest in organic farming and pesticide residue free agricultural produce would certainly warrant increased adoption of biopesticides by the farmers. Increasing concerns over environmental and health safety across the world would certainly create awareness among the farmers, manufacturers, policy makers and consumers to accept safer biopesticides for suitable pest management options.

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Insecticide Resistance and Case Histories

Abstract

Insecticides play a crucial role in the management of insect pests in order to reduce yield losses caused to high value and cash crops not only in India but also in the world. In the recent past few newer insecticide molecules have come in place to overcome hazardous issues with chemical pesticides. However, their usage is very limited and such molecules are not available for high value cash crops in case of pest outbreaks, resurgence etc. Insecticide resistance has got a long history wherein many insect pests have developed significant level of resistance making insecticides ineffective. Insecticide resistance in India started with development of resistance in Singhara beetle against DDT. Various resistance mechanisms adopted by insects to combat toxic effects are metabolic, altered target site sensitivity (mutations), reduced penetration, behavioural resistance etc. Pink bollworm developed resistance to Bollgard II in the country caused huge economic losses to cotton crop. Even recent outbreak of whiteflies in cotton in Punjab has also devastated the entire crop due to development of resistance. Although insecticide resistance management strategies have come in place for few pests, the insecticide resistance in insects in major crops made the various stake holders to think seriously for alternative pest control strategies to safe guard the crops from ravaging insect pests. Green pesticides, biopesticides and pesticides/ insecticides of biological origin have got greater attention as consumers are health conscious and demanding for organic products. Hence, there is paradigm shift in pest management strategies from chemical control to microbial control in India as well as in the world.

Keywords: Pest, Resistance, Insecticide, Mechanism, Biopesticides

Introduction

Insecticides are one of the key control measures to combat the insect pests for sustainable agricultural production in the world. Synthetic insecticides have been only strategy to control the resurgent and resistant insect population in of high value crops not only in India but also in the world. The advent of synthetic insecticides in the mid-20th century made the control of insect and other arthropod pests easy and much more effective, and such chemicals remain essential in modern agriculture despite their environmental issues. By preventing crop losses, raising the quality of produce, and lowering the cost of farming, modern insecticides increased crop yields by as much as 50% in some regions of the world. More than half of our crops would be lost to pests and diseases if pesticides are not employed. Between 26 and 40% of the world's potential crop production is lost annually because of weeds, pests and diseases (OECD-FAO 2012). Without crop protection, these losses could easily double. Insecticides enable farmers to produce safe, quality foods at affordable prices with abundance of nutritious, all-year-round foods, which are necessary for human health. Fruits and vegetables, which provide essential nutrients, are more abundant and affordable. Grains, milk and proteins, which are vital to childhood development, are more widely available because of lower costs to produce food and animal feed. Production of major crops has more than tripled since 1960, which was mostly due to pesticides (FAOSTAT) as in case of rice which feeds almost half the people on our planet doubled in production while the amount of wheat has increased nearly 160%. Insecticides have also been important in improving the health of both humans and domestic animals; malaria, yellow fever, and typhus, among other infectious diseases, have been greatly reduced in many areas of the world through their use. Pesticides include insecticides that are mainly used in agriculture or in public health protection programs in order to protect plants from pests, weeds or diseases, and humans from vector-borne diseases, such as malaria, dengue fever, and schistosomiasis (Alewu and Nosiri 2011). Besides, insecticides are being extensively used in sports fields, building bottoms, lawn development, public urban green areas etc to prevent unwanted insect pests such as termites (Hoffman et al. 2000; Nicolopoulou-Stamati et al. 2016). Despite tremendous benefits and advantages offered by insecticides to mankind, continuous use of insecticides over a longer period of time resulted in development of resistance, resurgence, residue and environmental issues, health hazards etc. Insecticide resistance become one of the major concerns in agriculture, public health sector and other fields in India as well as in the world

Insecticide and Insecticide Resistance

Insecticide is an agent that destroys insects as well as other small pests (such as mites or nematodes). While WHO panel of experts defined insecticide resistance as 'the development of an ability of a strain of insects to tolerate doses of toxicants which would prove lethal to the majority of individuals in a normal population of the same species' (Guedes 2017). Resistance in insects is usually a complex phenomenon with more than one mechanism operating simultaneously within the same insect strain (Oppenoorth and Welling 1976). The resistant phenotype of an insect that survives a dose of insecticide that would normally have killed it, is relatively monitored with direct insecticide bioassays. Pest control subjects the population to Darwinian selection and survival of the fittest and it attempts to kill the tolerant individuals lead to ever increasing doses and eventually resistant pest populations. As a result, the most difficult problems raised due to insensitive biochemical target conferring cross resistance to one or more classes of compounds formerly effective at that site. Insecticides are being widely used to control insect pests across the world which leads to high selection pressure on target insect over a period of time. Insecticides provide very good control of insects initially but over a period of time insects develop resistance by various mechanisms such as morphological, behavioural, ecological, environmental biochemical, genetical, molecular adaptations.

Importance of Insecticide Resistance and its Monitoring

Since the 1950s insecticide resistance has come into prominence around the globe as a key factor impacting the use and efficacy of a wide range of existing and new compounds for the control of insect and mite crop pests as well as vectors of human diseases. Insecticide resistance is also an important driver in the search for new insecticides, especially those with new modes of action. Within the crop protection industry, insecticide resistance was recognized as a concern as early as the late 1950s to early 1960s. The early industry response most often involved simply finding and using a different insecticide. Frequently the replacement products were in the same class of chemistry since there were few distinct classes of insecticides available during time. However, in some instances recommendations from industry, scientists included specific resistance mitigating measures such as moderation of use, alternation (rotation) of insecticides from different classes, and incorporation of biological control measures.

Mechanisms of Insecticide Resistance

Insect resistance to insecticides has been found to be mediated by various mechanisms in four different ways (Krathi et al 2002; Ju et al 2021, Liu 2015, Auteri et al 2018) :

- a) Metabolic resistance, due to an increased detoxification caused by the overexpression or conformational changes of the enzymes involved in the chemical insecticide metabolism, sequestration, and excretion. Cytochrome P450-monooxygenases, glutathione S-transferases, and carboxy/cholinesterases, microsomal mono-oxygenases, phosphotriester hydrolases, DDT-dehydrochlorinases are the main enzymes involved in this process.
- b) Altered target site sensitivity/mutation, caused by a modification of the chemical insecticide site of action reducing or preventing insecticide binding at that site. Mutations in the voltage sensitive sodium channel (Vssc) gene are one of the most common causes of target-site resistance. Insensitive acetylcholinesterases, insensitive sodium channels, insensitive GABA (γ-amino butyric acid) receptor are few such examples.
- c) Reduced penetration, due to modifications in the insect cuticle or digestive tract linings that limit the chemical insecticides absorption. However, the mechanism remains poorly understood, and its importance in Aedes species is yet to be confirmed.
- d) Behavioural resistance, which includes modifications in insect behaviour that help to avoid the lethal effects of chemical insecticides. This is considered as a contributing factor that allows the insects to avoid the lethal dose of an insecticide.

Insects metabolize insecticides to non toxic or less toxic forms through a process called 'detoxification' and sometimes to more toxic intermediates, a process called 'activation'. Substances that are completely water soluble (polar), and those that are completely insoluble in either water or fats, are excreted unchanged. Most insecticides, which are water insoluble (apolar) or fat soluble (lipophilic), are metabolised to polar compounds through a primary enzymatic conversion, mediated through 1) Oxidases, 2) Hydrolases or 3) Glutathione-S-transferases, resulting in watersoluble products that are subsequently converted to water soluble conjugates through a secondary non synthetic reaction. These conjugates are finally excreted. Apolar substances are converted to less lipophilic or polar metabolites by two reactions (Phase I and Phase II) in insects and many other organisms.

Phase I reactions are mainly carried out by two major groups of enzymes, the oxidoreductases and hydrolases. The oxidoreductases comprise of the cytochrome P450 dependent superfamily of monooxygenases, which introduce oxygen into or remove electrons from their substrates. Carbonyl reductases, alcohol dehydrogenases and aldehyde dehydrogenases remove hydrogen from, or add to the target molecule. The hydrolases hydrolyse esters, amides, epoxides or glucuronides. Typically the Phase I reaction introduces a functional group in a series of steps in lipophilic molecules. Phase II reactions are mainly carried out by the transferases. Glutathione S-transferases conjugate the electrophilic substrates, while the acetyl transferases, sulfotransferases, acyl-CoA aminoacid N-methyl transferases and UDP-glucuronosyl transferases metabolise the nucleophilic substrates. Insecticide metabolism in insects has been found to be catalysed mainly by monooxygenases, hydrolases and gluthathione-S-transferases. Generally, in resistant insects, the enzymatic detoxification is believed to be so rapid that the toxic molecule does not reach its site of toxic action

History of Insecticide Resistance

In India, insecticide resistance has been well documented by Mehrotra (1989) where Singhara beetle, Galerucella birmanica (1963), Tobacco caterpillar, Spodoptera litura (1965), Diamondback moth, Plutella xylostella (1968), Gram pod borer, Heliothis armigera (1986), aphids and jassids, Empoasca kerri (1986), Lipaphis erysimi (1986), Aphis craccivora (1986) have developed resistance to DDT, HCH, organophosphates (malathion and dimethoate), endosulfan etc. Subsequently, Helicoverpa armigera (1987) developed resistant to synthetic pyrethroids in cotton ecosystem where very high proportion of insecticides have been used before the introduction of Bt cotton. Insecticide resistance has been reported mostly in cotton ecosystem during the decade 1990-2000 which was the most difficult for cotton pest management due to excessive use of insecticides, especially synthetic pyrethroids that led to problems of high levels of resistance to pyrethroids and DDT in Helicoverpa armigera and Spodoptera litura in cotton and pulse growing regions of the country (Sekhar et al. 1996). Subsequent studies (Armes et al. 1996; Kranthi et al. 2002) showed that resistance to pyrethroids was ubiquitous and resistance in *H. armigera* to conventional insecticides such as methomyl, endosulfan and quinalphos was increasing in India. Due to unsatisfactory insect control on account of insecticide resistance, farmers were forced to spray repeatedly, most often with mixtures.

The outbreaks of whitefly during 1988 and recently on cotton in Punjab (2015) which destroyed 2/3 of cotton in India was due to indiscriminate

use of pyrethroids, elimination of natural enemies, favourable temperatures (Sundaramurthy et al. 1992), presence of a wide range of hosts such as vegetables, pulses, throughout year helps the whiteflies to survive and proliferate (Kranthi 2015). Moreover, development of resistance in whiteflies to synthetic pyrethroids and insect growth regulators like pyriproxyfen was reported. Very recently, Naveen et al. (2017) Studied the level of insecticide resistance to selected organophosphates, pyrethroids, and neonicotinoids in seven Indian field populations of Bemisia tabaci genetic groups Asia-I, Asia-II-1, and Asia-II-7. Asia-I and Asia-II-1 populations were showing significant resistance wherein LC⁵⁰ values were 7x for imidacloprid and thiamethoxam, 5x for monocrotophos and 3x for cypermethrin among the Asia-I, while, they were 7x for cypermethrin, 6x for deltamethrin and 5x for imidacloprid within the Asia-II-1 populations. A substantial increase in resistance ratios was observed in both the populations of Asia-I and Asia-II-1. It is evident that potential control failure was detected using probit analysis estimates for cypermethrin, deltamethrin, monocrotophos and imidacloprid in controlling whiteflies due to significant development of insecticide resistance. Insecticide resistance has been reported in public health in India in many insect vectors species.

Chinnababu Naik et al. (2018) reported the mean Resistance Ratio (RR) for cry1Ac against PBW was 47 during 2013 and the has increased to 1387 during 2017. A similar increasing trend was observed for cry 2Ab with a mean RR increase from 5.4 in 2013 to 4196 in 2017. Widespread infestation of pink bollworm in Bt cotton ranging between 40 – 95% accounting for estimated yield losses to the tune of 20-30 % have been reported from 16 major cotton growing districts of Maharashtra, a leading cotton producing state of Central India (Kranthi 2015). This was because of the evolution of resistance in PBW against bollgard II (cry1Ac & cry2Ab) of Bt cotton hybrid in India . The causes of resistance were insufficient refuge, extended growing season, lower expression levels of cry toxin in later crop growth stage etc lead to outbreak of PBW in Indian cotton (Fand et al. 2019). Experts primarily pointed that abundance of refuge varied among countries that might have played a key role in the striking differences in the incidence of the same pest species on the same crop and on the same toxins, without discounting the role of other differing factors like nature of hybrids and varieties, climate and production practice adopted in the three major cotton growing countries in the world. In addition, number of stored grain pests have also developed resistance to insecticides in India. Pesticide resistance in stored grain pests appeared comparatively later and first reported in flour beetle, Tribolium castaneum, in 1971 against DDT and malathion from Delhi followed by lindane and phosphine. *Sitophilus oryzae*, another serious pest of stored grain, becoming resistant to malathion originated from Kanpur in 1973 and also to lindane and phosphine. *Rhizopertha dominica, Ryzaephilus surinamensis, Dermestes maculatus* have also witnessed resistant to malathion, lindane and phosphine in various parts of the country (Mehrotra, 1989).

The pesticide resistance in India was first noticed in insect pests of public health importance and the concern about it led to an International Conference organised pointedly by the World Health Organisation and the Government of India in 1958 at New Delhi. Mosquitoes transmitting malaria and other vector diseases were the first to become resistant to pesticides. This was because of the large scale use of DDT in the National Malaria Control Programme / National Malaria Eradication Programme. The first report of DDT resistance in mosquitoes came in1952 from UP and Bombay in *Culex fatigans* a transmitter of filaria and has been reported to be resistant to both DDT and HCH in various parts of the country. The resistance in urban malaria transmitter, *Anopheles stephensi*, to DDT was reported first from Erode, Tamil Nadu in 1956. In fact *A. culicifacies* which accounts for more than 70% of the rural malaria is resistant throughout the country to one or the other pesticide used in Malaria Control Programme.

In the world, insecticide resistance has been reported in many occasions. Melander (1914) reported the first case of insecticide resistance to lime sulphur, an inorganic insecticide, in an orchard pest, the San Jose scale (Quadraspidiotus perniciosus) in the state of Washington. A treatment with lime sulphur killed all scales in one week in typical orchards, but 90 percent survived after two weeks in an orchard with resistant scales. Subsequently, the number of insecticide resistance cases grew exponentially following widespread use of DDT and other synthetic organic insecticides. Insects have evolved resistance to all types of insecticides including in-organics, DDT, cyclodienes, organophosphates, carbamates, pyrethroids, juvenile hormone analogs, chitin synthesis inhibitors, avermectins, neonicotinoids, and microbials. Since the first case of DDT resistance in 1947, the incidence of resistance has increased annually at an alarming rate. It has been estimated that there are at least 447 pesticide resistant arthropods species in the world (Callaghan, 1991). Insecticide resistance has also been developed by many insects to new insecticides with different mode of action like neoniconitoids. Resistance reported in thirteen orders of insects, yet more than 90 percent of the arthropod species with resistant populations are either Diptera (35 percent), Lepidoptera (15 percent), Coleoptera (14 percent), Hemiptera (14 percent), or mites (14 percent). The disproportionately high number of resistant Diptera reflects intense use of insecticides against mosquitoes that transmit disease,

and agricultural pests account for 59 percent of harmful resistant species while medical and veterinary pests account for 41 percent. Statistical analyses suggest that for crop pests, resistance evolves most readily in those with an intermediate number of generations (four to ten) per year that feed either by chewing or by sucking on plant cell contents.

1990's, neonicotinoids includes imidacloprid, clothianidin, and In thiamethoxam have been introduced into the global market as alternatives to organophosphates and carbamates to control sucking and other pests and they proved good for a while. But subsequently, neonicotinoids have proved the development of resistance in Myzus persicae and Phorodon humuli. The effects of imidacloprid on Nilaparvata lugens, tebufenozide on Plutella xylostella and Spodoptera exigua, thiamethoxam on Bemisia tabaci, trichlorphon on Bactrocera dorsalis, imidacloprid on Spodoptera litura, and emamectin benzoate on Chrysoperla carnea have been (Sahani and Pal 2021). The first report of neonicotinoid resistance was published in 1996, describing the low efficacy of imidacloprid against Spanish greenhouse populations of cotton whitefly. Later-generation, show stronger resistance (up to 17-fold in the first 15 generations) but >80-fold resistance after 24 generations, which has been confirmed in some populations of the whitefly (Bemisia tabaci) and the Colorado potato beetle (Leptinotarsa decemlineata).

Case Histories of Insecticide Resistance

India was one of the first country among third world countries to start a large scale use of synthetic pesticides for the control of insect pests of public health and agricultural importance. The modern era of vector control and plant protection in India started with the introduction of DDT in 1947 followed by HCH in 1949, organophosphates in 1953 and carbamates a little later. Despite the fact that these pesticides have brought immense benefits to the country, they also exhibited serious environmental consequences. It is interesting that DDT and HCH, which have been withdrawn from use in most of the advanced countries of the world, were still being used freely in India for public health.

San Jose Scale Resistant to Lime Sulphur

Melander's (1914) reported the first case of field-evolved insecticide resistance in San Jose scale to lime sulphur that certain populations of insects but not all of them were becoming less susceptible to sulphur-lime than they had been in the past though the chemical was documented to be very effective at killing scale insects previously. Surprisingly, it was found that 90% of the insects that he had sprayed in Clarkston had survived and even when he increased the amount of active ingredient by ten times, still 74% of them still survived. He was of the opinion that San Jose scale should have become acclimatized to a sulphur-lime environment. By consuming repeated small amounts of arsenic the body becomes immune to many times the normal lethal dose. Melander also predicted that entire populations would not become resistant as long as some non-resistant insects survived, because their non-resistant genes would be passed on to future generations. If only the resistant individuals survived to reproduce then resistant line might result after repeated sprayings. But always there are some scales missed by the spraying, and these, during the ten generations between sprayings, will produce a population in part, at least, non-resistant (Levin 2014).

Paradigm Shift to Biopesticides

The insecticide resistance in insects in major crops made the various stake holders to think alternative pest control strategies to safe guard the crops from ravaging insect pests. One such potential alternative is exploration of biopesticides of microbial origin for the management of insects. In India, the development of microbial entomopathogens as insecticides has involved notable successes and failures in the past two decades. India is a tropical and subtropical country with diverse pest and beneficial insect fauna, and with crop losses due to insect pests estimated at 17.5% (valued at US\$17.3 billion). Several classes of customarily used insecticides are now restricted or prohibited due to their harmful effect on the environment, human health, and non-target organisms. Concurrently, the past two decades have witnessed a rise in the use of microbial biopesticides based on entomopathogenic organisms in India.

The global biopesticide market was estimated at approximately \$3 billion, or 5% of total crop protection market, in 2013 and is expected to grow to more than \$4.5 billion by 2023 (Olson, 2015). The value of biopesticides as a part of integrated pest management (IPM) programs has led to the recent increase in their use in India; biopesticides were recently estimated to comprise about 4.2% of the Indian pesticide market (Das 2014). However, market growth has been restricted by slow adoption, limited resources for large-scale production, and challenges associated with regulation and commercialization (Singh et al. 2016). Undoubtedly, microbial biopesticides play vital role in controlling the desirable pests and gaining interest among the population with advantages like non toxic mechanism, eco-friendly nature, efficacy and suitability in the Integrated Pest Management programmes unlike synthetic insecticides.

Biopesticides, an alternative to chemical pesticides, are typically derived from living organisms, microorganisms, and other natural sources pose less risk to people and the environment and hence gain worldwide attention as a new tool to kill insects. Biopesticides are being widely used to manage biotic stresses as

a component of IPM under protected cultivation (Ramasamy and Ravishankar, 2018). On considering the international market of export commodities and health conscious of Western countries, the role of microbial biopesticides in pest management would address their importance among the growers and consumers which ultimately enhance the marketability of microbial biopesticides in India. Development and promotion of biopesticides usage in India need to be well addressed through promoting their manufacture at village level as an ancillary profession to agriculture. Registration process of biopesticides in India may be simplified without compromising quality and authenticity of the product. Policy decisions regarding production, development, promotion of biopesticides in India would definitely attract more scope in near future. A strict follow up of the policies pertaining to promotion and use of biopesticides will encourage inviting definite foreign exchange, besides producing healthy food commodities in India. Enormous scope for biopesticide market in India if the industry and extension functionaries convince farmers and pesticide manufacturing companies for better utilization of biopesticides in India as they are cheap, economical, viable, durable and effective.

Way Forward

The best way to overcome insecticide resistance is to reduce selection pressure and preserve the finite resources of new and useful compounds by adopting resistance management strategies in an integrated approach. Careful and systematic planning of insecticide application includes monitoring of resistance genes (or the associated enzymes or channels) in pest populations, as is now feasible for many of the mutated targets will help in partly in delaying the resistance. Resistance management is often necessary to shift to new compounds acting on novel targets that once again minimize selection pressure. This process of continually shifting approaches may ultimately be limited by a finite number of practical targets for pest control. Adopting the integrated pest management (IPM) approach usually helps with resistance management by retaining some susceptible populations along with resistant individuals.

The best way to delay onset of evolution of resistance in pests to minimize insecticide use and integration of chemical and non-chemical controls to seek safe, economical, and sustainable suppression of pest populations. The nonchemical approaches such include biological control by predators, parasitoids, and pathogens. Also valuable are cultural control through crop rotation, manipulation of planting dates to limit exposure to pests, and use of cultivars that tolerate pest damage and mechanical controls by exclusion using barriers and trapping. Tank mixing pesticides is the combination of two or more pesticides with different modes of action in order to improve individual pesticide application results and delay the onset of or mitigate existing pest resistance. Exploration and utilization of botanicals, biopesticides of biological origin to achieve target production as alternatives to synthetic chemical insecticides would serve the purpose. Educating, equipping and encouraging farmers to utilise biological pesticides to reduce the cost of protection, environmental pollution, without compromising yield and market price.

Conclusions

Although insecticides have been proved as one of the best management options, looking into growing demand for organic production and health consciousness of consumers across the world, adverse effect of environmental pollution and health hazards, biological based pest management strategies such as biological control, microbial control and have to be intensified and encouraged. Biopesticides, an alternative to chemical pesticides, are typically derived from living organisms, microorganisms, and other natural sources pose less risk to people and the environment and hence gain worldwide attention as a new tool to kill insects. Biopesticides are being widely used to manage biotic stresses as a component of IPM under protected cultivation. Development and promotion of biopesticides usage in India need to be well addressed through promoting their manufacture at village level as an ancillary profession to agriculture.

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Links

https://entomologytoday.org/2014/04/08/the-first-journal-article-on-insecticide-resistancewas-published-100-years-ago-this-month/downlaoaded dated 22 Dec 2021.

History and Development of Biological Control

Abstract

Chinese farmers were the first to apply biological control of agricultural pests when they used red ants to manage pests of fruit crops. Neem-based products were then used as fertiliser and as a barrier against pests of stored goods. Later, biopesticides surpassed macro-biocontrol agents in prominence and were added as one of the elements of integrated pest management. The interventions and directives of the United States Environmental Protection Agency and European Food Safety Authority were for way forward to the evolution of pesticides derived from naturally occurring organisms and plant materials. The development of biopesticides in agriculture across the world and India is reviewed in this chapter, along with the contributions of the Canada Department of Agriculture, Common Wealth Institute of Biological Control, Department of Biotechnology, New Delhi, Biotech Consortium India Ltd., New Delhi, etc. Since the Indian Council of Agricultural Research, New Delhi, launched its AICRP-Biocontrol programmes, a dramatic rise in the use of biopesticides has been observed in India.

Keywords: Microbial Biopesticide, History, USEPA, EFSA, ICAR initiatives

Introduction

The biocontrol use had almost completely disappeared due to the growth and success of the synthetic pesticide industry in the mid-1940s. The publication of Rachael Carson's 'Silent Spring' (Carson 1962) which condemned the use of agricultural pesticides and emphasized their harmful environmental effects on wildlife. Due to public outrage over this controversial book, there was a need for pesticide alternatives, which presents an opportunity for wider use of biological control (Barratt et al. 2010; Gay 2012). Many naturalists and environmentalists began looking for new insecticides with innovative chemical

structures that would have less harmful effects on people, animals, and the environment (Barratt et al. 2018).

In 1901, Japanese biologist Shigetane Ishiwata discovered spores of the bacterium, Bacillus thuringiensis (Bt) from a sick silkworm. This bacteria is still the most often used biopesticide today (Chen 2014; Glare et al. 2000). Sporeine, the first Bt product to be sold commercially, debuted in 1938. The extensive usage of biopesticides started in the 1950s in the US. A low level of research and development was maintained in the second half of the 20th century as a result of the widespread use of synthetic chemical insecticides and World War II. The Pacific Yeast Product Company created the submerged fermentation industrial process in 1956, allowing for the largescale manufacturing of Bt (Glare et al. 2000). In 1973, Heliothis NPV was granted exemption from tolerance and the first viral insecticide, Elcar received a label in 1975. In 1977, B. thuringiensis var. israelensis, which is poisonous to flies, was reported in 1977, while the strain tenebrionis, which is poisonous to beetles, was found in 1983. Following the demonstration by environmentalists and ecologists that widespread and repeated application of these synthetic chemicals could be ecologically detrimental, biological pest management was nevertheless expedited (Cook and Baker 1983).

Earlier, biocontrol agents like some predatory insects (red ant) and birds were engaged for insect pest management (Brahmachari 2004). Later few botanicals including various parts of neem tree (*Azadirachta indica* A. juss) and its extracts were tried as fertilizers and also to protect stored cereals from post-harvest losses (Isman 1997; Schmutterer 1985).

The concept of Integrated Pest Management (IPM) had come to the field of Agriculture during 1960s, in which judicious use of various methods of control was emphasized to overcome the ill-effects of chemical pesticides (Smith and Bosch 1967). Later on, based on the recommendation of US National Academy of Sciences, biological control with natural enemies and microbial biopesticides was included one of the components in IPM (Peshin et al. 2009). Control failure of few polyphagous cotton feeders including American boll worm, *Helicoverpa armigera* and generalized defoliator, *Spodoptera litura* with chemical pesticides alerted to switch over biological control, a safe, cost-effective, and eco-friendly method (Kranthi et al. 2002).

By the mid 1920s, entire British Empire was active in biological control work including Australia, New Zealand, Fiji, Canada, Bermuda and South Africa. In 1927, the Imperial Bureau of Entomology (IBE) created facilities for conducting biological control work in Farnham House Lab, England. It was under the control of W. R. Thompson in 1928 who initially concentrated on

natural enemies of insect pests and broadened to work on biological control of weed in 1929. In 1929, Canada Department of Agriculture (CDA) constructed a biological control lab at Belleville, Ontario. In 1940, this lab was moved to Ottawa, Canada where it became as Imperial Parasite Service. In 1947, it became independent and designated as Commonwealth Bureau of Biological control (CBBC). In 1951, it was renamed as Commonwealth Institute of Biological Control (CIBC). In 1961, the CIBC headquarters were transferred to Trinidad, West Indies. The CIBC identified two sub-stations in south East Asia, one at Bengaluru in India and another at Rawalpindi in Pakistan to undertake biological control research. In 1957, the India station of CIBC was established to initiate organized and systematic research in biological control.

Aristotle was the first to mention in his book "Historia Animalia" that honey bees suffered a disease which was later identified as foul brood disease. One chapter on diseases of Insects was included in the book entitled 'An Introduction of Entomology'. Disease can be defined as a departure of the insect from a state of health and was first noticed among domesticated insects. In Europe, Aristotle was the first to mention that bees suffered disease and in 1835, Agostino Bassi showed that animal disease could be caused by a microorganism, when he found that the fungus *Beauveria bassiana* causes the muscardine disease of silkworms. Early observations were largely concentrated on two domesticated insects, the honey bee and silkworm. Gradually these studies were extended to pest species too, and the concept of utilizing disease to control these insects was born.

In 1879, the Russian, Metchnikoff, conducted the first significant experiments in the destruction of injurious insects with micro organisms by infecting larvae of the beetle *Anisoplia austriaca* with the fungus *Metarhizium anisopliae*. The first commercial product, Sporeine, containing *Bacillus thuringiensis* was produced before 1938. After the second world war, several commercial firms in the USA, began to produce this bacterium. Today we can point to such achievements as the protection of over 50% of the cole crops from the cabbage looper, *Trichoplusia ni* in Southern California by *B. thuringiensis* in 1965 and 1966. The importance of efficient, eco-friendly methods for pest and disease control gained momentum. Steady growth of biological control has been reported in various eras including Ancient origins, North American Beginnings, California origins and 20th Century developments with several explorations and examples for successful management of many key pests of crops.

20th and 21st Century Developments

Many chemical pesticides were withdrawn from the market in the second half of the 20th century as a result of inappropriate application techniques used during World War II, such as aerial application (Howard 1935), which had numerous negative consequences, including acute or chronic toxicity (Hunt and Bischoff 1960), as well as other unfavorable effects like increased resistance in the target species (Mouches et al. 1986), the replacement of target species with more dangerous resistant species (Regnault-Roger et al. 1986), and contamination of different environmental compartments (Ellgehausen et al. 1980; Leduc et al. 1987). Authorities from the US Environmental Protection Agency (USEPA) and the European Food Safety Authority (EFSA) revised the pesticide laws when these issues arose in order to protect human and animal health as well as the environment from the risks associated with pesticides. They proposed the ideas of the ideal pesticide, which include i) a high selectivity to target species but a minimal toxicity to non-target organisms, ii) a high effectiveness at a low application rate, and iii) a low environmental persistence. Thus a new concept on 'Biopesticide' had evolved to fight with pests effectively but have minimal impacts on humans, animals, and the environment. Active biopesticide research has expanded in the latter decade of the 20th century along with a notable increase in publications (Shukla et al. 2019) (Fig. 1).

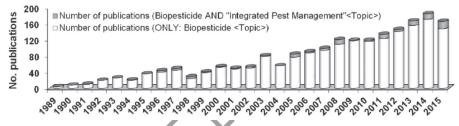


Fig. 1. Publications on biopesticide research from 1989 to 2015 (Web of Science 2015).

Around 1400 biopesticide products were sold globally at the start of the twentyfirst century, making up around 2.5% of the entire pesticide business (Marrone 2007; Chandler et al. 2012). From 2012 to 2017, the demand for biopesticides was predicted to increase at a faster compound annual growth rate (GAGR) of 16.1% (compared to 3%) than that of synthetic pesticides, resulting in an estimated \$ 5.2 billion global market in 2017 (Lehr 2010). Europe, Oceania, and Latin America accounted for 20, 20, and 10% of worldwide biopesticide consumption, respectively, whereas North America consumed roughly 40% of them (Leng et al. 2011). Numerous legislative initiatives for the sustainable use of pesticides were implemented, and they all emphasized how crucial it is to use less chemical pesticides overall to avoid potential environmental damage. When the All India Coordinated Research Project on Biological Control of Crop Pests and Weeds (AICRPBC) was established in 1977, the biological control effort received a further boost. In 1993, the AICRPBC was elevated even further to the position of Project Directorate of Biological Control (PDBC). The goal of the PDBC was to do basic and applied research on the biological control of crop pests and weeds across the nation. With 16 sites dispersed throughout the nation, PDBC serves as the central agency in India. The name of PDBC has been modified twice. PDBC was upgraded to become the National Bureau of Agriculturally Important Insects (NBAII) during the XIth 5-year plan (2009), and in the XIIth 5-year plan, it was renamed as National Bureau of Agricultural Insect Resources (NBAIR).

In 1989, the National Bio-control Network Program was also introduced by the Department of Biotechnology (DBT) in New Delhi. Ten R&D projects were launched at the beginning of the programme with a duration of five years (1989-1994). Over 200 projects were carried out in numerous national institutes and state agricultural universities (SAUs) once the programme was later expanded (Wahab 2004). IPM was included in the government of India's National Policy Statement in 1985, but the Department of Agriculture and Cooperation (DAC), Ministry of Agriculture, took a significant step by announcing a programme on "Strengthening and Modernization of Pest Management Approach in India in 1991-1992," along with the establishment and strengthening of biocontrol research at the regional level. Department of Biotechnology is one of the top financing organisations in India with programmes for biocontrol research (Singh et al. 2002; Sharma et al. 2003; Mishra et al. 2020). Currently, in addition to DBT, other funding organisations like the Department of Science and Technology (DST), New Delhi, and the Indian Council of Agricultural Research (ICAR) have also introduced a number of programmes with a major focus on prudent pesticide use in agriculture and supporting research on biopesticides. These government organisations are, however, also highly concerned with accumulating toxicological information regarding the use of biopesticides in diverse agro-climatic zones.

The first commercial biocontrol agent manufacturer in India was Bio-Control Research Laboratories (BCRL), a branch of Pest Control India (PCI) Limited working under a contract with the Plant Protection Research Institute (PPRI) (Manjunath 1992). Currently, *Trichoderma viride, Trichoderma harzianum*, and *Beauveria bassiana* are antagonistic bacteria and fungi that are manufactured and sold by the BCRL. Later, IPM was backed by the National Policy on Agriculture (2000) and the National Policy for Farmers (2007). In India, a total of 970 biopesticide formulations have been registered in Central Insecticide Board and Registration Committee (CIB&RC) as on 1.1.2020 in which *Trichoderma* and *Pseudomonas* based formulations dominate (http://164.100.161.213/divisions/cib-rc/bio-pesticide-registrant). A compendium consisting of 31 microbial formulations which are in different stages of registration and commercialization were reported to possess 3 to 25 months shelf-life at 25°C to 35°C (Saxena et al. 2021).

Conclusions

The history and expansion of biopesticide in agriculture around the world clearly showed that a phenomenal growth was reported during the second half of the 20th century and the beginning of the 21st century. The United States Environmental Protection Agency and the European Food Safety Authority's interventions, which involved revising the laws, greatly aided in the registration of plant protection products that contained less harmful ingredients to replace conventional pesticides or serve as a starting point for the synthesis of novel chemistries. The development of biopesticides was thought to be based on extraction from natural sources, chemical synthesis, and computational chemistry. Additionally, biopesticides are not harmless and are subject to the same laws and regulations as chemical pesticides (Villaverde et al. 2016). More than 900 biopesticide formulations have been certified through the Central Insecticide Board and Registration Committee, New Delhi. Over 31 bacterial and fungal-based biopesticide formulations are in pipeline to pass various stages of commercialization in India.

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Links

http://164.100.161.213/divisions/cib-rc/bio-pesticide-registrant



Biopesticides Classification and Their Formulations in India

Abstract

The fungal based biopesticides especially Trichoderma spp., are abundant in Indian market than other microbe based formulations. It is vice-versa in Western countries wherein Bacillus thuringiensis based products are being used in wide spread control of Lepidoptera pests of temperate crops. Talc and oil based biopesticide formulations are most common in India as they as fillers could be able to prolong the pathogenicity by enhancing shelf-life. List of biocontrol laboratories and important companies along with their products are tabulated and furnished to understand the scope of biopesticide in India.

Keywords: Biopesticide, Classification, Formulations, Biocontrol labs

Introduction

Three kinds of biocontrol products are being used in India which include 1) Microbial biopesticides, 2) Plant-derived botanical pesticides, and 3) Pheromones or other natural insect growth regulators. Microbial biopesticides consist majorly of fungi, bacteria, viruses or entomopathogenic nematodes as bioactive principles. In India, fungal based biopesticides share in the market is maximum (Mishra et al. 2020; Fig. 1,2,3) while *Bacillus thuringiensis* based products rank first in usage in USA, Europe etc. Among fungal biopesticides, *Trichoderma* (only two species) based products approximately around 355 are readily available in the Indian market for the field applications (Kumar et al. 2019). *Pseudomonas fluorescens* based products are in wide spread usage over *Bacillus* based products globally with reference to bacterial biopesticide (Mishra et al. 2020). In India, strains of *B. thuringensis, Bacillus sphaericus* and *Bacillus subtilis* are registered as biopesticides. Other non-spore forming bacteria like, *Serratia entomophila* and *Chromobacterium subtsugae* though have efficacy on limited range of insects are not evaluated systematically

(Martin et al. 2007). In India, two nucleopolyhedron viruses (NPVs) of *Helicoverpa armigera*, and *Spodoptera litura* are in use under field conditions in a smaller extent. Though the natural occurrence of granulovirus (GVs) of sugarcane pests in southern and northern states of India was reported, their commercial production are not yet started (Easwaramoorthy and Jayaraj 1987).

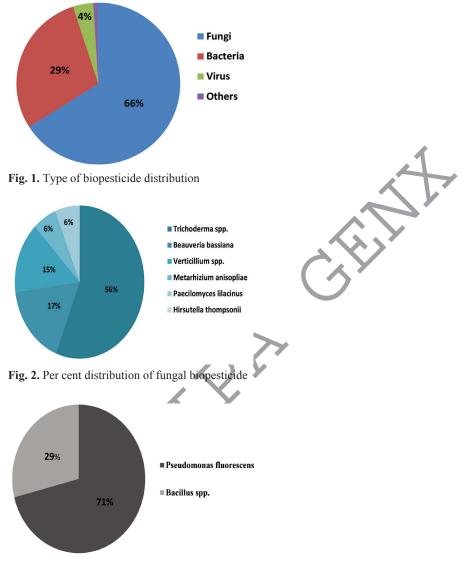


Fig 3. Per cent distribution of bacterial biopesticide (Mishra et al. 2020)

The fate of using two most effective entomopathogenic nematodes belonging to the family Heterorhabditis and Steinernema is questionable, however their applications against different soil-borne pests under field conditions are scarce (Sankaranarayanan et al. 2006) as no registered product is available in the market to date in India. Among the botanicals, neem-based products are in wide spread use in plant protection when compared to other plant-based products such as pyrethrum, eucalyptus leaf extract, and *Cymbopogon nardus* is less common at field level (Dougoud et al. 2019). Cotton, sugarcane, and rice are the three major crops wherein pheromone technology is being explored to monitor and mass trap pest population (Khergamker 2019). Recently, few pilot projects have been initiated in few parts of India to up-scale pheromone technology in the management of crop pests (Wadke 2018).

Common Method of Mass-production of Biopesticides

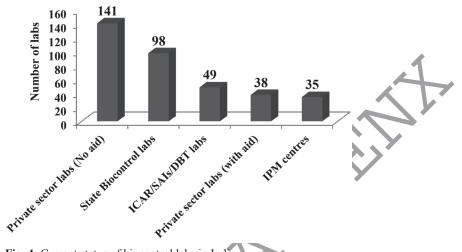
SAUs mostly follow solid-state fermentation (SSF) and sub-merged fermentation (SF) process for multiplication of microbe. At the level of villages, few indigenous techniques are followed for production. However, limited production and use of microbial biopesticide is experienced so far due to want of sophisticated machineries. Many private companies as of now posses the capacity to produce 10 to 2000 tons per annum in India (https://ncof. dacnet.nic.in/Opera tiona l_Guide lines/Guide lines_for_Capit al_Investment _Subsidy.pdf).

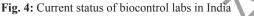
Biopesticide Formulations

CIBRC registration guidelines emphasized few quality parameters of biopesticide formulations for registration and before entering into market which include safe and effective, easy delivery with prolonged shelf-life. At present there are only few biocontrol products that strictly adhere to CIBRC guidelines. In India, wettable powder (WP), wettable granules (WG), suspension concentrates (SC) and aqueous suspension (AS) formulations are being used. Currently, biocontrol products are formulated in solid carriers which include talc, peat, lignite, clay, wheat husk, rice bran, grinded corn cob, fly ash and sawdust. Bacillus based products are being sold in the form of aqueous suspension, dust, WP and granules, charcoal, plaster of paris and fly ash (Tikar and Prakash 2017). Currently, the liquid formulations of Bt products and viral biopesticides are very popular in the market. New Nanotechnologybased biopesticide formulations such as nanoencapsulation and nanoemulsions can improve the large scale applicability of bioagents under field conditions (Koul 2019). However, these techniques are still in infancy and yet to be commercialized at the industrial level. However, the survival of infecting units of microbe in various carriers is remarkably debatable.

Biopesticide Mass-production Units in India

The number of bio-production units has currently increased to 361, of which 141 are in the private sector without GOI grant aids and 38 with GOI grant aids. Moreover, the Ministry of Agriculture and Farmers Welfare has assisted about 35 IPM centers to produce biopesticides since 2010 (Keswani et al. 2016). A total of 98 State Biocontrol Laboratories were established by the State Departments of Agriculture and Horticulture of Gujarat, Uttar Pradesh, Karnataka, Tamil Nadu, Andhra Pradesh and Kerala as well as the production of microbial pesticides by the Institutions of the Indian Council of Agricultural Research (Pathak et al. 2017; Mishra et al. 2020' Fig. 4).





Central and state agricultural universities and various ICAR institutes including Tamil Nadu Agricultural University (TNAU), Coimbatore, Tamil Nadu, ICAR-Central Plantation Crops Research Institute (CPCRI), Kayamkuklam, Kerala, Kerala Agricultural University (KAU), Kerala, ICAR-Indian Institute of Horticultural Research, Bengaluru, ICAR-Central Research Institute for Dryland Agriculture, Hyderabad and ICAR-Directorate of Oilseed Research (ICAR), Hyderabad are popular biopesticide production units in Southern parts of India. In the northeast, Assam Agriculture University and Central Agricultural University, Manipur are producing biopesticides against invasive pests. In north India, ICAR-Indian Agricultural Research Institute (IARI), New Delhi, Punjab Agricultural University (PAU), Punjab and G.B. Pant University of Agriculture and Technology (GBPUA & T), Uttarakhand are involved in the production of biopesticides. ICAR-Indian Institute of Sugarcane Research (IISR), ICAR-Central Institute for Subtropical Horticulture, and Directorate of Plant Protection Quarantine & Storage in Lucknow, are major government agencies which are mass producing biopesticides in Uttar Pradesh. Many ICAR sponsored Krishi Vigyan Kendras (KVK), State Government sponsored state biocontrol labs and National Agricultural Co-operative Marketing Federation of India (NAFED) are in the full time job in production of biopesticides.

In India, public sectors contribute 70% of the biopesticides production. Major companies are Biotech International Ltd., New Delhi, International Panaacea Ltd, New Delhi, Ajay Biotech (India) Ltd., Pune, Bharat Biocon Pvt. Ltd., Chhattisgarh, Microplex Biotech and Agrochem Pvt., Mumbai, Excel Crop Care Ltd., Mumbai, Govinda Agro Tech Ltd., Nagpur, Jai Biotech Industries, Satpur, Nasik, Ganesh Biocontrol System, Rajkot, Gujarat Chemicals and Fertilizers Trading Company, Baroda, Gujarat Eco Microbial Technologies Pvt. Ltd., Vadodara, Chaitra Agri-Organics, Mysore, Deep Farm Inputs (P) Ltd., Thiruvanandapuram, Kerala, Kan Biosys Pvt. Ltd., Pune, Indore Biotech Inputs and Research Pvt. Ltd., Indore, Romvijay Biotech Pvt. Ltd., Pondichery, Devi Biotech (P) Ltd., Madurai, Tamil Nadu, T. Stanes and Company Ltd., Coimbatore, Tamil Nadu, Harit Bio Control Lab., Yavatmal and Hindustan Bioenergy Ltd., Lucknow. Few Indian companies which work in biopesticde production in collaboration with foreign companies are Lupin Agro-chemicals, Mumbai; Sugar and distillery companies such as KCP Sugar and Industries Corporation Ltd., Andhra Pradesh, Rajshree Sugars and Chemicals Ltd., Tamil Nadu; New Swadeshi Sugar Mills, Bihar, and Bannari Amman Sugars Ltd., Tamil Nadu.

Biopesticide Registrant in India

As on 1.1.2021, a total of 970 biopesticides registered in India by CIB&RC under the 1968 Insecticide Act which include microbial biopesticides of *Bacillus thuringiensis* var. kurstaki (42), var. israelensis (22), var. sphaericus (05), var. galleriae (01), *Pseudomonas fluorescence* (196), *Bacillus subtilis* (04), *Trichoderma viride* (289), *T. harzianum* (51), *Ampyliomyces quisqualis* (02), *Beauveria bassiana* (106), *Metarhizium anisopliae* (30), *Verticillium lecani* (93), *Verticillium chlamydosporium* (03), *Helicoverpa armigera* NPV (30) and *Spodoptera litura* NPV (03) (Kumar et al. 2018; Keswani et al. 2019; http://ppqs.gov.in/divisions/cib-rc/biopesticide-registrant) (Table. 1).

S.No.	able 1: List of microbial biopesticide formulations registered in Indian S.No. Name of Combany Organism Tra	Iormulations registered in in Organism	uian Trade name	Target biotic stress
	M/s Rallis India I td	Racillus thurinoionsis	Bohit IIWP	L'enidontera insects
:	Bengaluru	a and shi min a min		
2.	M/s Sandoz (I) Ltd., India	<i>Bacillus thuringiensis</i> var. kurstaki	Deflin WG	Lepidoptera insects
3.	M/s Aventis Crop science I td Bengaluru	Bacillus thuringiensis var. israelensis (H_1A)	Vectobac 12 AS	Diptera insects
4.	nd Companey	Beauveria bassiana	Biopowder WP	Sucking insects, bollworms
5.	ries Ltd.,	NPV	Biovirus-H	Helicoverpa armigera
6.	ontrol India Ltd.,	NPV	Spodocide 0.50% AS	Spodoptera litura
	Bengaluru	NPV	Helicide 0.50% AS	Helicoverpa armigera
7.	M/s Anshul Agrochemicals,	Beauveria bassiana	Green Heal	Sucking insects, borers, bollworms
	Bengaluru	Beauveria bassiana	Beveroz-L	Sucking insects, borers, bollworms
		Beauveria bassiana	Almax	Sucking insects, borers, bollworms
		Pseudomonas Aorescens	Pseudomax	Soil and seed borne diseases
		Trichoderma viridi	Trichomax	Soil borne diseases, plant parasitic nematodes
8.	M/s Deepa Farm inputs	Pseudomonas florescens	Bio-Plus Pseudo	Soil and seed borne diseases
	Private Limited	Trichoderma viridi	Bio-Plus Tricho	Soil borne diseases, plant parasitic nematodes
		Verticilium lecanii	Bio-Plus Verticillium	Sucking pests, plant parasitic nematodes
		Metarhizium anisopliae	Bio-Plus Metarhizium	Beetles, soil arthropods
9.	M/s Kan biosys Pvt. Ltd.,	Trichoderma harzianum	Nemastin 1% WP	Root knot nematode
	Pune	Trichoderma viridi	Combat 1% WP	Root rot, damping off, wilt
		Beauveria bassiana	Brigade-B 1.15% WP	Rice leaffolder
		Pseudomonas florescens	Sudo 0.5% WP	Late leaf spot of groundnut
			~	

Table 1: List of microbial biopesticide formulations registered in Indian

S.No.	S.No. Name of Company	Organism	Trade name	Target biotic stress
	M/s Peak Chemical Industries Ltd., West Bengal	Paecilomyces fumosoroseus	Bardan	Spider mite, parasitic mites
_	(Metarhizium anisopliae	Moti	Termite
		Beauveria bassiana	Badsha	Sucking pests
		Verticilium lecanii	Victor	Parasitic nematodes, whitefly, thrips, aphids
	M/s Uttam Chemicals Industries. Raiasthan	Trichoderma harzianum	Trichoderma harzianum 1% WP	Parasitic nematodes
		Trichoderma viridi	Trichoderma viridi 1.5% WP	Nematicide, crop diseases
	M/s Ambic Organic, Surat	Paecilomyces fumosoroseus	Almite	Mites, DBM, sucking pests
	M/s Criyagen Agri & Biotech Pvt. Ltd., Bengaluru	Trichoderma viridi	Trichoderma	Soil borne fungus
	M/s Biotech International	Bacillus thuringiensis var.	BIOLEP WP	Helicoverpa, Spodoptera, DBM, borers, hairy
	Ltd., New Delhi	kurstaki		caterpillars, cut worms, army worms, leaf rollers & miners, skeletonizers & Defoliators
		Beauveria bassiana	BIORIN WP/AS	Helicoverpa, Spodoptera, DBM, leaf borer,
			Ć	hairy caterpillars, mites, spidermites, whiteflies, aphids, scale insects, locusts & colorado beetles
		Verticillium lecanii	BIOLINE WP/AS	Whitefly, green leaf hopper, thrips, mealy bug,
			Ś	brown plant hopper, leaf miner, aphids, mites, jassids
		Metarhizium anisopliae	BIOMET WP/AS	White grubs, termite, cut worm, caterpillars, semiloopers, sucking pests, mealybugs & aphids

S.No.	S.No. Name of Company	Organism	Trade name	Target biotic stress
		NPV	BIOVIRUS-H AS	Helicoverpa armigera
		NPV	BIOVIRUS-S AS	Spodoptera litura
	(Pseudomonas fluorescens	BIOMONAS WP/AS	Bacterial wilt, black rot, bacterial spot, sheath
	C			blight; blast, anthracnose, powdery & downy
				Mildew, panama wilt, panama wilt, sigatoka,
)			bacterial leaf spot
		Trichoderma viride	BIODERMA WP/AS	Root rot, stem rot, damping off, fusarium wilt
				and verticillium wilt, all types of leaf spot, leaf
				& blight
		Trichoderma harzianum	BIODERMA-H WP/AS	Root rot, stem rot, damping off, fusarium wilt
				and verticillium wilt, all types of leaf spot, leaf
				& blight
		Bacillus subtilis	BIOSUBTILIN WP/AS	Fusarium wilt, Macrophomina, damping off,
)		Pythium, Rhizoctonia Black scarf of Potato,
				root rot, leaf spot, powdery and downey mildew,
				bacterial spot & bacterial leaf blight
		Ampelomyces quisqualis	ARMOUR WP/AS	Powdery Mildew in pulses, vegetables, fruits &
		•		ornamental crops
		Paecilomyces lilacinus	BIONEMAT WP/AS	Root-knot nematodes, reniform nematodes,
				cyst nematode, golden cyst nematodes, citrus
				nematodes, lesion nematode
		Bacillus firmus	NEMATO CURE WP/AS	Root-knot nematodes, reniform nematodes,
				cyst nematode, golden cyst nematodes, citrus
				nematodes, lesion nematode
		Hirsutella thompsonii	NO MITE WP/AS	Various types of crop mite - Scoulet, purple, red
				spider mite
			7	

S.No.	S.No. Name of Company	Organism	Trade name	Target biotic stress
15.	M/s International Panaacea	Trichoderma viride	Sanjeevni 1.0% WP	Fusarium, Charcoal rot, Black scurf, Karnal
	Ltd, New Delhi			bunt, Silver leaf of plum & peach, Rhizoctonia,
	(Pythium, Schlerolims, Verticillium, Alternaria
		Trichoderma harzianum	Bioharz 2% AS	Fusarium, Charcoal rot, Black scurf, Karnal
				bunt, Silver leaf of plum & peach, Rhizoctonia,
				Pythium, Sclerolims, Verticillium, Alternaria
		Pseudomonas flourescens	Rakshak 1% WP	Soil and seed borne diseases
		Trichoderma viride	Bokashi Bran	Soil and seed borne diseases
		Trichoderma viride	Seed2plant	Soil and seed borne diseases
		Trichoderma viride	Tricho-PEP H 1% WP	Diseases
			Trichoz-P 1.5% WP	root rot, wilt, stem rot
		Paecilomyces lilaecinus	Nematofree 1% WP	Helicoverpa, Spodoptera, borers, hairy
				caterpillars. pest of vegetables and fruits,
				whitefly, aphids, DBM, scale insects, locust,
				Colorado potato beetles
		Pseudomonas fuorescens	Bactvipe 2% AS	Root rot, stem rot, collar rot, wilt, blights, leaf
		>		spots, anthracnose, Alternaria and downy &
				powdery mildews
		Ampelomyces quisqualis	MilGO 2% AS	Powdery mildew, Alternaria solani,
		Ŭ		Colletotrichum, Coccodes, Cladosporium,
				cucumerinum
		Beauveria bassiana	DAMAN 1% WP	Helicoverpa, Spodoptera, borers, hairy
				caterpillars, pest of vegetables and fruits,
				whitefly, aphids, DBM, scale insects, locust,
				Colorado potato beetles
		Bacillus subtilis	MILDOWN 2% AS	Pythium, Alternaria, Xanthomonas,
				Rhizoctonia, Botrytis, Scelerotiana,
			~	Phytophora

No.	S.No. Name of Company	Organism	Trade name	Target biotic stress
	M/s Ajay Biotech (India) Ltd, Pune	Bacillus thuringiensis var. Kurstaki	Bio Dart	Insecticide
	(Metarhizium anisopliae	Meta Guard WP/AS	Termites, white grubs, soil insects
		Beauveria bassiana,	AJAY VBM	Insect pests
	\mathbf{D}	Metarhizium anisopliae,		
		Verticillium lecanii		
		Paecilomyces spp.	Nemadart	Parasitic nematodes
		Consortia of microbes	TERMINIL	Termites
		Trichoderma viride	Trident 1.5% WP	Soil and seed borne diseases
		Trichoderma harzianum	Trichoshield	Soil borne fungus
		Trichoderma viride	ROM Trichoderma	Rhizome rot of turmeric and cardamom,
				fusarium wilt of banana, wilt disease of pepper,
				beetelvine, chillies, tomato and vegetables,
				Phytophthora, Pythium, Sclerotium
	M/s Bharat Biocon Pvt. Ltd.,	Pseudomonas fluorescens	Pseudocon 0.5% WP	Crop diseases
	Chhattisgarh	Trichoderma harzianum	TRICHOCON 1% WP	Crop diseases
		Beauveria bassiana	BASICON 1.15%	Lepidoptera insects
		Verticillium lecanii	VERTICON 1.15%	Plant parasitic nematodes
		Metarhizium anisopliae	METACON 1.15%	Beetle pests, locust
	M/s Jai Biotech Industries,	Verticillium lecanii	Vertimust 1.1% WP	Sucking pests
	Satpur, Nasik	Beauveria bassiana	Beauvera 1.15% WP	Lepidoptera pests
		Trichoderma viride	Jaimold 1% WP	Sclerocium, Rhizoctonium, Pythium, Fusarium
			*	

<u> </u>	S.No. Name of Company	Organism	Trade name	Target biotic stress
19.	M/s Ganesh Biocontrol	NPV	Biokills	Spodoptera litura
_	System, Rajkot	Pseudomonas fluourescens Monas	Monas	Rhizoctinia, Sclerotini, blights & Alternaria,
	(Ascochyta, Cercospora, Macrophomina,
_				Xanthomonas, Erwinia
_		Beauveria bassiana	BASS 1.5% WP	Helicoverpa, Spodoptera, DBM, borers, hairy
_)			caterpillar of vegetables & fruit plants, mites
_				& spidermites of vegetables & ornamentals,
-				whiteflies on cotton & vegetables, aphids &
				scale insects, locust, potato and Coffee pod -
				borer
	M/s Gujarat Eco Microbial	Trichoderma viride	NR-III	Soil & air borne pathogens, sucking pest
_	Technologies Pvt. Ltd., Vadodara	Trichoderma viride	TRIOJEET	Damping off, wilt, root rot
21.	M/s Indore Biotech Inputs	NPV	Helicop AS	Helicoverpa armigera
	and Research Pvt. Ltd.,	Metarhizium anisopliae	BIO-MAGIC	Grub, termite, fruit flies, hopper, wire worms,
_	Indore			vegetable worms, aphids, jassids
_		Beauveria bassiana	BIO-WONDER 1.15%	Hairy insects, aphids, white flies, mealy bugs,
_			WP	grasshoppers, thrips, stem borer, termites,
				beetles, caterpillars
_		Verticillium lecanii	Vercitile 1.15% WP	Aphids, whiteflies, thrips, mealy bugs, scale
-				insects, leaf hopper, mango hopper
_		Bacillus thuringiensis var.	Cezar 0.5% WP	Castor semilooper, soybean and gram pod borer,
_		kurstaki	へ	Spodoptera litura, Bihar hairy caterpillar, sphinx
_		, , ,		
_		Paecilomyces lilacinus	BioAce 1% WP	Plant parasitic nematodes
_		Trichoderma viride	Biohit WP	Pythium, Fusarium oxyforum, Rhizoctonia
_				solani, Alternaria, Sclerotinia rolfsia
_		Pseudomonas fluorescens	Biomonarch	Macrophomina, Fusarium, Rhizoctonia, Sclerotium, Pythium

S.No.		Organism	Trade name	Target biotic stress
22.	M/s Romvijay Biotech Pvt.	Hirsutella thompsoni, Verticillium lecanii	Biomite	All species of mites
		Beauveria, Bacillus subtilis ROM Grub kill	ROM Grub kill	Beetles, borers
		Metarhizium anisopliae	ROM Meta kill	Beetles, borers
		Beauveria bassiana	ROM Beevicide	Beetles, borers
)	Paecilomyces lilacinus	ROM Pelicide	Plan parasitic nematodes
		Verticillium lecanii	VERELAC	Sucking pests
		Trichoderma viride	ROM Trichoderma	Wilt, rot, plant parasitic nematode
		Paecilomyces lilacinus	ROM TRY PAE MIX	Wilt, plant parasitic nematode
		Ampelomyces quisqualis	ROM No-Mildew	Mildew diseases
	>	Pseudomonas fluorescens	ROM Pseudomonas	Foliar and soil borne diseases
23.	M/s Devi Biotech (P) Ltd.,	Trichoderma viride	Boom Derma 1.5% WP	Damping off, wilt, collar rot, root rot, leaf
	Madural, Tamil Nadu	$\mathbf{\hat{b}}$		blights spots
		Paecilomyces lilacinus	Boom Nemo 1% WP	Root knot nematodes, cyst nematodes, reniform
				nematode, burrowing nematode, citrus
				nematode, golden cyst nematode and lesion
				nematodes
		Pseudomonas fluorescens	Boom Monas 1% WP	Root rot, wilt, blast, sheath blight, damping off,
				leaf spot and rhizome rot
		Verticillium lecanii	Boom Vert 1.5% WP	Aphids, thrips, mealy bugs, white flies, jassids,
				Hoppers, scales and all types of mites.
		Beauveria bassiana	Boom Bass 1.15% WP	Root grubs, boll worms, Spodoptera, coffee
				berry borers, pod borers, hoppers and weevils.
			F	
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S.No.	S.No. Name of Company	Organism	Trade name	Target biotic stress
24.	M/s T. Stanes and Company	Verticillium lecanii	BIO CATCH 1.15% WP	whiteflies, jassids, aphids, thrips, mealybugs
	Ltd., Coimbatore, Tamil Nadu	Beauveria bassiana	Bio-Power 1.15% WP	Borers, cutworms, root grubs, leafhoppers. whitefly, aphids, thrips, mealybug
		Metarhizium anisopliae	Bio Magic 1.15% WP	Leaf hoppers, grasshoppers, root grubs, corn
				root worms, bugs, beetles, palm weevils, borers,
)			cutworms, termites
		Paecilomyces lilacinus	Bio Nematon 1.15% WP	Root knot nematodes, burrowing nematodes,
				cyst nematodes, lesion nematodes
		Trichoderma harzianum	Bio Wrap 1% WP	Root-knot nematode, wilt disease of tomato,
				okra crops
		Entomopathogenic	Crown	Root grub
		nematode		
		Streptomyces spp.	Stanomyte 1.5% LF	Mites
25.	M/s Harit Bio Control Lab.,	NPV	Helistop	Helicoverpa armigera
	Yavatmal	Trichoderma viride	Haritderma 1% WP	Damping, wilting, root spots, leaf spots and
				blights
		Verticillium lecanii	Versatile 1% WP	Sucking pests
		Beauveria bassiana	Wow 1.5% WP	Leaffolders
26.	M/s Bannari Amman Sugars	Bacillus licheniformis	LEAF GUARD	Actinopelte Leafspot, Alternaria, Leafspot, leaf
	Ltd., Tamil Nadu			blight, Anthracnose, leaf blotch, Drechslera ink
				spot, Bipolaris Leaf spot, Rhizoctonia blight
		Trichoderma viride	ROOT CARE	Soil borne diseases
		T. harzianum	くと、	
		Pseudomonas fluorescens	Pseudo Care	Crop diseases
		Bacillus subtilis	LEAF CARE	Fungal diseases
			ł	
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Table 2	Table 2: List of biopesticides in pipe-line for registration and licensing in India	n pipe-line for registra	ttion and licens	ing in India		
S.No.	S.No. Entomopathogenic Formulation fungi	Formulation	Shelf life	Trade name	Target pests	Dose
-i	Beauveria bassiana (Bals, Criv.) Vuill. (1912) (Bb-5a)	Oil formulation (1× 12 months at 10 ⁸ cfu/ml) 25-35°C	12 months at 25-35°C	Shatpada Aphid Kill	Chilli and brinjal aphids, <i>Aphis</i> 5 ml/lit. of water at 15 days gossypii Glover, 1877, cabbage interval aphid, <i>Brevicoryne brassicae</i> (Linnaeus, 1758); cowpea aphid, <i>A. craccivora</i> C. L. Koch, 1854	5 ml/lit. of water at 15 days interval
2.	Isaria fumosorosea Wize (1904) (Pfu5)	Talc $(1 \times 10^{8} \text{ cfu/g})$; 12 months at Shatpada oil formulation $25-35^{\circ}\text{C}$ Rugose $(1 \times 10^{8} \text{ otu/m1})$	12 months at 25-35°C	Shatpada Rugose Whitefly Kill	Coconut and oil palm Rugose spiraling whitefly, <i>Aleurodicus</i> rugioperculatus Martin	2-3 foliar spray at 5 ml or 5 g/lit. of water at 15 days interval
Э.	Lecanicillium lecanii R. Zare & W. Gams, 2001 (VI-8)	Oil formulation $(1 \times 10^8 \text{ cfu/ml})$	12 months at 25-35°C	Shatpada Sucking pest Hit	Chilli aphids, A. gossypii, cowpea aphid, A. craccivora	Three foliar sprays at 5 ml/lit. of water at 15 days interval
4	<i>Metarhizium</i> <i>anisopliae</i> (Metchnikoff) Sorokin (1883) Ma 4	Talc $(1 \times 10^{\circ} \text{cfu}/\text{ml})$	12 months at 25-35°C	Shatpada Grubicide	Sugarcane white grub, <i>Holotrichia</i> spp.	Soil application twice in a year during June/July, July/ August at 30 days interval at 2.5 kg mixed with 250 kg FYM/ha
5.	M. anisopliae Ma 35	Talc $(1 \times 10^8 \text{ cfu/g});$ oil formulation $(1 \times 10^8 \text{ cfu/ml})$	12 months at 25-35°C	Shatpada Larvicide	Maize fall armyworm, <i>Spodoptera frugiperda</i> (J. E. Smith)	Three foliar sprays at 5 ml or 5 g/lit. of water at 20,30,40 days after sowing
6.	Trichoderma reesei Simmons, 1977 CST-T-3 CST-T-3	Wettable powder $(1 \times 10^8 \text{ cfu/g})$	12 months at 25-35°C	Fusicont	Fusarium wilt of banana, <i>Fusarium oxysporum</i> f. sp. <i>cubense</i> E. F. Sm., W. C. Snyder & H. N. Hansen (1940) Tropical race 4 and race 1	Soil drenching at 1 lit/, four times on 3 rd , 5 th , 9 th , 12 th month after planting

Dose	Wilt of chickpea, lentil, pea, pigeonpea; damping off seedling mortality in papaya; Target fungi (Rhizoctonia, Sclerotium, Sclerotinia, Husarium, Pythium, Ralstonia, Macrophomina, Bipolaris ,Seed treatment (10 g/kg seed) seed)	<i>Fusarium oxysporum</i> f. sp. Seed treatment (4 g/kg seed); soil application (1 kg/ha with 50 kg FYM) before sowing	Macrophomina phaseolina Seed treatment (10 g/kg (Tassi) Goid. (1947) in seed with jiggery) and soil legumes and oils seed crops application (1 kg/ha with 40 kg FYM	Rice leaf folder, Foliar spray at 2 g/lit. of <i>Cnaphalocrocis medinalis</i> water (Guenée, 1854)	Rice leaf folder, <i>C. medinalis</i> Foliar spray at 2 g/lit. of water	Phytophthora seedling blight, Seed treatment at 1 ml/kg Macrophomina root rot and seeds, foliar spray at 1-2 ml/ Eusarium wilt of safflower and lit. of water gray mold of castor, Alternaria
name Target pests			Mishrit Maru Macrophomina phaseolina sena (Tassi) Goid. (1947) in legumes and oils seed crop:			
e Trade name	hs at Bio-Pulse	s at Maru Sena 1		s at NRRI-BBLF	s at NRRI-Malf	hs at Triguard
Shelf life	() 12 months at 25-35°C	4 months at 55°C	4 months at 550C	8 months at 25-35°C	8 months at 25-35°C	1 18 months at 25-35°C
genic Formulation	Talc $(1 \times 10^7 \text{cfu/g})$	Carrier based $(1 \times 10^{8} \text{ cfu/g})$	Carrier based $(1 \times 10^8 \text{ cfu/g each})$	Talc $(1 \times 10^{9} \mathrm{cfu/g})$	Talc $(1 imes 10^9 \mathrm{cfu/g})$	Liquid suspension concentrate (1 × 10 ⁹ cfu/ml)
S.No. Entomopathogenic fungi	T. harzianum Rifai, (1969) + Bacillus amyloliquefacens Priest et al., 1987	T. harzianum AZNF 5	T. harzianum (ICAR- CAZRI AZNF-5 and Bacillus firmus Bredemann and Werner 1933, ICAR- CAZRI AZ-1	B. bassiana RF6	M. anisopliae TF 19	T. harzianum* Th4d
S.No.	7.	8.	.6	10.	11.	12.

S.No.	S.No. Entomopathogenic fungi	genic Formulation	Shelf life	Trade name	Target pests	Dose
13.	T. harzianum* Th4	Wettable powder (1 × 10° cfu/ml)	18 months at 25-35°C	Triguard Th-P	Phytophthora seedling blight, Macrophomina root rot; Fusarium wilt of safflower and castor, Aspergillus root rot in groundnut	Seed treatment at 10 g/kg of seeds
14.	<i>T. asperellum*</i> Samuels, Lieckf. & Nirenberg 1999 Ta DOR 7316	Wettable powder (1 × 10° cfu/ml)	18 months at 25-35°C	Triguard Ta-P	Phytophthora seedling blight, Macrophomina root rot; Fusarium wilt of safflower and castor	Seed treatment at 10 g/kg seeds
15.	B. bassiana (ITCC 4513)	Liquid suspension concentrate $(1 \times 10^{12} \text{ cfu/m1})$	25-30°C Bb-L	Mycoguard Bb-L	<i>Helicoverpa armigera</i> (Hubner) in pigeonpea	Two to three foliar sprays at 0.3 ml/lit of water at 10 days interval
16.	T. harzianum* ICAR-IIHR Th-2	Wettable powder (2 × 10° cfu/g)	10 months at 25-55°C	Arka Krishi Vriddhi	Meloidogyne incognita (Kofoid & White, 1919), Fusarium oxysporum f. sp. vasinfectum, S. rolfsii, F. solani (Mart.) Sacc. (1881) in brinjal, tomato, carrot, okra	Seed treatment at 20 g/kg seed, nursery bed treatment at 50 g/m ² for transplantable crops, soil application at 5 kg/ha after enrichment in 5 tons FYM before sowing or transplanting
17.	T. viride* ICAR- IIHR Tv-5	Wettable powder (2 ×10° cfu/g)	10 months at 25-35°C	Arka Khishi Veera	M. incognita, Ralstonia solanacearum (Smith 1896), Erwinia carotovora Winslow et al., 1920, Fusarium oxysporum f. sp. vasinfectum, Fusarum oxysporum f. sp. lycopersici, F.solani	Seed treatment at 20 g/kg seed, nursery bed treatment at 50 g/m ² for transplantable crops, soil application at 5 kg/ha after enrichment in 5 tons FYM before sowing or transplanting

uthogenic	S.No. Entomopathogenic Formulation fungi	Shelf life		Target pests	Dose
	(2 × 10 ⁶ cfu/g)	10 months at 25-35°C	Arka Krishi Rakshak	<i>M. incognita</i> in brinjal, tomato, carrot, okra	Seed treatment at 20 g/kg seed, nursery bed treatment at 50 g/m ² for transplantable crops, soil application at 5 kg/ha after enrichment in 5 tons FYM before sowing or transplanting
	Liquid formulation $(5 \times 10^{11} \text{ ofu/ml})$	3 months at 25-35°C	Manjari Vineguard	Grapes powdery mildew	Soil drenching at 2 ml/lit. of water
	Liquid (5 × 10 ⁸ cfu/ ml)	3 months at 25-35°C	Manjari Rakshak	Grapes powdery mildew	Foliar spray at 2 ml/lit. of water
	Wettable powder (10 ⁸ cfu/g)	25 months at Pusa 5 SD 25°C		<i>F. oxysporum</i> f. sp. <i>ciceris, S.</i> rolfsii, <i>S. sclerotiorum</i> (Lib.) de Bary (1884) in chickpea; <i>R. solani</i> J.G. Kühn 1858, <i>R. bataticola</i> (Taub.)Butl. in chickpea and mugbean, <i>F.</i> oxysporum f. sp. <i>lycopersici</i> in tomato, <i>P. ultimum, R. solani</i> in fresh bean, major seed borne fungal pathogens	Seed treatment at 4 g/kg of seeds
	Wettable powder (2 × 10 ⁶ cfu/ml)	10 months at 25-35°C	ARKA Krishi Kawach	10 months at ARKA Krishi <i>Meloidogyne incognita</i> in 25-35°C Kawach brinjal, tomato, carrot, okra	Seed treatment at 20 g/kg seed, nursery bed treatment at 50 g/m ² for transplantable crops, soil application at 5 kg/ha after enrichment in 5 tonns FYM before transplanting or sowing

S.No.	S.No. Entomopathogenic fungi	genic Formulation	Shelf life	Trade name	Target pests	Dose
23.	Bacillus thurigiensis var. kurstaki	Liquid $(1 \times 10^8 \text{ cfu})$ ml)	12 months at 25-35°C	Shatpada Armour	Maize fall armyworm	Two to three foliar sprays at 10 ml/lit of water at 25, 35, 45 days after sowing
24.	<i>B. thur</i> ingiensis var. kurstaki	Liquid $(1 \times 10^{\circ} \text{ cfu}/12 \text{ months at Shatpada}$ ml) 25-35°C Terminatt	12 months at 25-35°C	Shatpada Terminator	H. armigera, Plutella xylostella, Chilo partellus, Cnaphalocrocis medinalis, Leucinodes orbonalis, Amsacta albistriga	Two to three foliar sprays at 20 ml/lit of water at pre- flowering and post flowering stages
25.	Pseudomonas fluroescens	Tarc based (1 × 10 ⁸ cfu/ml)	12 months at 25-35°C	Shatpada All Rounder	<i>Thrips</i> spp., in capsicum and Fusarium will of red gram	Foliar application at 20 g/ lit of water at 20,30,40,50 days after transplanting for the management of thrips in capsicum; soil application in the root zone during 25,40,55 days after sowing at 2.5 kg/ha for management of red gram wilt; mix 2.5 kg of formulation in 250 kg farmyard manure and apply
26.	B. albus	Talc based (1 × 10 ⁸ cfu/ml)	12 months at Sharpada 25-35°C Master Blaster	Shafpada Master Blaster	S. frugiperda, Tuta absoluta, Fusarium oxysporum f. sp. cucumerinum	Foliar application at 20 g/ lit of water at 20,30,40,50 days after sowing for fall armyworm and tomato pin worm; soil application in the root zone during 25,40,55 days after sowing at 2.5 kg/ha for management of cucumber wilt; mix 2.5 kg of formulation in 250 kg farmyard manure and apply

S.No.	S.No. Entomopathogenic fungi	genic Formulation	Shelf life	Trade name	Target pests	Dose
27.	P. fluorescens	Talc based $(1 \times 10^8$ cfu/ml)	12 months at 25-35°C	Eco-Pesticide	Spot blotch of wheat, sheath blight of rice and wilt of tomato and chickpea	Seed treatment (10 g/kg seed)
28.	B. firmus	Carrier based (1 × 10 ⁸ cfu/ml)	6 months at 25-35°C	Maru sena 3	Macrophomina phaseolina in legumes and oil seed	Seed treatment (30 g/kg seed with jiggery solution) before sowing
29.	<i>B. thuringiensis</i> var. kurstaki	Liquid suspension concentrate (1 × 10 ⁿ cfu/m1)	24 months at Bioguard 25-35°C Bt-L	Bioguard Bt-L	Spodoptera litura in soybean	Two foliar sprays at 3 ml/lit of water at 10 days interval.
30.	P. fluorescens	wder (2	10 months at 25-35°C	ARKA Krishi Samarakshak	ARKA Krishi Meloidogyne incognita, Samarakshak Ralstonia solanacearum, Erwinia carotovora, Fusarium oxysporum f. sp. vasinfectum, Fusarium solani in brinjal, tomato, carrot, okra	Seed treatment at 20 g/kg seed, nursery bed treatment at 50 g/m ² for transplantable crops, soil application at 5 kg/ha after enrichment in 5 tonns FYM before transplanting or sowing
31.	P. fluorescens	Liquid (2 × 10 ⁸ cfu/ ml), carrier based (2 × 10 ⁸ cfu/ml)	T0-12 months at 25-35°C	ARKA krishi All Rounder and ARKA Plant Growth Booster	Meloidogyne incognita, Fusarium oxysporum f. sp. vasinfectum, Fusarium solani in several vegetable and fruit crops	Seed treatment at 20 g or 20 ml/kg seed, substrate treatmen with 10 ml or 10 g/kg of cocopeat, soil application at 5 kg or 5 li/ ha after enrichment in 5 tons FYM before transplanting or sowing
*Licens	*Licensed to private companies Source: Saxena et al. (2021)	s Source: Saxena et al	. (2021)		4	

Conclusions

Government regulations and the detrimental effects of chemical pesticides force a shift to alternate plant protection measures. As a result, microbial biopesticide, one of the environmentally friendly techniques, has become more significant in agriculture both globally and in India. Although a few factors, such as quality control and the identification of effective organisms, predispose the market and widespread use of biopesticide, central and state government initiatives, such as the establishment of assisted and non-aided biocontrol laboratories and intense R&D activities, support the growth of biopesticide steadily.

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Links

http://ppqs.gov.in/divisions/cib-rc/biopesticide-registrant

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Nano-Biopesticides for Management of Insect Pests of Crops

Abstract

A type of pesticide known as a "biopesticide" is derived from natural sources such as plants, animals, minerals, and microbes. The employment of cuttingedge scientific tools will be extremely beneficial for enhancing the efficacy, greater applicability and adaptability, and storability of biopesticides. One such rapidly rising scientific discipline, nanotechnology, has considerable potential applications in agriculture, including the creation of carriers for pesticides, plant growth regulators, biofertilizers, nano-sensors, insecticides, food packaging materials, and gene transfer, among other things. Nanoparticles fall under the category of ultrafine particles and range in size from 1 to 100 nm. These nano-particles have special uses in various industries, including the production of nano-biopesticides. They differ from their bulk material due to their small size, shape, reactivity, and increased surface area to volume ratio. Without sacrificing on safety and health risks, nano-biopesticides offer higher accuracy in reaching their potential efficacy against target pests. This chapter covers several nano-biopesticides, their history, their use as biopesticides, modes of action, and their effectiveness against the pests they are intended to control.

Keywords: Nano-particle, Nano-biopesticide, AgNP, SiNP, Botanicals, Semiochemical

Introduction

Biopesticides are pesticides that are produced naturally by plants, animals, microbes, and other minerals. These represent less of a harm to humans and the environment than chemical insecticides. Nanotechnology is a rapidly developing scientific topic that has numerous uses in numerous industries, including agriculture. The transport of plant hormones, seed germination, water management, transfer of target genes, nano barcoding, nano sensors,

and controlled release of agrichemicals are now being investigated as applications for nanotechnology in agriculture (Worrall et al. 2018). For their diverse applications in numerous industries, many scientists have modified nanoparticles for their size, shape, porosity, and/or surface tension, among other properties.

Nano-particles are a subcategory of ultrafine particles that range in size from 1 to 100 nm. These nano-particles have special uses in numerous industries due to their distinctive characteristics, which include small size, shape, reactivity, and a greater surface area to volume ratio. These nano-particles have numerous and varied uses in agriculture, including as pesticide transporters, plant growth regulators, biofertilizers, nano-sensors, insecticides, food packaging materials, and gene transfer agents, among others. Nano-particles give agricultural insect pest management techniques fresh dimensions. For targeted, controlled distribution of the active component in a biopesticide, nano-particles may be utilised. Compared to conventional biopesticides, nano-biopesticides provide a number of benefits.

Nano-Biopesticides

To increase their effectiveness, expand their potential applications, and solve numerous drawbacks, the various biopesticides, including microbial biopesticides, biochemical biopesticides, and plant-incorporated protectants, can be produced as nano-based biopesticides. In Table 1, a few of the nano-biopesticides were listed.

Table 1: Nanomaterials appli	applied to bio-pesticide and their functions (An et al. 2022)	leir functions (An et al. 2	2022)	
Bio-pesticide category	Active ingredients Material(s) (AIs)	Material(s)	Function	Reference
Microbial pesticides (Bacterial pesticide)	Bacillus thuringiensis (Bt)	Nano-tubular sodium titanate	Bacillus thuringiensis Nano-tubular sodium Effective in controlling cotton leafworm (Bt) titanate	Zaki et al. (2017)
Microbial pesticides (fungal pesti- cides)	Beauveria bassiana (Bals.)	Silica nano-particles and carbon fibers	<i>bassiana</i> Silica nano-particles Improving mortality to larvae of potato Hersanti et al. (2020) and carbon fibers	Hersanti et al. (2020)
Plant-derived pesticides	Essential oils	ted		Gahukar et al. (2020)
Agricultural antibiotics	Avennectin	Poly (ethylene glycol)-carboxyme- thyl cellulose (PEG- CMC)	Poly (ethylene Improved the anti-UV ability & increased glycol)-carboxyme- the biocompatibility of the Avermectin thyl cellulose (PEG-CMC)	Zhu et al. (2020)
	Validamycin and thifluzamide	and Polylactic acid	The nano-particles prepared by compounding Validamycin with chemical pesticides showed better control of rice sheath blight, which was 4.2 times more effective than the control	Cui et al. (2020)
Biochemical pesticides	Gibberellic acid (GA)	Layered hydroxides (L1	double Promoted plant growth	Hafez et al. (2018)
Plant immunity elicitor- inducing antibacterial agents	Chitosan & Zinc oxide	C h i t o s a n encapsulated zinc oxide nanocomposite	an As an efficient biocompatible elicitor to Asgari-Targhi zine improve agronomic traits of crops (2021)	Asgari-Targhi et al. (2021)
			A	

Botanical Based Nano-Biopesticides

Some of the nano-biopesticides are based on the botanical biopesticides such as essential oils, neem oil, neem powder etc., were found effective in manging many insect pests. The antifeedant and larvicidal activity of nano-PONNEEM[®]-encapsulated biopesticide tripolyphosphate cross-linked chitosan nanocarriers was reported against Helicoverpa armigera (Paulraj et al. 2017). Adel et al. (2018) tested the nano-emulsion of Mentha piperita essential oil against stored grain pest, Tribolium castaneum and reported that it was very much effective in controlling the pest with highest percent mortality in comparison to essential oil without nano-emulsion in wheat. Further, they also noticed that nano-emulsion significantly enhanced the germination percentage of wheat seeds. Similarly, Yang et al. (2009) reported the highest mortality (up to 80%) of Tribolium castaneum when treated with nano-biopesticides loaded with garlic essential oils. The list of nano-carriers used for biopesticides/active ingredients that target crop pests is provided in Table 2.

In an exclusive study, Palermo et al. (2021) tested the nano formulations prepared from eight commercial essential oils (*Pimpinella anisum*, *Artemisia vulgaris*, *Foenicum vulgare*, *Allium sativum*, *Lavandula angustifolia*, *Mentha piperita*, *Rosmarinus officinalis*, and *Salvia officinalis*) for their acute toxicity and repellence against confused flour beetle, *Tribolium confusum* and reported that all the nano-emulsions were found best repellent over time. The highest acute toxicity was noticed in garlic nano-emulsions with maximum mortality. Bidyarani and Kumar (2019) encapsulated the rotenone, a naturally occurring pesticide in the roots of Fabaceae plants, in zein nanoparticles by antisolvent precipitation method. They evaluated nano-encapsulated rotenone against plant pathogens *Pseudomonas syringae* and *Fusarium oxysporum* and reported excellent antimicrobial activity.

Biopesticide	Nanoparticle	Crop	Biopesticide Target insect pest	Reference
Garlic essential oil	Polyethylene glycol	Rice (harvested)	Rice (harvested) Red flour beetle (<i>Triboleum castaneum</i>)	Yang et al. (2009)
Azadirachtin	Chitosan	1	Tobacco cutworm (Spodoptera litura) culture ovarian cell lines SI-1	Lu et al. (2013)
Azadirachtin	Chitosan		1	Feng et al. (2012)
α-pinene and Linalool	Silica	Castor	Tobacco cutworm (S. litura), Castor semi-looper (Achaea	Rani et al. (2014)
			Janata)	
Abamectin	Porous silica			Wang et al. (2014)
Anacardic acid	Layered double hydroxide	Mustard	Tobacco cutworm (S. litura)	Nguyen et al. (2015)
Avermectin	Polydopamine	Cotton and corn	1	Jia et al. (2014)
Avermectin	Polydopamine	1	1	Sheng et al. (2015)
Suaeda maritima-based herbal coils	Silver	Cotton	Tobacco cutworm (S. litura)	Suresh et al. (2018)
PONNEEM	Chitosan	Cotton	Cotton bollworm (Helicoverpa armigera)	Paulraj et al. (2017)
Avermectin	Polydopamine	Cucumber and broccoli	Aphids	Liang et al. (2018)
Avermectin	Castor oil-based polyurethane	Corn leaves		Zhang et al. (2018)
Azadirachtin	Zinc oxide and chitosan	Groundnut	Bruchid (Callosobruchus serratus)	Jenne at al. (2018)
Carvacrol Linalool	Chitosan	I	Mite (Tetranychus urticae)	Campos et al. (2018)
Geraniol	Chitosan/Gum Arabic	ı	Whitefly (Bemicia tubaci)	De Oliveira et al. (2018)

Biopesticide	Nanoparticle	Crop	Target insect pest	Reference
Satureja hortensis essential oil	Chitosan/TPP	-	Mite (T. urticae)	Ahmadi et al. (2018)
Geraniol and R-citronellal essential oils	Zein	1	Mite (T. urticae)	Oliveira et al. (2018)
Nicotine	Chitosan/TPP	1	House fly (Musca domestica)	Yang et al. (2018)
Carvacrol Linalool	Chitosan	Bean	Corn earworm (<i>H. armigera</i>), mites (<i>T. urticae</i>)	Campos et al. (2018)
Avermectin	porous ho llow silica nanoparticles	Brassica oleracea	Plutella xylostella larvae	Kaziem et al. (2018)
	P.Y.	Gy Gy	A.	

Nano-formulations of Semiochemicals

One such naturally occurring semiochemical that is commonly employed to control insect pests is pheromones. They are somewhat unstable in nature as a result of isomerization, photooxidation, autooxidation, and volatility, among other processes (Deepa et al. 2013). In order to increase the effectiveness of pheromones in real-world settings, slow and controlled release formulations are crucial. Nano-formulations are the greatest options for delayed and controlled release of pheromones. By immobilising the pheromone into the nano gel, Deepa Bhagat et al. (2013) created a nano gel of methyl eugenol, a pheromone used to manage the fruit fly pest, *Bactrocera dorsalis*, and tested the gel's effectiveness in the field. Additionally, they discovered that pheromones based on nanogels were stable at room temperature and exhibited a reduced rate of evaporation, making handling and shipping simpler. When immobilised into nano-gels, methyl eugenol's shelf life was increased, and fruit fly pest trap catches were substantially higher than with methyl eugenol alone.

Abd El-Wahab et al. (2020) investigated the catchability of the red palm weevil, *Rhynchophorus ferrugineus*, and found that aggregation nano-gel pheromone traps attracted considerably more adult beetles than conventional pheromone traps in two seasons (55.33 and 46.33 adults/trap). White grubs (*Holotrichia consanguinea*) will be caught in ground nuts using a nano-gel formulation of the aggregation pheromone, methoxy benzene, created by Deep Bhagat et al. (2020). By immobilising the aggregation pheromone in a matrix and creating a viscoelastic semi-solid mass, a nanogel formulation was created. Additionally, they examined its effectiveness and discovered that the nanogel trap may capture up to 17.5 adult beetles daily.

Nano-Biopesticides Derived from Plants

Number of nano-biopesticides containing different nanoparticles were reported by many researchers. Plants serve as excellent sources for various nanoparticles which can be used as nano-biopesticides against various insect pests. The list of plant-derived nano-biopesticides, their application and nanoparticles present were presented in Table 3.



1able 3. Plant derived Nano-biopesticides and their efficacy against various insect pests (Krishnamurthy et al. 2020)	o-piopesuiciaes and me	II VIIIVUVY UBUILLY			
Plantname	Common name	NPs Present	Part of plant	Application	Reference
Acorus calamus	Sweet flag	Au, Ag	Root	Insecticide	Ganesan et al. (2015)
Agave americana	Sentry plant	Ag	Leaf	Whiteants	Ahmad et al. (2016)
Ageratum conyzoides	Chick weed	Ag	Leaf	Liceinhair	Wardani et al. (2019)
Albizialebbeck	Lebbeck	Ag, Ni, Fe	Seed, leaf, bark, root	Insecticide	Umar et al. (2010)
Albiziaprocera	White siris	Zn, Ag, Cu	Leaf	Insecticide	Jayakumarai et al. (2010)
A loese cundifior a	New castle in chicken	Cd, Ag	Sap	Insecticide	Tippayawat et al. (2016)
Alysicarpus bupleurifolius	Alyce clover	Ag, P	Whole plant	Bed bugs, whiteants	Kasithevar et al. (2017)
Anacardium occidentale	Kaaju	Au, Ag, Cu, Pt	Shell oil	White ants, insecticide	Begum et al. (2018)
Anamirtacocculus	Indian berry	Zn, Ag, Au	Fruit	Insecticide	
Annona reticulate	Wild-sweetsop	Ag	Leaf, seed, bark	Body lice, insecticide	Parthiban et al. (2019)
Annona squamosa	Sugar apple	Ag, Fe	Seed, stem, bark, leaf, fruit	Body lice, insecticide	Vivek et al. (2010)
Arisaemator tuosum	Whipcord cobra lily	Ag, Au, Zn	Tubers	Insecticide	Kumar et al. (2018)
Aristolochia bracteolate	Worm killer	Ag	Juice	Insecticide	Doss (2015)
Artemisia japonica	Mug wort	Ag, Au, Fe	Whole plant	Insecticide, housefly repellent	Yu et al. (2019)
Artemisia nilagirica	Worm wood	Ag	Leaf	Insect repellent, prevent moths	Vijayakumar et al. (2013)
Azadirachta indica	Neem tree	Ag, Cu	Whole plant	Insecticide, rice and wheat weevil	Ahmad et al. (2016)
Bambusa arundinacea	Bambusa bambos	Ag	Shoot	Kill mosquito larvae	Kataria et al. (2017)
Biden spilosa	Black jack	Ag, Au	Leaves	Aphids	Kyomuhimbo et al. (2019)

Plantname	Common name	NPs Present	Part of plant	Application	Reference
Blumeae riantha	Buradi	Zn	Whole plant	Mosquito repellent	Benelli et al. (2017)
Boswellia serrata	Indian frankincense	Ag	Gum	Fumigation repel houseflies, mosquitoes	Kora et al. (2012)
Brassica campestris	Field mustard	Ag, Zn	Seed oil	Beetles	Khan et al. (2018)
Butea monosperma	Sacred tree	Ag, Au, Zn	Seed flower extract	White ants	Das et al. (2018)
Calotropis procera	Rubber bush	Ag, Zn, Ni, Fe	Leaf	Larvicidal	Gawade at al. (2017)
Canna bissativa	Kumbhi	Au, Ag	Whole plant leaf	Bugs & pests	Singh et al. (2018)
Capsicum annuum	Hot pepper	Cu, Ag, Au	Fruit, leaf	Thrips, aphids, white flies	Yuan et al. (2017)
Careya arborea	Karanda	Ag	Root, bark and leaf	Several	Nair et al. (2015)
Carica papaya	Papaya	Ag, Zn	Leaves, seeds	Several	Rathnasamy et al. (2017)
ta	Hemp	Ag	Root and bark	Vet worms in wounds	Joshi et al. (2018)
Cassia hirsute	Cassia	Ag, Zn	Bark	Insecticide	Adesuji et al. (2016)
Cassytha filiformis	Love vine	Ag, Cu, Mg	Whole plant	Insecticide	Nasrollahzadeh et al. (2018)
Catunaregam spinosa	Mountain pomegranate	Sn, Zn, Ni	Fruit	Insecticide	Haritha et al. (2016)
Cinnamomum camphora	Camphor tree	Ag, Au, Pt, Pd	Bark powder	Protect clothes against insects	Huang et al. (2006)
Citrus limon	Lemon	Ag, Au	Drieddeaf	Wheat weevil, flour beetle	Sujitha et al. (2013)
Commiphora wightii	Indian bdellium tree	Ag	Resin	Mosquito repellent	Sarkar (2017)
Cordia latifolia	Sebestan plum	Ag	Leaves	Maize weevil, butterfly	Ioset et al. (2000)

Plantname	Common name	NPs Present	Part of plant	Application	Reference
Coryphaum braculifera	Talipot palm	Cu, Ag	Young fruit	Insect repellent	Abdel-Wahab et al. (2019)
Croton roxburghii	Croton	Ag	Seed	Insecticide	Panda et al. (2010)
Cucumis melo	Muskmelon	Ag	Leaf	lice	Haryani et al. (2018)
Cucumis sativus	Cucumber	Ag, Cu, Zn	Rhizome	lice and insects	Zhao et al. (2014)
Curcuma longa	Turmerio	Ag, Zn	Rhizome	Drive away ants	Shameli et al. (2012)
Cuscuta reflexa	Amarbel	Cu, Ag	Whole plant	lice	Naghdi et al. (2018)
Cymbopogan nardus	Citronella grass	Ag	Whole plant	Mosquito repellent	Kamarudin et al. (2019)
Derriss candens	Gewel vine	Ag	leaf, bark	Insecticide	Firdhouse et al. (2013)
Derris trifoliate	Karanjvel	Ag	Bark	Insecticide	Kumar et al. (2017)
Desmodium triflorum	Tick clover	Ag, Cu, Au	Whole plant	Insecticide	Ahmad et al. (2011)
Dioscorea hispida	Asia ticbitteryam	Ag	Bark	Insecticide	Ashri et al. (2014)
Duranta erecta	Brazili ansky flower	Ag, Zn	Whole plant	Insecticide	Ravindran et al. (2016)
Euphorbia antiquorum	Spurge	Ag	Milkyjuice	Maggot sin wound	Rajkuberan et al. (2017)
Euphorbia dracunculoides	Dragon spurge	Ag	Latex	Kill slice	Annamalai et al. (2013)
Euphorbia thymifolia	Thyme leaf	Pd	Whole plant	Flies, mosquitoes	Nasrollahzadeh et al. (2016)
Fioria vitifolia	Grape leaved mallow	Ag	Root, bark	Kill slice	Ghosh et al. (2015)

Plantname	Common name	NPs Present	Part of plant	Application	Reference
Glorio sasuperb	Flame lily	Ag, Au, Ce, Cu ,Pt, Pd	Leaf	Liceinthe hair	Ashokkumar et al. (2013)
Haldina cordifolia	Kadam	Ag	Bark	Insecticide	Khan et al. (2019)
Hard wickiabinate	Anjan	Zn	Wood	Insecticide	Gunaselvi et al. (2010)
Harpullia arborea	Tulip wood	Zn, Sn	Bark	Leech repellent	Mohan et al. (2018)
Holarrhena pubescens	Indrajao	Ag	Flower, seed	Insecticides	Venkata Subbaiah et al. (2013)
Hyptissua veolens	American mint	Ag, Cu	Twig	Repel bed bugs	Elumalai et al. (2017)
Kalanchoe integra	Never die	Au, Cu	Leaf	Insecticide	Patel et al. (2019)
Lagenandra ovate	Malayans word	Ag	Whole plant	Insecticide	Bokaeian et al. (2015)
Lavandula bipinnata	Lavender	Zh	Whole plant	Insect repellent	Shaikh et al. (2014)
Lavendulalawii	Lavender	Ag	Wholeplant	Insect repellent	Kulkarni et al. (2013)
Leonotis nepetifolia	Klip dagga	Ag	Leaf	House fly repellent	Al-Sheddi et al. (2018)
Leucas aspera	Thumbai	Ag, Ce, Cu	Whole plant	Insecticide	Malleshappa et al. (2015)
Lippia javanica	Fever tea	Ag	Leaf	Insecticide	Kumar et al. (2016)
Madhu calongifolia	Ilippai	Ag, Cu, Au	Seed, seed, oil, cake	Worm killer, insect repellent	Sharma et al. (2019)
Melaleuca leucadendron	Caju puttree	Au, Ag	Oil	Mosquito repellent	Souza et al. (2017)
Melia azadarach	Chinaberry	Ag	Fruit, seed	Insecticide	Anbu et al. (2016)

Plantname	Common name	NPs Present	Part of plant	Application	Reference
Melia volkensii	Melia	Ag	Fruit pulp	Termites	Kamau et al. (2016)
Millettia extensa	Benth	Ag	Root	Insecticide	Panda et al. (20160
Mimosa pudica	Shame plant	Ag, Zn, Fe, Au, Leaf Cu	Leaf	Veterinary wound maggots	Fatimah et al. (2016)
Mundulea sericea	Cork bush	Zn	Seed, root, bark	Insecticide	Chaithong et al. (2006)
Nigella sativa	Black seed	Ag, Zn	Seed	Pesticide	Amooaghaie et al. (2015)
Ocimum americanum	Hoary basil	Ag	Whole plant	Insecticide	Anuradha et al. (2014)
Ocimum gratissimum	Ramtulsi	Ag, Au	Whole plant	Insect repellent	Das et al. (2017)
Ocimum kilimandscharicum	Camphor Basil	Ag	Leaves, flower	Mosquito, fleas	Selvarani et al. (2016)
Ocimum tenuiflorum	Tulsi	Ag	Whole plant	Insect repellent	Singhal et al. (2011)
Peganum harmala	Wild rue	Ag, Zn	Root	Mosquito repellent	Fazlzadeh et al. (2017)
Psidia punctulata	Mpepe	Ti, Cu	Leaves	Lice, fleas, mites	Zinjarde et al. (2011)
Pongamia pinnata	Indian beech	Ag	Seed, root, seed oil	Repellent, insecticide	Paul et al. (2018)
Riccinus communis	Castor bean	Ag, Au	Seed oil	Flies repellent, rice moth, rice weevil	Ojha et al. (2017)
Ruta graveolens	Herbof grace	Zn, Ag	Whole plant	Insects	Lingaraju et al. (2016)
Sarcostm maviminale	Caustic vine	Ag	Leaf	White ants	Kannan et al. (2018)
Securidacalon gepedunculata	Violet tree	Ag	Whole plant	Stored grain pest	Ojewole et al. (2008)

Plantname	Common name	NPs Present	Part of plant	Application	Reference
Senna didymobotrya	Popcorn senna	Ag	Leaves	Nematodes	Vijayakumari et al. (2018)
Solanum nigrum	Sodom apple	Ag	Fruit, leaf	Insecticides	Venkat Kumar et al. (2017)
Strychno snuxvomica	Poison nut	Zn, Au, Ag	Fruit, seed	White ants	Steffy et al. (2018)
Strychnos spinosa	Monkey orange	Ag	Whole plant	Insecticides	Isa et al. (2014)
Symplytum officinale.	Common comfrey	Ag	Leaf, root	Insecticides	Singh et al. (2018)
Tagetes minuta	Wild marigold	Ag	Leaf, flower	Insecticides	Shahzadi et al. (2015)
Tanacetum cinerariifolium	Pyrethrum	Ag, Au	Flower	Bees and insects	Kitherian et al. (2016)
Tephrosia purpurea	Fish poison	Ag, Au	Wood, roots	Cotton and woollen cloth moths	Srikar et al. (2016)
Tithonia diversifolia	Marigold	Ag	Leaf, flower	Insecticides	Tran et al. (2013)
Trachylo biumammi	Ajwain	Ag, Zn	Seed	Mosquito repellent	Chouhan et al. (2017)
Trigonella foenumgraecum	Fenugreek	Au, Ag	Seed	Insect repellent	Aswathy et al. (2012)
Vernonia amygdalina	Bitter leaf	Ag	Leaf	Insecticides	Widyaningtyas et al. (2019)
Vernonia anthelminticum	Kalijiri	Zn, Au	Seed	Fleas	Karthikeyan et al. (2008)
Vitex negundo	Chaste tree	Ag, Zn	Leaf	Insect repellent	Prabhu et al. (2013)
Vitex trifolia	Arabianlilac	Ag, Zn, Au	Leaf	Insect repellent	Elumalai et al. (2015)

Nanoparticles as Nano-Biopesticides

With the growing advancement of science and awareness about environment and pesticide free food materials, eco-friendly measures are employed for insect pest management practices. Nanotechnology offers one such solution to eco-friendly control of insect pests in the form of nano-particles. Lot of evidence proved that many nano-particles offers as insecticide/acaricide against a range of insect pests across the genera. Some of the nano-particles used as biopesticides are listed in Table 4.

Tested nanomaterial (dose or concentration)	Insect target	Morphological damages and/ or mode of action	References
Various green and microbial synthesized Ag, Au, and ZnO nanoparticles	Aedes aegypti, Anopheles stephensi	Midgut, epithelial cell, and cortex damages, with accumulation of nanoparticles in the midgut. Shrinkage in the abdominal region, thorax shape changes, midgut damages, loss of lateral hairs, anal gills and brushes	Banumathi et al. (2017); Kalimuthu et al. (2017), Sundararajan and Kumari (2017), Abinaya et al. (2018), Ishwarya et al. (2018)
Ag nanoparticles preparedusing Cassia fistula extract($LC_{50} =$ 3.6 and 1.7 mg/l, respectively)	Aedes albopictus, Culex pipiens pallens	4^{th} instar larvae showed a decrease of total protein levels; nano Ag also reduced acetylcholinesterase and α - and β -carboxylesterase activities	Fouad et al. (2018)
Ag nanoparticles fabricated using salicylic acid and 3,5-initrosalicylic acid(1–12 ppm)	Aedes albopictus	4 th instar larvae showed a decrease of total proteins, esterase, acetylcholine esterase, and phosphatase enzymes.	Ga'al et al. (2018)
Ag nanoparticles (0.2, 0.5, and 1 mg/l)	Chironomus riparius	GST genes up- or down regulated, according to tested concentration and duration of exposure, highest mRNA expression was in delta3,Sigma4 and Epsilon1 GST class	Nair and Choi (2011)

 Table 4: Mode of action of nano-particles as biopesticides (Benelli, 2018)

Tested nanomaterial	Insect target	Morphological damages and/	References
(dose or		or mode of action	
concentration)			
Ag nanoparticles (up	Chironomus riparius	Down regulation of the	Nair et al.
to 4 mg/l)		ribosomal protein gene	(2011)
		(CrL15) regulating ribosomal	
		assembly, thus protein	
		synthesis. Up regulation	
		of gonadotrophin releasing	
		hormone gene (CrGnRH1)	
		and Balbiani ring protein	
		gene (CrBR2.2), which can	
		indicate the activation of	
		gonadotrophin releasing	
		hormone mediated signal	
		transduction pathways and	
		reproductive failure.	
Ag nanoparticles	Chironomus riparius	Expression of the ecdysone	Nair and Choi
(0.2, 0.5, and 1 mg/l)		receptor gene was up or	(2012)
		Down regulated according to	
		the exposure time	
Ag nanoparticles	Chironomus riparius	Up regulation of Mn	Nair et al.
(0.2, 0.5, and 1 mg/l)		superoxide dismutase;	(2013)
		transcript levels of catalase,	
		phospholipidhydroperoxide	
		glutathione peroxidase 1	
		and thioredoxin reductase	
		1 upregulated. Boosted expression of Delta-3,	
		sigma-4, and epsilon-1	
		classes of glutathione	
		S-transferases	
Ag nanoparticles (<	Drosophila	Loss of melanin cuticular	Armstrong et
50 mg/l)	melanogaster	pigments, reduced vertical	al. (2013)
55 1115/17	metunoguster	flight ability, reduced activity	ui. (2013)
		of Cu-dependent enzymes	
		(tyrosinase and Cu-Zn	
		superoxide dismutase);	
		nanoAg coupled with	
	*	membrane-bound Cu	
		transporter proteins lead	
(ر)		sequestration of Cu,	
		mimicking Cu starvation	
	1		1

Tested nanomaterial	Insect target	Morphological damages and/	References
(dose or	insect target	or mode of action	references
concentration)			
Ag nanoparticles (25–50 µg/ml nanoAg 4.7 nm and250–1000 µg/ml nanoAg 42 nm)	Drosophila melanogaster	Lack of mutagenic and recombinogenic activity. However, both nano-Ag 4.7 and 42 nm evoked pigmentation defects and locomotor ability decrease in adult flies	Ávalos et al. (2015)
Ag nanoparticles	Drosophila	Accumulation of reactive	Mao et al.
(10–50 µg/ml)	melanogaster	oxygen species (ROS) in the fly tissues leading to ROS- mediated apoptosis, DNA damage, and autophagy; activation of the Nrf2- dependent antioxidant pathway	(2018)
Ag and TiO ₂	Drosophila	Progeny loss and a decrease	Philbrook et
nanoparticles $(0.005$ to $0.05\%)$	melanogaster	in developmental success	al. (2011)
Ag nanoparticles	Spodoptera litura and	Nano-induced oxidative	Yasur and
(500 to4000 mg/l)	Achaea janata	stress in moth larval guts,	Usha-Rani
		with enhanced antioxidant	(2015)
Ag nanoparticles synthesized using the <i>Punica</i> granatumpeel extract ($LC_{50} = 19.21$ µg/larva)	Spodoptera litura	enzyme levels Reduction of amylase, protease, lipase, and invertase activities; gut microflora and the extracellular enzyme production decreased, along with weight, pH, and total heterotrophic bacterial population	Bharani and Namasivayam (2017)
Nanostructured Al ₂ O ₃ (60–500 ppm)	Sitophilus oryzae	Bind to the beetle cuticle due to triboelectric forces, sorbing its wax layer by surface area phenomena, resulting in insect dehydration	Stadler et al. (2017)
Au nanoparticles $(87.44 \ \mu g/gin the diet)$	Blattella germanica	Disrupted reproduction and development	Small et al. (2016)
Au nanoparticles fabricated using latex of <i>Jatropha</i> <i>curcas</i> (500–1000 µl)	Aedes aegypti, beetles, and Mealy bugs	Triggered trypsin inhibition	Patil et al. (2016)

Tested nanomaterial (dose or concentration)	Insect target	Morphological damages and/ or mode of action	References
Carbon black and multiwalled nanotubes (3.3 and 3.1 mg, respectively)	Drosophila melanogaster	Strong adherence of the nanomaterials to the fly body parts, leading to impaired motor functions and insect mortality	Liu et al. (2009)
Graphene oxide nanoparticles (0.1 µl per 100 mg ofinsect's body weight)	Acheta domesticus	Increased enzymatic activity of catalase and glutathione peroxidases, as well as heat shock protein (HSP 70) and total antioxidant capacity levels	Dziewięcka et al. (2016)
Carbon-dot-Ag nanohybrid(LC_{50} values from 0.30 to0.76 ppm)	Anopheles stephensi, Culex quinquefasciatus	Deformation of larval body, presence of Ag(2.93%) in the tissues of treated mosquitoes, cuticle, and cellular organization damages	Sultana et al. (2018)
Polystyrene nanoparticles (20–500 µg/ml)	Insect cells (BACULOSOMES®)	Inhibited the enzymatic activity of CYP450 isoenzymes in BACULOSOMES®	Fröhlich et al. (2010)
SiO ₂ nanoparticles	Different species, with special reference to stored product pests	Physio-sorbed by the insect cuticular lipids, causing major damages, followed by the insect's death	Barik et al. (2008), Debnath et al.(2011), Athanassiou et al. (2018)
SiO ₂ nanoparticles (LudoxTMA) (\geq 34 mg/l)	Bombus terrestris	Midgut epithelial injury in intoxicated workers	Mommaerts et al. (2012)
TiO ₂ nanoparticles (5 μg/ml)	Bombyx mori	Upregulation of pi3k and P70S6K [rapamycin (TOR) signalling pathway]; 4 cytochrome P450genes (20-hydroxyecdysone biosynthesis),were up-regulated; 20-hydroxyecdysone biosynthesiswas stimulated; reduced development and	Li et al. (2014)
		moulting duration were noted	

Ag Nanoparticles (AgNPs)

Many researchers studied the green synthesis of AgNPs and their efficacy in controlling many agricultural and household insect pests. Devi et al. (2014) synthesized AgNPs from leaf aqueous extract of Euphorbia hirta L. (Malpighiales: Euphorbiaceae) and tested against larvae and pupae of cotton bollworm, *H. armigera* and observed the susceptibility of all stages. Similarly, Marimuthu et al. (2011) synthesized AgNPs from leaf aqueous extract of Mimosa pudica L. (Fabales: Fabaceae) and tested on larvae of mosquitos C. guinguefasciatus and A. subpictus and larvae of the tick Rhipicephalus microplus Canestrini (Acari: Ixodidae) and found their susceptibility to AgNPs. Kantrao et al. (2017) synthesized AgNPs from leaf extracts of the Peepal tree, Ficus religiosa and the banyan tree, Ficus benghalensis and tested on H. armigera and found that AgNPs modulated gut protease activity in larvae of H. armigera. Vinayagamoorthi et al. (2015) synthesized AgNPs from agueous extract of Sargassummuticum (Yendo) Fensholt (Fucales: Sargassaceae) and tested on 4th instar larvae of the common castor, Ariadne merione (Cramer) (Lepidoptera: Nymphalidae) and observed the physiological and anatomical abnormalities in the larval body.

Silica Nanoparticles (SiNPs)

SiNPs are most studied nanoparticles either as nanocarriers or as biopesticide in one or the other form (fungicide, bactericide, pheromone, plant growth regulator) against number of insect pests both under field and storage conditions (Barik et al. 2008). The mode of action of SiNPs is similar to that of bulk silica where in SiNPs are physio-sorbed by the cuticular lipids destroying the protective barrier and thereby causing insect to death. World Health Organization (WHO) declared use of amorphous silica as nanobiopesticide is safe to humans (Athanassiou et al. 2017). The surface charged SiNPs (3-5 nm) were successfully used for management of insect pests agricultural and veterinary importance across the taxa (Ulrichs et al. 2005). Debnath et al. (2011) reported the application of SiNPs caused 100% mortality of adults of storage pest rice, rice weevil *Sitophilus oryzae* (L.) (Coleoptera: Curculionidae). Fouad et al. (2016) reported that application of SiNPs (600 ppm) along with Jasmonic acid at rate 1.141 µM/plant significantly reduced the tomato fruit damage by *Tuta absoluta* larvae.

El-Samahy et al. (2015) reported that 70.11 and 60.56% reduction in larvae of *Spodoptera littoralis* (75 and 60 g/fed) in sugarbeet due to application of SiNPs. Similarly, El-Helaly et al. (2016) tested the SiNPs at 200, 300, 400 and 500 ppm along with bulk silica and diazinon against *S. littoralis* in squash and

reported that 73.07, 79, 72, 87.88 and 89.82% mortality of larvae at respective doses. Shoaib et al. (2018) tested the SiNPs against *P. xylostella* larvae @ 1 mg cm⁻² and reported that mortality percentage increased from 58% and 85% at 24 and 72 h after treatment and further noticed that the larval death was due to desiccation, body wall abrasion, and spiracle blockage.

Conclusions

Nanotechnology offers grater applicability in various fields of agriculture and allied scineces. It has tremendous role in insect pest management strategies such as nano carrier, nano emulsions, nano pesticides and so on. Biopesticides are one such class of compounds which offer environmentally friendly, residual and pollution free control of insect pests in various crops. Nano form of biopesticides offer greater advantage over traditional biopesticides in achieving maximum potential in controlling target insect pests. More studies are required on effect of nanobiopesticides on natural enemies and their role in tri-trophic interactions. With this, it can be concluded that nanobiopestcodes are the best alternatives for chemical pesticides and traditional biopesticides for successful and sustainable management of insect pests of various crops.

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Impact of Biopesticides Application on Crop Quality and Environmental Quality

Abstract

Synthetic organic insecticides are posing serious health hazards to human beings and other non-target organisms across the world. The consumers are well aware of the ill effects of insecticides and there is a great demand for pesticide free agricultural produces. In few occasions, Indian consignments such as chillies, mango, vegetables, wheat have been either rejected by the importing countries or in the international markets due to pesticides residues, quarantine pests etc. Insecticides of biological origin such biopesticides are gaining importance as they are environmentally safe, leaving no residues in products etc. To date more than 12 biopesticides have been registered in India for their use in pest management. Many more are in pipeline to combat the yield losses caused by pests. Some of the biopesticides such EPN's have controlled white grubs without compromising soil and environmental quality. Biopesticides such as Bacillus subtilis GAL and Bacillus sp. have well preserved the mango juice over a period of 15 days which was as good as chemical preservative there by improved the shelf life of perishable fruits. The government of India has given greater emphasis on natural farming/zero budget farming, organic farming where biopesticides play crucial role in management of pest populations. In this chapter, we have narrated the economic importance of biopesticides, international trade issues, impact of biopesticides on crop quality, soil and environmental quality.

Keywords: Biopesticides, Soil, Environment, Quality, Contamination

Introduction

Insecticide based plant protection in India as well as in the world resulted in wide variety of environmental and health issues. Insecticides although gave

satisfactory control of target pests initially, have posed serious issues such as resistance, resurgence and residue and also contamination of water bodies, food chain leading serious ill effects in humans, mammals, soil beneficial microbiota etc. After thorough examination of scientific evidences and facts on ill effects of synthetic organic insecticides, much emphasis was given for biological control including parasitoids, predators, entomopathogens, biopesticides, microbial control etc. in order to overcome above ill effects. The concept of biopesticides have come up as an alternative to the indiscriminate use of harmful synthetic insecticides which are being extensively used in organic and natural/zero budget farming. Biopesticides are formulations made from naturally occurring substances like animals, plants, microorganisms and include living organisms, their products or byproducts that control pests by non-toxic mechanisms in an ecofriendly manner. The biopesticides such as botanicals, Entomopathogenic fungi, Bt, NPV, EPN, PI, etc. which are widely used in the global market including India. Biopesticides may be categorized into three major groups: plant-incorporated protectants (PIPs), biochemical, and microbial biopesticides. While microbial biopesticides use microorganisms (bacteria, fungi, viruses or protozoans) as active-ingredient, biochemical pesticides are naturally occurring substances from plants and animals. PIPs are produced naturally on genetic modification of a crop plant, such as Bt cotton. Such transgenic plant produces biodegradable protein with no harmful effect on animals and human beings, and thus curtails the use of hazardous pesticides. PIPs may be more effective and economical strategies in the developing countries to help produce more food, feed and forages in an environmentally safer manner. However, the pesticides of biological origin have also been reported to cause relatively less side effects which are being discussed in this chapter.

Merits of Biopesticides

The biopesticides are more preferred in today's health conscious world due to following advantages.

- Biopesticides are generally less toxic than chemical pesticides often target specific pests
- Little or no residual effects hence pose less risks to human health and environment
- Have wide acceptability for use in the organic farming
- Many biopesticides have a zero or low re-entry and handling interval
- Some microbial biopesticides can reproduce on or near to the target pest / disease, giving some self-perpetuating control

- The risk of pests and disease developing resistance to biopesticides is often considered to be low
- They often have good compatibility both with biological pest control agents (natural enemies) and conventional chemical pesticides, so can be readily incorporated into IPM
- They can also be useful as a second line of defence or supplementary treatment
- Relatively less costly
- Enhanced crop quality
- Adequately degradable
- No harmful residues remain in food, fodder and fibers
- Growing market preferences

Constraints in Exporting Agricultural Commodities

India is one of the largest producers of a number of agriculture commodities and the EuropeanUnion (EU) is one of the largest export markets for India. India is seeing growth in the export of agricultural commodities like cereals, non-basmati rice, wheat, millets, maize, and other coarse grains and the largest markets for India's agricultural products are the US, China, Bangladesh, the UAE, Vietnam, Saudi Arabia, Indonesia, Nepal, Iran, and Malaysia.Demand for Indian cereals was robust in 2020-21, with shipments sent to several countries for the first time, such as rice to countries like Timor-Leste, Puerto Rico, and Brazil. Similarly, wheat was despatched to countries such as Yemen, Indonesia, and Bhutan, and other cereals have been exported to Sudan, Poland, Bolivia. In financial year 2021, fresh fruits were the leading horticulture product exported from India (56 billion Indian rupees). Over 956 thousand metric tons of fruits were exported that year from the south Asian country. Organic exports that include products such as cereals and millets, spices and condiments, tea, medicinal plant products, dry fruits, and sugar grew 51 per cent year on year to \$1,040 million. However, the pesticide residues in the commodities are the major bottlenecks in exporting the quality products to international market. Pesticide residue problems have affected exports of basmati rice which is the key traditional export product to the EU, due to stringent norms imposed for chemicals such as Tricyclazole and Buprofezin, extensively used in rice cultivation in India. Testing by the Export Inspection Council (EIC) has been made mandatory for basmati exports to the EU, which led to a decrease in the number of alerts.

In recent years a number of Indian agricultural products have been facing rejection and export bans in the EU due to standards related to food quality, safety and health, sanitary and phytosanitary (SPS) measures. The products such as mangoes, grapes and eggplants in which Indian exporters have faced rejections or bans in the EU and other markets in the past for SPS issues such as fruit flies or thrips infestation. Among the pesticides/chemicals, aflatoxins had the maximum notifications in Basmati rice, followed by Carbendazim, Acephate, Triazophos, Hexaconazole and other miscellaneous pesticides (such as bromide, chlorpyrifos, ochratoxin and profenofos).. EU rejected table grapes consignments from India in 2010, leading to a slowdown in the industry. Reduction in the chlormequat chloride limits in grapes from 0.05mg/kg to0.01mg/kg in the year 2016 which hampered the export of grapes.Saudi Arabia actually cited pesticide levels beyond its own MRLs to block shipments of green chillies and cardamom from India. In the EU, on the other hand, Indian export products that have faced issues on MRL levels in 2020 include:

- Sesame Seeds: Ethylene Oxide (insecticide)
- Chillies: Chlorothalonil (fungicide)
- Frozen curry leaves: Chlorpyrifos (pesticide)
- Frozen diced red chilli puree: Methamidophos, monocrotophos, acephate, propargite and triazophos
- Basmati Rice: Thiamethoxam, tricyclazole and buprofezin

Impact of Biopesticides on Crop Quality

Crop quality is of utmost importance to both growers and consumers. Plant physiology is highly responsive to the prevailing environmental conditions that plays a critical role in both quantity and quality. Active management of plant physiology plays an important role in crop productivity, and biopesticides, particularly those in the plant growth regulator category (PGRs), are key tools in this regard. Characteristics such as fruit size, taste, texture, shape, colour, firmness and shelf life can all be enhanced by careful use of plant growth regulators. In addition, some PGRs can give a boost to plant health by increasing the root mass or enhancing resistance to pests and disease. PGRs have the added benefit of being non-toxic. No harmful residues remain to delay handling or consumption. Crop quality and yield largely determine a grower's income. Biopesticides provide dealers with products that can markedly improve crop quality and yield by preventing pest damage and promoting physiological benefits in plants, including increased fruit size and enhanced colour. Dealers who supply biopesticides and encourage their innovative use are on the forefront of yield-and-profit enhancement practices. Most bio-based pest management products are listed for use in organic farming, providing those growers with compelling pest control options to protect yields and quality. A heterogeneous representation of target products, such as winter guava, mango, apple, mandarin, kiwifruit, strawberry, pepper fruit, red-fleshed table grape, pineapple, cherry fruit, papaya, plum needs much attention in consumer point of view due to ill effects of synthetic pesticides.

Bacillus subtilis GA1 and Bacillus sp. have well preserved the mango juice over a period of 15 days which was as good as chemical preservative. Biopesticides serves as good preservatives and attract consumer preferences than chemical (Kohi et al. 2020). Biopesticides such as T. harzianum T22 and 6PP are able to improve crop yield and increase the total amount of polyphenols and antioxidant activity in the grapes by reducing the powdery mildew fungi indicting the improvement of crop quality (Pascale et al. 2017). Besides, enhancement of corn yield was reported in several commercial which has been considered as a direct effect of an increased root and foliar systems (Harman 2000). The PGPR activity is induced by Trichoderma can be explained by an upregulation of photosynthesis related proteins and a higher photosynthetic efficiency, enhanced the plant nutrient uptake mechanism and increased plant nitrogen use efficiency etc (Harman et al. 2004). Even biopesticides such as foliar spray of neem oil @1.5% along with tree pruning significantly improved fruit physical quality and cosmetic appearance of mandarin (Aftab et al. 2021). Arbuscular mycorrhizal fungi play major role in biological control of plant diseases owing to their capabilities of amelioration crop yields by multiple role as bio-pesticides and plant growth promotion (Nelson 2004). Mycorrhiza can be seen as an assurance against quality deterioration caused by stress factors and also positive effects not only on plant growth, but also on plant quality that include improved product quality of lettuce, tomato, pepper and strawberries (Baum et al. 2015). Preharvest foliar spray of fungal culture filtrates from Aspergillus niger and Rhizopus oryzae improved the plant defence mechanism, with also enhanced quality and shelf life of date fruit in India (Bhatt and Jampala 2020). Pre-harvest treatment with Metschnikowia fructicola for the control of postharvest rots not only reduced the fruit rots significantly but also improved the fruit quality strawberry quality (Karabulut et al. 2004; Sellitto et al. 2021). Botanical and microbial fungicides are effective for suppressing botrytis fruit rot in strawberry alone or in rotation with synthetic fungicides thereby increased the quality for fetching higher market value (Dara 2020).

Impact of Biopesticides on Environmental Quality

Sustainable use of agro-pharmaceuticals, together with the demand for more environment-friendly production systems are the need of the hour in health-conscious consumer world. A growing public interest in the search for alternative approaches to chemical control in biotic stress management is very much required. The time-tested indigenous technical knowledge (ITK) of using natural materials for the control of pests has been very effective which need to be practiced. Biopesticides pose less threat to the environment and human health. They are generally less toxic than chemical pesticides, often target specific, have little or no residual effects and have acceptability for use in organic farming. Use of botanicals is now emerging as one of the important means to be used in protection of crop produce and the environment from pesticidal pollution, which is a global problem. There is less danger of biopesticide impact on the environment and water quality and they offer a more environmentally friendly alternative to chemical insecticide. Biopesticides have long been attracting global attention as a safer strategy than chemical pest control, with potentially less risk to humans and the environment. To this end, co-operation between the public and private sectors is required to facilitate the development, manufacturing, and sale of this environmentally friendly alternative.

Case Studies

Entomopathogenic Nematodes (EPN) for Crop and Soil Health

Entomopathogenic nematodes (EPN) of families Heterorhabditidae and Steinernematidae are microscopic, non-segmented roundworms that are obligate parasites of insects and have become important in biological control and integrated insect pest management as biopesticides. Soil insect pests including white grubs, cutworms, termites, root grubs, etc., cause 24-40% yield losses in sugarcane, maize, arecanut, cardamom, groundnut, potato, banana, guava, turmeric, pulses, vegetables, grasses, lawns etc., and direct plant loss to the tune of 20-60% in arecanut, sugarcane, cardamom, banana, groundnut, turmeric, guava, soybean etc. Due to continuous depletion of forest cover and organic carbon, summarily attributed to anthropogenic and geological events, the soilborne insect pests are increasingly causing a serious threat. Many synthetic chemicals like OP, carbamates, neonicotinoids, fumigants etc., are in indiscriminate use with little effect on the target pest, but causing soil and water body contamination, residual effects on soil biota, human and animal wellbeing, soil health and productivity. Farmers are desperately looking for ecologically safe, sustainable and on-farm recyclable green technologies, alternative to soil-contaminating synthetic chemicals, to secure their crop losses, soil health and productivity. EPN infect many different types of soil insects, and their life stages, larval, pupal and adult forms of Lepidopteran, Coleopteran and Dipteran pests.

ICAR-National Bureau of Agriculturally Insect Resources, Bengaluru has developed and scaled up novel WP formulations of EPN consists of infective juveniles capable of controlling soil born insect pests like whitegrubs and other insects having a shelf life of 10-12 months. These formulations are easy easyto transport, application and safety during transport and storage. The WP formulation of nematodes developed is effective for controlling a variety of obnoxious and cryptic soil insect pests that are hardy to pesticides on a number of crops, including arecanut, banana, cashew, sugarcane, potato, maize, groundnut, redgram. The impact of this technology has been assessed in large scale by producing 1200 tonnes WP formulation of EPN to cover an area of 20,000ha for the management of whitegrubs & other soilborne insect pests. Very encouraging results were obtained indicating the impact of this green technology for the management of whitegrubs in above crops. Thse WP formulations were licenced to 12 firms so that farmers use it for the management of whitegrub in above crops very effectively by securing soil and environmental health (EPA, 2016).



Conclusions

Biopesticides are gaining importance across the globe for their environmental safely, crop quality, soil quality and consumer preference. Biopesticides have significantly improved fruit quality (nutritional) in various fruit crops, vegetable crops etc besides suppressing insect pest populations in crop ecosystems for sustainable production. They also enhance the export quality of the agricultural products and fetch very high prices in the international market. Therefore, farmers should be encouraged to produce residue free and good quality agricultural products/commodities in phased manner. Many more biopesticides with good shelf life that control wide variety of major insect pests may be introduced to market to overcome hazardous effects of synthetic pesticides. Farmers may also be convinced by demonstrating merits of biopesticides over conventional synthetic insecticides for increasing adoption rate.

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7

Indian Biopesticides Market, Consumption, Growth and Opportunities

Abstract

In India, the market for biopesticides has reportedly grown at a quick and rapid rate (23%) over the previous ten years, whereas the market for chemical pesticides has only grown by 2%. However, the usage of biopesticides has not yet reached the same level as that of chemical pesticides, but it is predicted to do so between 2040 and 2050. There is reportedly a 30% difference between the demand for and consumption of microbial biopesticide in India. Anytime the use of biopesticide is encouraged to a broad adoption by stakeholders through appropriate development of awareness, further market sizing is conceivable. Maharashtra state consumed the most biopesticide formulations, whereas Chhattisgarh, Haryana, and other Indian states consumed the least. According to crop consumption, cereal crops receive the most, followed by pulses, oilseeds, fibre, fruits, and vegetables. The market and consumption of biopesticides in India and globally are reported.

Keywords: Biopesticide, Demand and Consumption, State-wise, Crop-wise

Introduction

A tremendous change in the Indian biopesticide industry and market has reflected on increased global trade in agricultural commodities, a healthy revolution in the consumers and stack-holders with adequate understanding on negative effective of chemical pesticides in plant protection. Currently biopesticides comprise approximately 3-5% of the Indian pesticide market, with at least 15 microbial species and 970 microbial formulations registered through the Central Insecticides Board and Registration Committee (CIBRC). As of 2017, over 200 products based on entomopathogenic fungi (*Beauveria bassiana, B. brongniartii, Metarhizium anisopliae, Lecanicillium lecanii* and

Hirsutella thompsonii) and nematicidal fungi (*Purpureocillium lilacinum* and *Pochonia chlamydosporia*) are registered for use against various arthropods and plant parasitic nematodes. Regarding bacteria, over 30 products based on *Bacillus thuringiensis* (Bt) subsp. kurstaki are registered against bollworms, loopers and other Lepidoptera, while 12 based on *Bt* subsp. israelensis and three with *Bt* subsp. sphaericus are being used against mosquitoes.

Two viruses are registered, namely *Helicoverpa armigera* nucleopolyhedron virus (22 products) and *Spodoptera litura* NPV (5 products) for use against bollworms and armyworms. Four entomopathogenic nematode species consisting wettable powder formulations of *Heterorhabditis indica* developed by the ICAR-National Bureau of Agricultural Insect Resources, Bengaluru which have been distributed on a large scale to control white grubs and other sugarcane pests. Biopesticide research in developing countries like India though in infant stage, but evolving rapidly, and focusing on indigenous entomopathogens. Despite enormous regulation, quality-control issues and limited large-scale production facilities, investment in domestic fermentation technologies, improved delivery systems, and promotion of biological control through private and public initiative will increase the share of microbial biopesticides in the country.

Biopesticide Market: Global and Indian Perspectives

Biopesticides are organic substances used to control pests that are derived from plants, animals, microbes, and some minerals. Only one entomopathogenic bacteria, *Bacillus thuringiensis*, is the source of about 90% of the microbial biopesticides currently on the market (Kumar and Singh 2015). Currently, biopesticides only account for a small portion of the overall crop protection business, with a value of roughly \$3 billion globally, or 5% of the total crop protection market (Marrone 2014; Olson 2015).

There are more than 200 items accessible on the United States (US) market, compared to only 60 comparable products on the European Union (EU) market. Although the global market for these pesticides appears to need to expand further in the future if these products are to play a significant role in replacing chemical pesticides and lessening the current over-reliance on them, even though the use of biopesticides is rising globally by almost 10% annually (Kumar and Singh 2015). However, it should be noted that the EU uses the same regulations for evaluating biopesticides as they do for synthetic active substances. As a result, several new provisions in the law were needed, and new guidelines were also created to make it easier for potential biopesticide products to be registered (Czaja et al. 2015). The EU currently has less

registered biopesticide active chemicals than the US, India, Brazil, or China. The greater complexity of EU-based biopesticide laws is connected to the comparatively low level of biopesticide research in the EU (Balog et al. 2017).

With compounded annual growth rates of more than 15%, biopesticides are expected to outgrow chemical pesticides in terms of growth (Marrone 2014). Between the late 2040s and the early 2050s, it is anticipated that the market size for biopesticides will equal that of synthetics, but there are significant uncertainties surrounding the rates of uptake, particularly in regions like Africa and Southeast Asia, which account for a significant portion of the flexibility in those projections (Olson 2015). In recent years, biopesticides have grown in popularity and are thought to be safer than traditional pesticides. Biopesticides have the potential to reduce the usage of conventional pesticides as essential elements of IPM programmes because they are effective in small amounts and breakdown quickly without leaving harmful residues. However, it should be noted that while there may be situation-specific exceptions to the aforementioned qualities, they do not negate the overall norm.

Indian market is a house to hundreds of biopesticides that are duly registered by the Central Insecticides Board and Registration Committee (CIB&RC), but quality control is a major problem in most of these products. Extensive research on biopesticides in national laboratories and State Agricultural Universities has clearly demonstrated the efficacy of biopesticides for management of pests and diseases. Regardless of the persistent government programs and initiatives, the consumption of biopesticides in India has remained relatively low, for several years in past especially since 2000s. The recent years have witnessed the introduction of nanotechnology mediated biopesticides. Nanoparticles mediated biopesticides have shown considerate potential in alleviating the problems associated with conventional pesticides. The market has attained speedy growth over the period of six years from FY'2013 to FY'2019. The potential benefits of using biopesticides in agriculture and public health programs are considerable. This has tremendously escalated the consumption for biopesticides in the country over the years resulting in a double digit (https://www.researchandmarkets.com/reports/5003583/ growth CAGR biopesticide-market-trends-forecast-and#rela3-5214644). In FY'2019 the revenue generated through microbial pesticides consisted of a microorganism (e.g., a bacterium, fungus, virus or protozoan) as the active ingredient which contributed to a majority of the proportion in the overall Biopesticides market. Microbial biopesticides are eco-friendly pests management solutions and have high specificity due to which share of microbial biopesticides has contributed highest share in terms of revenue in FY'2019. Invertebrate pathogenic microorganisms employed as active substances in pest management are

recognized as generally safe for the environment and non-target species, in comparison with synthetic chemicals. Botanical/biochemical and PIP was observed to capture the remaining volume share in the FY' 2019.

With improved seed vigor and introduction of systematic disease resistance, the demand for this fungal symbiont has remained high and rendered a majority share in the Indian biofungicides market in FY'2019. During FY'2019, *Bacillus thuringiensis* var. kurstaki contributed to a majority of the bioinsecticides sold in India. Due to its high effectiveness and quicker results, it is preferred over any other bioinsecticide present in the Indian market and thus, contributed the highest share of in the Indian Bioinsecticide market in FY'2019. The application of biopesticides is spread across several crops in agriculture. The share of cereals, pulses and oilseeds has been recorded the highest, and has commanded a major portion of the overall bio-pesticide consumption in India during FY'2019. Flower, spices and tea constituted the smallest market share in terms of revenue generated.

In India, the demand for indigenous biopesticides has dominated the overall biopesticides market during FY'2019. Imported biopesticides which mainly includes Bacillus, semiochemicals and others have accounted for the rest of the market. The consumption of biopesticides was dominated by western region in FY'2019 followed by South, East, North and North Eastern region. The Major companies in the market of biopesticides are EID Parry, T Stanes, Fortune Biotech, Excel Crop Care, International Panaacea Ltd, Biotech International, Kan Biosys, PCI, PJ Margo, Prathibha Biotech and Zytex Biotech. Price, quality and distribution network are some of the critical parameters on the basis of which companies compete in the organized segment.

India Biopesticide Market

In India, the usage of biopesticides is growing at a faster pace than that of the chemical pesticides. According to the Directorate of Plant Protection, Quarantine and Storage, Ministry of Agriculture and Farmer Welfare, in the last 10 years, consumption of bio-pesticides increased by 23%, while that of chemical pesticides grew only by 2%. The total demand by various states/UTs of India was reported as 59,458 MT technical grade of biopesticides which was 8795 MT at 2014-15 and 10,852 MT at 2019-20 with a sharp decline at 2018-19 (9725 MT) (Fig. 1). However, there was a gap between demand and consumption of biopesticides in India in which only 70.2 % of total demand was consumed under various crops, grown in different states of India (Fig. 2,3).

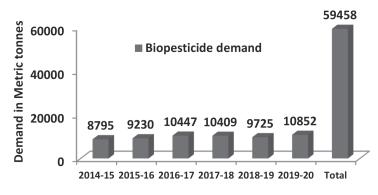


Fig. 1: India biopesticide demand during 2014-2020

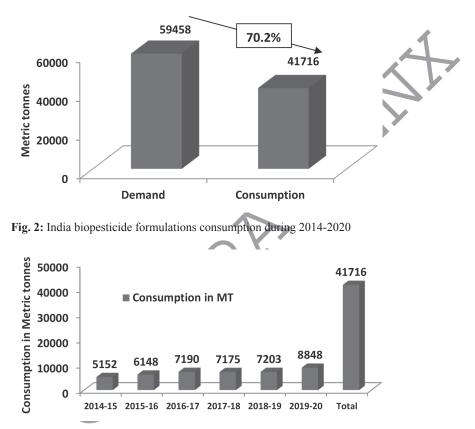


Fig. 3: Total demand and consumption of biopesticide formulations in India during 2014-2020

The total consumption of biopesticides was computed as 22,404 MT technical grade during 2014-2020 which was 9.07% of total chemical pesticides consumed (2,41,969 MT) (Fig. 4).

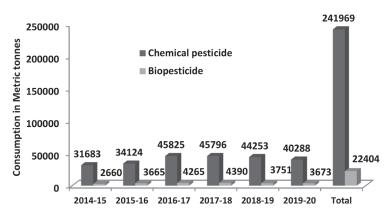


Fig. 4: Comparative analysis on consumption of chemical pesticides and biopesticides in India during 2014-2020

Among the states, Maharashtra was leading to absorb maximum quantity of biopesticides (6630 MT) during 2014-2020, followed by West Bengal (5433 MT), Kerala (4257 MT), Karnataka (3160 MT), Tamil Nadu (2816 MT), Madhya Pradesh (2737 MT), Chhattisgarh (2549 MT) and Haryana (2250 MT) (Fig. 5) (https://ppqs.gov.in/statistical-database).

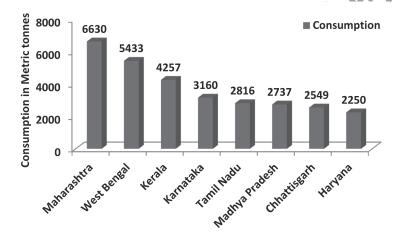


Fig. 5: State-wise consumption of biopesticide formulations during 2014-20 in India The 9.4% of total cultivated areas (4.86 million ha) in cereals, pulses, oilseeds, fibre, furits, vegetables, plantation and other crops during 2014-2020 was reported to be covered with biopesticides in India. Cereal crops consumed the maximum quantity of biopesticides (4876 MT), followed by oilseeds (4434 MT), vegetables (3980 MT), pulses (2905 MT), fruits (1819 MT) etc., during 2014-2020 (Fig. 6)

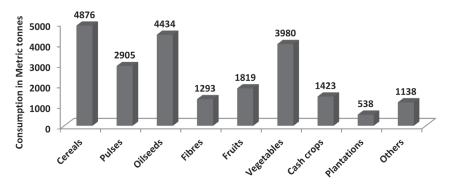


Fig. 6: Crop-wise consumption of biopesticide formulations during 2014-20 in India

Global Biopesticide Market Growth

Global biopesticides market has occupied in a small fraction of the total global crop protection market and it was estimated as \$ 1.72 billion in 2014; \$ 1.89 billion in 2015; \$ 2.09 billion in 2016; \$ 2.31 billion in 2017; \$ 2.55 billion in 2018; \$ 2.81 billion in 2019; and \$ 3.09 billion in 2020. The biopesticide market is anticipated to contribute \$ 3.42 billion in 2021; \$ 3.77 billion in 2022; \$ 4.5 billion in 2023; and \$ 8.19 billion in 2025 (Fig. 7). At 2020, the compound annual growth rate (CAGR) of global biopesticide market was approximately 3-5% of the total crop protection market (Marrone 2014, Olson, 2015; Kumar et al. 2018; Damalas and Koutroubas 2018) while the market was anticipated to grow by 8.64 % at 2023; 9.7% at 2015-2023; 10.3% at 2014-2022; 15% at 2019-2024; 16% at 2020-2025 (https://www.researchandmarkets.com/reports/5003583/biopesticide-market-trends-forecast-and#rela3-5214644).

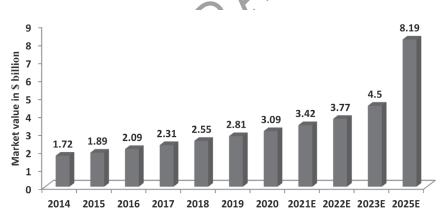
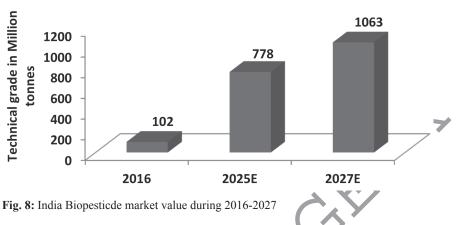


Fig. 7: Global Biopesticide Market Value during 2014-2025

India Biopesticide Market Growth

The India biopesticides market generated revenue of \$102 million in 2016 and is anticipated to contribute \$778 million by 2025, growing at a CAGR of 25.4% (Fig. 8). The growth rate of biopesticide market in India varied in different periods of report and the CAGR was reported/ anticipated to be 9.3% at 2013-2018; 16.4% at 2013-2019; 7.3% at 2016-2026; 25.4% at 2017-2025; 10.3% at 2018-2024; 13.1% at 2019-2027; 25.1% at 2016-2025 (https://inkwoodresearch. com/reports/india-biopesticides-market-forecast- 2017-2025) (Fig. 8).



Conclusions

As restrictions have tightened in recent years, the pipeline of new chemistry has significantly decreased. Products are being pulled off the market because they no longer adhere to the severe standards. As a result, a smaller selection of chemical remedies continues to target numerous pests in a small number of staple crops. These effects, which have always been clear in the market for pesticides, are now more clear than ever. Future market growth for biopesticides will be closely correlated with biological control agent research. There are few comprehensive and systematic studies on the preliminary research that a number of scientists from various research institutes have conducted in the area. Therefore, it is crucial to improve businesses and research institutions' cooperation on this issue. The agriculture industry can and should profit from the coexistence of biopesticides and chemical pesticides as it appears that biopesticides cannot yet totally replace chemical pesticides. In this context, it is anticipated that large-scale industrial development will be facilitated by speeding the practical application of research findings.

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Links

https://ppqs.gov.in/statistical-database

- https://inkwoodresearch.com/reports/india-biopesticides-market-forecast-2017-2025
- https://www.researchandmarkets.com/reports/5003583/biopesticide-market-trends-forecastand#rela3-5214644
- https://www.researchandmarkets.com/reports/5003583/biopesticide-market-trends-forecastand#rela3-5214644

ALP'P



Biopesticides Research and Development Activities

Abstract

In the creation of prospective biopesticides, bacteria and fungi are the main focus among the beneficial microbes utilised in biological control. Even if there is more research and development being done on the discovery of microorganisms and subsequent commercialization, the slow kill rate, short shelf life, resistance to microbe etc., are seen as a setback in the biopesticide industry. Several attempts have been made to increase the rate of kill, including developing Bt-based transgenics, transferring the genes encoding the production of toxins from scorpion, spider, etc. into the genome of helpful microbes etc., that increased the rate of kill by several folds. To perfect such genetically modified microbe based biopesticides, additional research is necessary.

Keywords: Beneficial microbes, Biopesticide, Genetic improvement

Introduction

Worldwide, various biopesticides have been created and are in use, including viruses, microorganisms (bacteria, fungi, etc.), microorganism derived products, animal derived goods (pheromones, hormones, insect-specific toxins, etc.), plant derived products, and genetically modified organisms (Nicholson 2007; Erlandson 2008; Mazhabi et al. 2011; Islam and Omar 2012). Microbial biopesticides are the greatest class of broad-spectrum, pest-specific among all currently employed biopesticides (safe to non-target organisms and eco-friendly). Thirty member nations of the Organization for Economic Co-operation and Development (OECD) offer more than 200 microbial biopesticides (Kabaluk and Gazdik 2007). According to Kiewnick (2007), there are 21 microbial biopesticides registered in the European Union (EU), 22 in Canada, and 53 in the United States. However, reports of the

items registered for usage in Asia vary (Thakore 2006). Overall, microbial biopesticide registrations are increasing globally, the expansion of various technologies has increased the scope for more products and the change in the trend to develop microbial products is definitely on the rise (Bailey et al. 2010 and Kristiofferesen et al. 2008; Shukla 2019).

Entomopathogenic Bacteria

Bacillus that forms crystalliferous spores (Bacillus thuringiensis), obligate pathogens (Bacillus popilliae), prospective pathogens (Serratia marcesens), and facultative pathogens (Pseudomonas aeruginosa) can all be classified as biopesticide-producing bacteria. Due to their efficacy and safety, spore formers have been the most commonly used for commercial purposes. Bacillus sphaericus and B. thuringiensis are the most often used microorganisms. A unique, secure, and efficient tool for controlling insects is B. thuringiensis (Roy et al. 2007). It is largely a pathogen of lepidopterous pests like rice stem borers and the American bollworm in cotton. Bt releases poisons when consumed by insect larvae, damaging the pest's midgut and ultimately killing it. The strains of the subspecies kurstaki, galeriae, and dendrolimus are the primary sources for the manufacturing of Bt preparations. Other bacterial species have limited effect on pest management, although there are commercial products based on Agrobacterium radiobacter, B. popilliae, B. subtilis, Pseudomonas cepacia, Pseudomonas chlororaphis, Pseudomonas flourescens, Pseudomonas solanacearum, and Pseudomonas syringae.

Insect Viruses

More than 700 viruses that infect insects have been discovered, the majority of which originate from Lepidoptera (560), followed by Hymenoptera (100), Coleoptera, Diptera, and Orthoptera (40) (Khachatourians 2009). A dozen or more of these viruses have been made available for commercial usage as biopesticides. The RNA-containing reoviruses, cytoplasmic polyhedrosis viruses, nodaviruses, picrona-like viruses, and tetraviruses, as well as the DNA-containing baculovruses (BVs), nucleopolyhedrosis viruses (NPVs), granuloviruses (GVs), acoviruses, iridoviruses, parvoviruses, polydnaviruses, and poxviruses are used in insect management. However, NPVs and GVs have been the primary categories utilised in pest management. These viruses are efficient against insects that consume plants and are used extensively around the world to control pests in vegetables and field crops. Their use has significantly reduced the populations of gypsy moths, pine sawflies, Douglas fir tussock moths, and pine caterpillars in forest settings. Potato tuber moth is controlled by *Phthorimaea operculella* GVs in stored tubers, and codling

moth is managed by Cydia pomonella GVs on fruit trees (Arthurs et al. 2008). Insects including cabbage moths, corn earworms, cotton leaf worms and bollworms, beet armyworms, celery loopers, and tobacco budworms can also be controlled with virus-based solutions. Target-specific viruses called baculoviruses can infect and kill a variety of significant plant pests. When they are used against lepidopterous pests of cotton, rice, and vegetables, they are especially effective. Their use has been constrained to small areas because of the challenges associated with their large-scale manufacture. They are not available commercially in India, but are being developed on a modest scale by various IPM institutions and state agricultural departments. Natural baculoviruses have been successfully used to preserve crops and forests, but from an agro-industrial standpoint, they are ineffective insecticides and have a number of potential drawbacks (Possee et al. 1997; Inceoglu et al. 2006). Compared to chemical insecticides, they have a slower rate of mortality (from five days to more than two weeks) and have a narrower host specificity, limited field stability, susceptibility to UV exposure, short shelf life, and higher production costs.

Entomopathogenic Fungi

Trichoderma harzianum, Trichoderma viridae, Streptomyces griseoviridis, Verticillium chlamydosporium, Beauveria bassiana, Metarhizium anisopilae, Nomuraea rileyi, Paecilomyces farinosus, and Verticillium lecanii are some of the most often employed species and many of them have received global commercialization. An efficient fungicide against root rot that is transmitted through the soil is *Trichoderma*. It is especially important for dry land crops like chickpeas, groundnuts, black gram, and green gram that are prone to various diseases. Trichoderma based biopesticide is simple to make and only needs a fundamental understanding of microbiology. For the management of soil- and seed-borne diseases, this bio-fungicide is advised for use as a seed treatment, soil application, soil drenching, root dip technique, etc. Important crop diseases which are well managed with Trichoderma based biopesticides are Armillaria, Botrytis, Chondrostereum, Colletotrichum, Dematophora, Diaporthe, Endothia, Fulvia, Fusarium, Fusicladium, Helminthosporium, Macrophomina, Monilia, Nectria, Phoma, Phytophthora, Plasmopara, Pseudoperonospora, Pythium, Rhizoctonia, Rhizopus, Sclerotinia, Sclerotium, Venturia, Verticillium, and wood rot fungi. Many Trichoderma strains, mainly T. harzianum, T. viride and T. virens (formerly Gliocladium virens) play vital role in plant diseases management (Singh 2014). Additionally, recent studies suggest that Trichoderma strains may be used to handle abiotic stresses as salt and drought (Shukla et al. 2012; Rawat et al. 2011). In addition, green muscardine fungus (*Metarizhium anisopliae*), white halo fungus (*Beauveria bassiana*) and *verticillium lecanii* based biopesticides are also popular in management of beetle pests, Lepidoptera pests, sucking pests etc.

Entomopathogenic Nematodes

The entomopathogenic nematodes (EPN), which control weevils, gnats, white grubs, and numerous species of the Sesiidae family, are another group of biopesticide (Klein 1990; Shapiro-Ilan et al. 2002; Grewal 1990). Insects feeding in enigmatic settings such as soil-borne pests and stem borers are kept under control by this interesting EPN. Nematodes from the genera Steinernema and Heterorhabditis, which attack hosts as infective juveniles (IJs), are frequently used in pest management (Kaya and Gaugler 1993; Koppenhofer and Kaya 2002).

Protozoans

The use of protozoan pathogens as biopesticide agents has not been particularly effective, despite the fact that they naturally infect a wide variety of pests and cause chronic and crippling effects that lower the target pest populations. Taxonomically speaking, protozoa are split into various phyla, some of which have entomogenous species. In-depth research has been done on microsporan protozoans as potential inclusions in integrated pest management plans. For many insect species, microsporidia are the disease-causing intracellular parasites that are ubiquitous and necessary. Because they target lepidopteran and orthopteran insects and tend to kill hoppers more frequently than any other insect, two genera, Nosema and Vairimorpha, offer some potential (Lewis 2002). According to research on the microsporidium Nosema pyrausta, which infects the European corn borer Ostrinia nubilalis, a spore is consumed by a larva of the European corn borer, which then germinates in the midgut, extrudes a polar filament, and injects sporaplasm into a midgut cell. The sporaplasm multiplies and creates additional spores, which can spread infection to other tissues. Infected midgut cells shed their spores into the gut lumen, where they are eliminated to the maize plant with the animal's waste. The infection cycle is repeated in the midgut cells of the new host as a result of these spores, which are still viable, being ingested during larval feeding. If a female larva is affected. Nosema is vertically transmitted to the filial generation. The developing overtes and ovarian tissue get infected with N. pyrausta as the infected larva grows into an adult. When the larvae hatch, they are infected with *N. pyrausta* since the embryo is already contaminated within the yolk. *N. pyrausta* is maintained in naturally occurring populations of the European corn borer by both horizontal and vertical transmissions.

Resistance to Microbes

The development of resistance has been observed most frequently in B. thuringiensis among the numerous families of microbial pathogens. At least 16 insect species have been discovered recently that are resistant to B. thuringiensis. Noctuid species like Spodoptera frugiperda, Busseola fusca, and H. zea have been found to have developed resistance to 8-endotoxins in the field (Tabashnik et al. 2009). The majority of reports of the development of resistance in *Plutella xylostella* field populations come from the nations that employ Bacillus thuringiensis widely, including China, Japan, the Phillipines, Malaysia, India, and North America. We now have B. thuringiensis Bt cotton and B. thuringiensis maize available in 13 and nine countries, respectively, grown on 42.1 million ha of land (Shelton et al. 2008). Genetic engineering was thought to be a useful tool to avoid this resistance problem where microbial genes from *B. thuringiensis* were transferred to plants to produce transgenics. In terms of microbial pest control, the introduction of such transgenics was hailed as a miracle cure; nevertheless, field resistance in H. zea as a result of an increase in the frequency of resistance alleles is concerning (Tabashnik et al. 2008). The field-evolved insect resistance to B. thuringiensis crops and various aspects related to resistance monitoring methods have been comprehensively reviewed recently (Tabashnik et al. 2009); obviously more prominent in lepidopterans (Downes et al. 2010; Huang et al. 2011). Factors associated with field resistance are the failure to use high dose B. thuringiensis cultivars and lack of a sufficient refuge. While implementation of the high-dose/ refuge insect resistance management strategy has been successful in delaying field resistance to Bt crops (Huang et al. 2011), Gene pyramiding is another approach used to try and address the emerging resistance problem (Zhao et al. 2003; Manyangariwa et al. 2006). Pyramiding is the stacking of various genes to cause the transgenic plant to express numerous toxins. However, gene pyramiding must be sustained and shouldn't result in numerous resistances or cross-resistances. Multiple resistance cannot be completely ignored because doing so would render these techniques useless in the end. In order to maintain the efficacy of pyramided B. thuringiensis crops, it is critical to account for the potential implications of such cross-resistance in resistance management plans. Pink bollworm has asymmetrical cross-resistance between B. thuringiensis toxins Cry1Ac and Cry2Ab (Tabashnik et al. 2009).

Gene pyramiding may not be a sustainable tactic per se, according to recent research, thus management plans must also include other tactics including refugia, the employment of predators and parasitoids, and crop rotation schemes (Zhao et al. 2003; Tabashnik et al. 2009). Soon, RNA interference-based transgenic plants that control insects will be a reality (Baum et al. 2007; Mao

et al. 2007), expanding the potential applications of transgenics and reducing the negative effects of resistance. Recent research has demonstrated that toxin-binding proteins like *cadherin* increase the toxicity of *B. thuringiensis* (Soberon et al. 2007). In contrast to the usual *B. thuringiensis* toxins, these binding proteins help toxin oligomerization and hence change the toxin, which can avoid resistance. The experiments show that *cadherin* gene silencing using RNA interference in *M. sexta* reduces the toxicity of *B. thuringiensis* toxin Cry1Ab. *M. sexta* and *Pectinophora gossypiella* that were resistant to *B. thuringiensis* were killed by the toxins that possessed *cadherin* deletion mutations (Soberon et al. 2007).

Recently, resistance in a baculovirus in the field has been found in Europe where *Cydia pomnella* GV is one of the main components of the codling moth control. *C. pomonella* GV in apple orchards has led to a high degree of resistance in some populations (Sauphanor et al. 2006; Frisch et al. 2007). This is the first documented instance of field resistance to a commercially applied baculovirus (Eberle and Jehle 2006). Apparently, this is either the result of the overuse of the product or the predominant control strategy applied. However, there do not seem to be any reported examples of field development of resistance to entomopathogenic fungi or nematodes (Shelton et al. 2007). However, there is evidence to demonstrate the existence of natural resistance mechanisms in insects against fungi (Wilson et al. 2001) and nematodes (Kunkel et al. 2004), suggesting that resistance to these pathogens cannot be summarily ignored.

Genetic Improvement of Insect Pathogens

Entomopathogenic Bacteria

The goal of genetically modifying microbial pathogens is to increase their potential to cause disease by enhancing toxin production, reproduction, and transmission rates. One strain of *B. thuringiensis*, for instance, exhibits insecticidal action against both coleopteran and lepidopteran insects as a result of genetic modification (Lereclus et al. 1992). Genetic modification can potentially increase *B. thuringiensis* activity on crop foliage or in soil treatments. For instance, the Cry34 and Cry35 families of crystal proteins from *B. thuringiensis* operate as binary toxins with action against the western maize rootworm, *Diabrotica virgifera virgifera*. Pairings Cry34A/Cry35A are busier than pairs Cry34B/Cry35B. The binary Cry34/Cry35 *B. thuringiensis* crystal proteins are closely linked to one another, are found throughout the environment, and have sequence similarities that are consistent with their ability to affect their target organisms' membranes. Plant pests and rootworms can be effectively controlled by modified Cry35 proteins, which have had

their segments, domains, and motifs swapped with those of other proteins to increase their insecticidal activity (Schnepf et al. 2007). Similar to this, the *B. thuringiensis* Cry8Bb1 toxin polypeptide was developed to feature a proteolytic protection site that renders it insensitive to a plant protease, aiding in the toxin's protection from any proteolytic inactivation. Modified Cry8Bb1 has been used for controlling corn rootworms, wireworms, boll weevils, Colorado potato beetles and the alfalfa weevils (Abad et al. 2008).

A new study demonstrates the presence of the *Bacillus enhancin-like (bel)* gene in the genomes of the *B. cereus* group, which has the potential to boost the insecticidal action of biopesticides based on *B. thuringiensis* and transgenic plants derived from *B. thuringiensis* genes (Fang et al. 2009). Bel genes produce peptides that resemble viral enhancin protein by 20-30%. As they destroy the peritrophic matrix of insect midguts, these proteins are known to strengthen viral infections. The mortality rate was 2.2 times higher when *Bel* and *Cry1Ac* were combined (Fang et al. 2009).

Insect Baculoviruses

Since the slow mortality rate of wild-type baculoviruses makes them impractical to utilise, numerous methods have been devised to increase the baculovirus's ability to kill by introducing genes encoding insect hormones, enzymes, or particular toxins (Kamita et al. 2005; Li and Bonning 2007; Gramkow et al. 2010). Maeda was the first to create a genetically altered baculovirus that expressed a gene encoding a hormone successfully in 1989 (Maeda 1989). This baculovirus produced the gene for a diuretic hormone, which led to water loss in Bombyx mori larvae and interfered with the insect's normal physiology. This modified BmNPV had a 20% quicker kill rate than the parent BmNPV's wild-type counterpart. This work developed a novel idea and laid the groundwork for later usage of baculoviruses to eradicate insects. Other enzymes and hormones were tested to alter baculoviruses in the years that followed.

Recombinant baculoviruses have demonstrated promise as more effective insect pest controllers. However, it is important to consider how utilising such viruses may affect the environment. Baculoviruses are not contagious to non-target creatures, including beneficial insect species, predators, and parasitoids of lepidopteran larvae, according to the findings of several research (Boughton et al. 2003; Sun et al. 2009). According to Hartig et al. (1991), recombinant AcMNPV baculovirus expressing AaiT was not infectious to adherent mammalian cells, and recombinant HaSNPV expressing AaiT was not pathogenic to fish, birds, or other vertebrates in any way (Sun et al. 2002). A

recombinant baculovirus does not possess any selected ecological advantages over the wild-type baculovirus, according to numerous investigations conducted both in the field and in greenhouse environments (Cory et al. 1994; Black et al. 1997; Lee et al. 2001). Additionally, there is a negative selection towards recombinant baculoviruses, which causes the wild-type to swiftly displace them (Georgievska et al. 2010; Zwart et al. 2010). The likelihood that the cloned gene will transfer from the recombinant baculovirus to another creature has also been conjectured. Although this is theoretically possible, it has never been demonstrated because of variables that prevent or restrict the occurrence of this genetic recombination (Inceoglu et al. 2001). Combination viruses have the potential to be more effective insect pests.

Recombinant baculoviruses were effectively used to express juvenile hormone esterase (Hammock et al. 1990), eclosion hormone (Eldridge et al. 1991), and prothoracicotropic hormone (O'Reilly et al. 1995). Only those producing juvenile hormone esterase, however, significantly outperformed parent wildtype baculoviruses in terms of insecticidal efficacy (El-Sheikh et al. 2011a). Juvenile hormone esterase controls the hormone, therefore when it is over expressed, the hormone's concentration falls. This causes the insect to stop feeding and pupate (El-Sheikh et al. 2011b). The effective utilisation of recombinant baculoviruses expressing this enzyme is severely hampered by the juvenile hormone esterase's brief half-life in the hemolymph. However, numerous attempts have been made to increase in vivo stability in order to make it more effective (Hinton and Hammock 2003; Inceoglu et al. 2006; Kamita and Hammock 2010).

Baculoviruses that have undergone genetic modification to express toxins have been widely used in the past. The first successful insertion of toxin genes into baculoviruses was reported in the late 1980s (Carbonell et al. 1988; Tomalski et al. 1991; Ooi et al. 1989). Since then, most studies have concentrated on comprehending arthropod-specific venoms produced by mites, spiders, or scorpions (Inceoglu et al. 2006). The Androctonus australis insectspecific toxin (AaiT) was the first and most effective insecticide expressed in baculoviruses (MacCutchen et al. 1991; Maeda et al. 1991; Stewart et al. 1991). When compared to the parent wild-type baculovirus, using a recombinant Bombix mori baculovirus (BmNPV) expressing AaiT accelerated the death of silkworm larvae by up to 40%. (Maeda et al. 1991). Another study utilising a different baculovirus expressing AaiT revealed that Manduda sexta larvae were paralysed many hours before death, increasing pesticidal efficacy (MacCutchen et al. 1991). The efficacy of baculoviruses that express AaiT was further validated in field experiments. (Cory et al. 1994; Sun et al. 2002; Sun et al. 2004).

Although AaiT has been the focus of numerous studies and is thought to be the best model peptide neurotoxin for enhancing the insecticidal activity of baculoviruses (Inceoglu et al. 2006; Sun et al., 2009), other scorpion toxins such as those from Leiurus quinquestriatus quinquestriatus, Leiurus quinquestriatus hebraeus, and Buthus marten (Kopeyan et al., 1990; Zlotkin et al. 1993; Moskowitz et al. 1998; Froy et al. 2000; Tang et al. 2011), spiders Agelenopsis aperta, Dighetia canities, Tegenaria agrestis and Araneus ventricosus (Prikhodko et al. 1996; Hughes et al. 1997; Jung et al. 2012), or straw itch mite, Pyemotes tritici is another source of powerful toxins that, when expressed in baculovirus, are active against insect pests and may one day be employed as biopesticides (Tomalski and Miller 1991; Lu et al. 1996; Burden et al. 2000). Another method for quickening the death of the baculoviruses is to delete an endogenous gene, such as the gene encoding the baculovirusencoded enzyme ecdysteroid UDP-glucosyltransferase (O'Reilly and Miller 1991). Because ecdysteroids are hormones that regulate larval-pupal moulting and eating behaviour, infection with an egt deletion mutant baculovirus results in a reduction in food consumption and an early mortality (Eldridge et al. 1992; Wilson et al. 2000; Cai et al. 2010; Georgievska et al. 2010).

The insect sodium channel is the molecular target of the majority of these neurotoxins (Cestele and Catterall 2000; Casida and Durkin 2013), which is also the main target of insecticides of the pyrethroid class. However, since their individual binding sites on the channel do not overlap, there is a chance of creating a synergistic effect that would permit the employment of both pyrethroids and baculoviruses that express toxins at the same time (McCutchen et al. 1997). The newest strategy involves the expression of the crystal protein gene from *Bacillus thuringiensis* in *Autographa californica* mNPV. In comparison to the parent wild-type AcMNPV, this recombinant baculovirus has demonstrated a high insecticidal activity against *Spodoptera exigua* and *Plutella xylostella* (Jung *et al.* 2012; Shim *et al.* 2013).

Entomopathogenic Fungi

Metarhizium anisopliae and *B. bassiana*, two commonly employed entomopathogenic fungi, have undergone substantial study for the clarification of pathogenic mechanisms and alteration of the genes of the pathogens to increase biocontrol efficacy (St. Leger et al. 2010). In the genome of *M. anisopliae*, extra copies of the gene encoding the controlled cuticle-degrading protease Pr1 were introduced and over expressed. Compared to the parent wild-type strain, the offspring strain decreased tobacco hornworm (*M. sexta*) survival time by 25%. (St. Leger et al. 1996). The scorpion toxin (AaIT) expressed in the *M. anisopliae* strain ARSEF 549 illustrates the astonishing

extent to which pathogenicity can be boosted. At 22-fold lower spore dosages than the wild type, the modified fungus produced the same mortality rates in *M. sexta*, and survival times at some concentrations were 40% lower (Wang et al. 2007). Similar outcomes have been seen with mosquitoes, where the LC_{50} was reduced by 9 times, and coffee berry borer beetles, where the LC_{50} was reduced by 16 times (Pava-Ripoll et al. 2008).

Entomopathogenic Nematode

Artificial selection has proven successful in boosting infectivity and nematicide resistance in entomopathogenic nematodes (Griffin 1993). With relation to host penetration and reproductive potential, the strain selection has demonstrated an improvement in fitness. The possibility of examining whether a selection strategy might enhance the control of root pests has been made possible by the recent revelation that maize roots harmed by the western corn rootworm release a crucial attractant for insect-killing nematodes (Hiltpold et al. 2010). After 10 to 25 selection cycles, a diverse population of *Steinernema feltiae* was produced for desiccation tolerance and host-seeking capacity. Antificial selection for one characteristic, however, may come at the expense of other crucial traits like contagiousness, establishment, and/or field persistence. In the near future, it may be possible to produce GM nematodes with higher storage stability, more resilience to environmental challenges, and greater biological control potential using data from the sequenced genomes of EPN (Sandhu et al. 2006; Ciche et al. 2007; Bai et al. 2009).

Conclusions

Despite having a slower death rate than chemical pesticides, wild-type beneficial organisms have demonstrated to be an effective long-term solution in particular situations, such as forest ecosystems. However, the parent wild-type microbe does not kill insects as quickly as recombinant microbes do, which is a severe drawback. These recombinant microorganisms are generally made of toxin genes from scorpions or spiders. Hopefully, recombinant microorganisms will get more market share worldwide. Numerous papers state that there is no evidence showing that genetically modified organisms pose a greater harm to animals and the environment than do organisms of the natural variety.

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9

Biopesticide Commercialization World-wide Regulation, Policies for Registration and Use of Biopesticides

Abstract

Biopesticides have become viable substitutes for man-made chemical pesticides in recent years. They are less expensive and don not endanger agro-ecosystems. Because of this, their demand and production are rising globally as well. In-depth examination reveals that there is no consistent regulatory approach that can streamline their regulation and registration procedure because the laws and regulations governing their usage and development differ from one country to the next. In spite of various effort by several international organisations like the Organization for Economic and Co-operative Development (OECD), International Organization for Biological Control (IOBC), and European and Mediterranean Plant Protection Organization (EPPO), some flexibility in biopesticide regulation has been offered, it still falls short of chemical pesticides, which have a strong market and well-established, non-overlapping legislation. World-wide regulation policies on biopesticide commercialization including registration and field use, limitations in regulations and modified regulations required are discussed in brief to understand the growth of biopesticides across the world.

Keywords: Biopesticide, Commercialization, Regulation policies Registration, Use

Introduction

The single piece of legislation under the Indian Government that regulates the import, manufacturing, sale, transportation, distribution, and use of all varieties of insecticides, including biopesticides, is the Insecticide Act (1968). Various parameters like shelf life, cross-contamination, moisture content, and packaging are significant factors that must be addressed before a biopesticide is registered. According to Organization for Economic Co-operation and Development's (OECD) recommendations, CIB simplified the protocols and listed the infrastructure needs for manufacturing of biopesticides. Information required to generate toxicity calls a strenuous effort. The demand for meteorological data, though, adds load on manufacturers and suppressed them from growing businesses. For instance, isolated microorganisms from one agroclimatic zone, possessing biocontrol property may or may not result the same findings in another agroclimatic zone. According to Rabindra (2005), Keswani et al. (2016), and available at http://ppqs.gov.in/divisions/ cib-rc/guidelines, new biopesticides should go through provisional/temporary registration under either 9(3B) or 9(3) section of the Insecticide Act 1968 by providing information on moisture content, shelf life, commodity potency with reference to LC_{50} , toxicity, secondary non-pathogenic microbial.

Regulation Policies of Biopesticide Registration

Generally speaking, the organisms chosen for insect management are effective only against the target insect. Therefore, it is assumed that there is a minimal chance of hurting non-target creatures, such as people. Before authorizing the widespread use of biopesticides, it is required to conduct certain standardized safety tests that will support the presumption and provide evidence of their efficacy. As a result, the Food and Agriculture Organization of the United Nations has established criteria, accordingly several countries have also developed their own guidelines for licensing biopesticides (Kulshrestha 2004). Perhaps the most difficult aspect of biopesticides is their registration. The number of registered biopesticide products has increased recently, but this number could increase further if the registration process is standardised globally. There are many different authorities and laws emerging to control biopesticides, but very little fatitude is offered. The laws observed in various nations and continents around the world are outlined in this section.

China

To regulate pesticide use and manufacturing in China, the Regulation on PesticideAdministration law was enacted in 1997. The law requires biopesticides to be registered before they can be sold (Kabaluk et al. 2010). Among other ministries, the Chinese Ministry of Agriculture (MOA) is authority of pesticide registration, manufacture, and commercial administration (Fang 2014). The Ministry of Agriculture's Institute for the Control of Agrochemicals (ICAMA 2008) is the apex regulatory body on monitoring the registration of pesticides, including biopesticides. The General Administration of Quality Supervision, Inspection and Quarantine of the People's Republic of China only permits registered and approved businesses to submit applications for the registration of pesticides (Kabaluk et al. 2010). Through financial support for insect control in forests, the Chinese Ministry of Forestry promotes the use of biopesticide. The good agricultural practices was inculcated among farmers by encouraging the use of biopesticides.

India

Government of India made many changes in regulations and laws to promote biopesticide manufacturers for registration. The Integrated Pest Management (IPM) initiative was overseen by the National Agricultural Technology Project (NATP) from 1998 to 2005, and the National Farmer Policy (2007) also supported the use of biopesticides in agriculture. By streamlining the licencing and regulating process for biopesticides, the Insecticide Act (1968) encouraged increased development and application of biopesticides. Under this act, the Central Insecticides Board (CIB) and the Registration Committee (RC) both functioned as extremely powerful entities for biopesticide regulation (www. cibrc.nic.in) (Kabaluk et al. 2010). The Apex Advisory Committee, or CIB, is composed of professionals from all relevant areas and fields. The CIB has simplified the criteria and data requirements for registration as well as the minimal infrastructure needs for the manufacture of biopesticides based on the OECD recommendations (NAAS 2013). After carefully examining and confirming claims on their bio-efficacy and safety for both humans and animals, the RC issues registrations. A key factor in the promotion of biopesticides is the National Agricultural Research System in which many ICAR institutes and State Agricultural Universities are involved (www.icar.org.in) (Rabindra 2005).

Africa

In order to create systems for the registration and regulation of biopesticides in the control of pests and diseases, some African nations adopt a variety of standards. Some African nations are taking the initiative to build their capabilities to control microbial pesticides. A regional inventory of the regulatory environments was conducted in 2012 by six country representatives from the West African region, including Kenya, Uganda, Ethiopia, Tanzania, Nigeria, and Ghana as part of the commercial Products (COMPRO II) project, which is run by the International Institute of Tropical Agriculture (IITA). The project's goal is to make biopesticides and biofertilizers more strictly regulated (Simiyu et al. 2013).

South Africa

The use, sale, and registration of biological control agents are governed by laws and regulations in South Africa. In accordance with Act 36 of 1947, the Department of Agriculture, Forestry, and Fisheries (DAFF) (www.daff.gov.za) regulates the registration of biological medicines (DAFF 2010).

European Union

In terms of the use and production of biopesticides, it is the second-largest continent. Microorganisms, plants, and pheromones were all governed under the EU's 1991 Directive 91/414/EEC, which was initially designed for chemical pesticides (Regnault-Roger et al. 2012). While new plant protection legislation was added in the EU in 2009, the following four pieces of legislation are also included: (1) Regulation (EC) No 1107/2009, (2) Directive 2009/128/ EC, (3) Directive 2009/127/EC, and (4) Regulation (EC) No 1185/2009. The Directive 91/414 was amended by 2001/36/EC (EC 2001) and 2005/25/EC (EC 2005) to add the specific requirements for microorganisms. As of 2011, all member states must abide by the new Regulation (EC) No. 1107/2009, which takes the place of Directive 91/414/EEC (Meeussen 2012). The registration of biopesticides in EU nations appears to be more challenging than elsewhere in the world because the dossier must be submitted along with results of environmental and toxicological testing, as well as an efficacy assessment. According to Regulation (EC) No. 1107/2009, product registrations are handled by three zones based on geographic and climatic factors (Hauschild 2012). Denmark, Estonia, Latvia, Lithuania, Finland, and Sweden are in Zone A (North); Belgium, Czech Republic, Germany, Ireland, Luxembourg, Hungary, the Netherlands, Austria, Poland, Romania, Slovenia, Slovakia, and the UK are in Zone B (Central); and Bulgaria, Spain, Greece, France, Italy, Cyprus, Malta, and Portugal are in Zone C (South). Plant protection product (PPP) applicants must submit their registration dossier to a "Zonal reporters member state" (zRMS), which reviews the dossier. Regulation (EC) No. 283/2013, which was recently adopted, implements Regulation (EC) No. 1107/2009 for establishing data-related concerns (EC 2013).

USSR (formerly)

The Russian Agricultural Control regulates the state registration of microbiological pesticides in Russia (RAC). In addition to managing pesticide registration, RAC oversees pesticide usage, manufacture, sale, transportation, storage, disposal, advertising, import, and export (Kabaluk et al. 2010). The Russian Agricultural Academy (RAN), which houses the All-Russian Institute for Plant Protection (VIZR) in St. Petersburg, is involved in the registration

procedure as well as research and development of biopesticides (Kabaluk et al. 2010).

United Kingdom

The Chemicals Regulatory Directorate (CRD)/Pesticide Safety Directorate (PSD) (http://www.hse.gov.uk/pesticides/) is the primary regulatory authority in the UK in charge of plant protection products, including biopesticides. Pesticides, biocides, detergents, and other chemicals are regulated by the CRD, a new Directorate of the Health and Safety Executive (HSE), in accordance with the Registration, Evaluation, Authorization and Restriction of Chemicals Act (REACH). Agricultural pesticide registration is handled by the Department of Environment, Food, and Rural Affairs (DEFRA) entity known as PSD (DEFRA 2006). The UK regulatory framework was created on a chemical pesticide model, which might have prevented the commercialization of biopesticides (ACP 2004). The biopesticide scheme was created in 2003 as a significant initiative, and its primary goal was to increase the production of biopesticides (http://www.pesticides.gov.uk/environment.asp). In order to register and regulate biopesticides, this approach introduced the position of "biopesticide champion" in 2006 (Chandler et al. 2011).

USA

A sizeable component of the worldwide biopesticide market is in the United States. According to USEPA (2010), the EPA in the USA has a comprehensive and complicated regulatory system for the registration and regulation of biopesticides, and this system has registration requirements that are different from those of other regulatory systems (Harman et al. 2010; Chandler et al. 2011). The Office of Chemical Safety and Pollution Prevention (OCSPP) and the Office of Pesticide Programs (OPP) are in charge of regulating biopesticides, and OPP is divided into three divisions that are involved in pesticide registration: the Antimicrobial Division, the Registration Division, and the Biopesticides and Pollution Prevention Division (BPPD) (Matthews 2014). EPA typically mandates the use of biopesticides since they pose fewer dangers than chemical pesticides.

The Insecticides Act of 1968, which was endorsed by the pesticide registration committee in India, set the rules for the registration of biopesticides. The research and commercialization of pest control solutions involves a number of stakeholders, including scientists, regulators, marketers, and end users. Although some members of this chain are frequently involved from the very beginning of the development process, there are still many problems to be solved. For example, marketers may frequently disagree with regulators and scientists, leaving end users perplexed about alleged flaws in the finished product (Damalas and Koutroubas 2018; Satapathy 2018).

Biopesticide Registration Protocol in India

Regulations for biopesticide registration and further marketing were framed during the 357th Meeting of CIB&RC held on 10th August, 2015. Important regulations are hereunder.

- 1. The earlier registrants of the strain/inventor of the strain has to deposit one sample containing at least one kg of product/formulation to the Secretary, CIBRC that should be subject to 16 SR-DNA/Gene code sequencing/finger printing for creating data bank of all the strains by ICAR-NBAIM, Mau.
- 2. Registration of already registered strains of biopesticides:
 - a. Requirement of the data/information to be submitted for getting permanent registration under section 9(3) /9(3B)
 - b. Form-I duly filled in along with requisite registration fee of Rs. 100 as per existing requirement.
 - c. Already approved Label leaflets of the product/strain
 - d. Testimonial/documents about the company as per existing requirement.
 - e. Undertaking about the strain from the inventor of the strain or first registrant or subsequent registrant of the strain or the applicant.
 - f. One sample (minimum one kg) for pre-registration verification (PRV) through Central Insecticides Laboratory
 - g. Another sample (minimum one kg) for pre-registration verification (PRV) of Gene code sequencing/16 SR-DNA/finger printing along with a demand draft (as per invoice obtained as testing fee from NBAIM, Mau) in favour of NBAIM, Mau as testing fee for Gene code sequencing/16 SR-DNA/finger printing.
- 3. Registration of new strain of the biopesticides:
 - a. The applicants for registration of new strain has to submit all the data as per existing guidelines for registration under section 9(3)/9(3B) for all the disciplines. Two samples have to be submitted to the Sectt. of CIB&RC; one for pre-registration verification (PRV) from Central Insecticides Laboratory as per product specification requirement & another sample to be used for pre-registration verification (PRV) of Gene code sequencing/16 SR-DNA/finger printing along with a demand draft (as per invoice obtained as testing fee from NBAIM,

Mau) in favour of NBAIM, Mau as testing fee for Gene code sequencing/16 SR-DNA/finger printing

- b. Minimum infrastructure required for production and registration of biopesticides:
- c. Verification of the infrastructure and technical competency of the applicants already registered under section 9(3B) and applying for registration u/s 9(3) and/9(3B) extension has to be conducted by a team constituted by the Secretary (CIB&RC) for the purpose.
- d. Minimum CFU count and nominal concentration strength of the formulation to be continued as per existing guidelines
- e. Verification of shelf life of strain and verification of product
- f. Submission of photographs for veracity of research, test and trails

Policies on Biopesticide Use

The National Agriculture Policy of India from 2000 placed a strong emphasis on farmers receiving timely and appropriate supplies of agricultural inputs, including biopesticides. In accordance with the "Zero Budget Natural Farming" (ZBNF) initiative put forth by the Food and Agriculture Organization of the United Nations (FAO), which promotes the use of locally obtainable natural fertilisers and biopesticides as well as farmer-owned seeds for organic farming, the Government of India has taken the necessary coordinated action (https://www.fao.org/agroecology/detail/en/c/443712/).

The marketing of biopesticides to farmers is the responsibility of the Ministry of Agriculture and Farmers Welfare and the Department of Biotechnology (DBT), in addition to the Central Integrated Pest Management Centre (CIPMC), Faridabad, the National Centre for IPM (NCIPM) under the Indian Agricultural Research Council, and the Directorate of Biological Control (Alam 1994). The Department of Biotechnology (DBT), in addition to the aforementioned regulatory bodies, funds research into the development of biopesticides (Sinha and Biswas 2008). Both the National Accreditation Board (NBA) and the National Agricultural Research System (NARS) conduct quality control testing on biopesticides and train state agricultural departments in these techniques.

The registration procedure for biopesticide products seems to be impeding their commercialisation. In order to enable quick registration of biopesticide products based on justifiable standards, regulatory agencies should encourage the use of safer technology in the creation of commercial products. Additionally, the regulatory framework should support the growth of small and mediumsized biopesticide companies, enabling them to provide consumers highquality products and giving growers trustworthy tools for the cost-effective control of pests. Data requirements for biological goods are frequently derived from those for synthetic chemical products. However, risk assessment for biopesticides ought to be based on pertinent scientific knowledge rather than synthetic chemical criteria. In order to reflect the nature of the various categories of biopesticide active ingredients, it is required to modify the standards. Data standards and instructions for biopesticides are now accurately modified (Isman 2014). The major problem for the biocontrol sector seems to be the length of submission procedures at both the EU and Member State levels. In order for new products to succeed on the market, faster processes and the enforcement of deadlines are essential. The high expense associated with registering new medicines is another barrier to the commercialization of novel products (Pavela 2014).

Legislation that prohibit the use of conventional pesticides like dicrotophos, azinphos ethyl, and ammonium sulphate, among others, that have been passed by governments of countries like India, Germany, and other European nations. These laws are projected to help the biopesticides market grow as a result of the extraction of pesticides from natural resources such as animals, plants, microbes, and particular minerals. It was predicted that the ban on a certain class of chemical pesticides might affect crop exports from India to other countries, particularly Europe. For instance, the Agricultural and Processed Food Products Export Development Authority between the United States and India has decided to end their collaboration as of 2020 APEDA. All organic businesses in India that seek to export to the US after July 2022 needed a certificate provided by a USDA-accredited certifier in order to export the generated crops. This requirement took effect after an 18-month transition period. For instance, the export amount of rice declined from 2018 to 2019 as a result of limits on specific chemicals, and it was projected that the use of biopesticides might assist rice growers increase the export volume in 2020. The usage of biopesticides was therefore expected to rise during the anticipated period as a result of rules regarding the use of chemicals for crop protection. (https://www.researchandmarkets.com/reports/5175605/india-biopesticidesmarket-growth-trends-and#rela2-5214644).

Limitations in Regulations of Biopesticide Registration

The primary issue with biopesticide regulation, which is a systemic one, is that it is based on the models used for traditional chemical pesticides (Greaves 2009). In the EU system, regulatory failure, according to Chandler et al. (2008), results from the use of an ineffective synthetic pesticide paradigm and

a lack of regulatory innovation. In addition, the evaluation of biopesticides and their registration for commercial use are also drawn-out processes. The sector complains that the current registration period is expensive and timeconsuming, notably for microbial biological control, and that the EU system takes a long time to process registrations (Bailey et al. 2010). As an illustration, the average time required in the EU was 75 months as opposed to 28 months in the USA (Hokkanen and Menzler-Hokkanen 2008). The US approach is flexible, and it occasionally invites applicants to pre-submission meetings where the applicant is informed on which investigations are required, based on available literature and preliminary data (Mubyana-Jhon and Taylor 2015).

Asia's biopesticide production system is underdeveloped and underutilised as a result of a number of institutional, social, and technical barriers that prevent the commercial production of innovative biopesticides (NAAS 2013). There is a difficulty with quality control in developing nations like Asia and Africa, which makes it difficult for farmers to have confidence in their products. Only an effective regulatory structure will be able to remedy this issue. Even though India is producing and using biopesticides, the increase is still lagging behind that of chemical pesticides. In a study, Rabindra (2005) projected that less than 10% of the identified need is being met by existing production of microbial pesticides. The CIB has registered around 500 biopesticides, which are available on the Indian market. However, quality control is a significant problem for the majority of the products (NAAS 2013).

Even though data requirements are becoming more transparent and standardised for more effective regulatory procedures, Mensink and Scheepmaker (2007) contend that insufficient guidance on the evaluation and use of biological products prevents premarket evaluations of the environmental safety from being carried out. It is difficult to establish an evaluation method that is equally fair to both biopesticides and chemical pesticides since regulatory authorities are aware that biopesticides are fundamentally different from chemical pesticides and should not be evaluated with the same standards of safety and efficacy (Bailey et al. 2010). One issue is that regulatory mechanisms only evaluate individual items, although the nature of microbial pesticides is extremely complicated and varied (Hubbard et al. 2014). In addition to providing guidelines, Ravensberg (2011) provided advice on how to compile a dossier and what sources to consult in order to better comprehend the exact requirements of the authorities. A data requirement is rather widespread in the USA and Canada, however the EPA does not call for a comprehensive dossier of efficacy and phytotoxicity data, while PMRA does.

The regulation of biopesticides with diverse modes of action is another complicated problem. For instance, Trichoderma species that are utilised as biopesticides against soil borne plant pathogenic fungi can parasitize such fungi in the soil; they can also create antibiotics (Ghisalberti and Sivasithamparam 1991; Vey et al. 2001) and enzymes that break down fungal cell walls (Bech et al. 2015). Trichoderma compete with soil borne pathogens for carbon, nitrogen, and other resources (Limon and Codon 2004). They can also encourage plant development by producing chemicals that are similar to auxin (Vinale et al. 2008; Nega 2014). Some Trichoderma products have been marketed as plant growth promoters rather than plant pesticides (Nega 2014), which has allowed them to avoid regulatory review of their effectiveness and safety (Bailey et al. 2010). Pseudomonas is in a same situation. Fluorescent Pseudomonas can be employed for both biocontrol and encouraging plant development (Negi et al. 2005; Mehnaz 2013; Tewari and Arora 2014). There aren't any specific regulatory controls in place to prevent this, though. To effectively use biopesticides, there are a number of technological and regulatory gaps that must be filled in order to reduce the use of chemical pesticides and to advance the use of biopesticides (Kumar 2015).

Interventions in Regulations of Biopesticide

In order to increase the production of agrobiologicals on a global scale, innovation in the current biopesticide control framework is essential (Arora et al. 2012). Currently, the regulatory environment differs by country; some have developed systems, some are making progress in their regulatory frameworks, and a few do not have adequate rules for biopesticides (Simiyu et al. 2013). In order to effectively control pests, the regulatory structure in place should be environmentally benign, scientifically sound, and technologically advanced (Greaves 2009). The issue of why biopesticides aren't utilised more frequently could be resolved by cutting registration fees and doing away with effectiveness requirements (Greaves 2009). Another sensible strategy for improving biopesticide regulation is for nations to enact laws on a worldwide scale by holding conferences, workshops, and meetings to raise the status of biopesticides (Mishra et al. 2015).

Exogenous pressure, such as government action, and endogenous pressure, such as pressure within regulatory organisations, are some variables that may encourage the necessary regulatory improvements (Greaves 2009). There is a need for guidelines to encourage the collaboration of businesses and research institutes because several institutions have conducted some preliminary research about the industrialization of biopesticides and institutional changes may be significant; however, no systematic reports have yet been published

(Leng et al. 2014). The innovative strategy for the manufacturing and marketing of biopesticides depends heavily on the global harmonisation of biopesticide regulatory rules, and the OECD is crucial for this harmonisation at the global level (Holm et al. 2005). The World Health Organization (WHO) and the OECD have an impact on pesticide control, and their participation is crucial (NAAS 2013). In order to assist its member nations in harmonising the methods and procedures used to analyse biological pesticides, the OECD project on biopesticides was launched in 1999 (Sigman 2005).

More than 70 emerging and transition economies have working links with the OECD, which now comprises 34 member countries (http://www.oecd. org/chemicalsafety/pesticides-biocides/). The OECD's working group on pesticides is made up of the I Registration Steering Group (RSG), (ii) Risk Reduction Steering Group (RRSG), and (iii) Biopesticides Steering Group (BPSG). Through the creation of working documents and guidance, the BPSG has made significant strides toward harmonisation and work sharing (Richards and Kearns 1997). The OECD group's headquarters are in Paris, France, and they work closely with EU governments to carefully examine the risks that biopesticides pose to people and the environment (http://www.biopesticideindustry alliance.org/).

The OECD, Food and Agriculture Organization (FAO), and EU have all focused their emphasis on pesticide control globally, in general and in specific (Greaves and Grant 2011; FAO 2012). An intergovernmental organisation in Paris called EPPO is financed by contributions from its member nations (www.eppo.int). The International Organization for the Control of Noxious Animals and Plants (IOBC) examined the rapid global spread of the use of microbial pesticides and improvements in their regulatory systems in 2010 (IOBC 2010). Various organisations, including the OECD, North American and European governments, have made significant strides toward promoting harmonisation for biopesticide legislation and facility developments for work sharing between governments (AGBR 2015). As the most significant international organisations for biopesticide regulation and innovation, the BPSG of the OECD, the FAO, the European Commission (EC), the IOBC, the EPPO, the North American Plant Protection Organization (NAPPO), and NAFTA have all worked together (AGBR 2015).

Conclusions

Worldwide, the commercialization of biopesticides is expanding quickly, however the increase is not proceeding as anticipated due to a lack of appropriate laws and other restrictions. Effective regulation can also stop fake biopesticides from being sold. The regulation of biopesticides is a barrier to their manufacturing and commercialization. The aforementioned explanation makes clear that different countries have different regulation criteria. It is suggested that a common regulatory system be developed to overcome this obstacle. In order to examine the potential dangers related to microbial biopesticides, it is also necessary to increase communication and information exchange between regulatory agencies, scientists, and enterprises. Guidelines for assessing the efficacy, quality, and field testing of biopesticides also need to be updated because they are frequently carried out by non-experts, particularly in poor nations. Establishing regulatory organisations is necessary to ensure quick registration of biopesticide products with justifiable regulations and open processes, as well as to support the adoption of new, safer technology in the creation of commercial products. The standard for regulating should be the same across all nations and should be based on the nature of agro-biologicals rather than chemical pesticides.

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Links

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Promotion of Biopesticides in India Role of Government and Growers

Abstract

Biopesticides are cutting-edge crop protection agents that shield crops from a wide range of pests and pathogens in an environmentally responsible way. They outperform synthetic pesticides in a wide range of ways, including target specificity, reduced toxicity, and biodegradability. Despite this, they are underrepresented in the crop protection industry, accounting for only 3.5 percent of the worldwide pesticides market. Biopesticides have a lot to offer for the development of sustainable agriculture, despite their slow adoption in the commercial pesticide industry. Understanding the main obstacles and constraints that affect the market for biopesticides can help in the development of innovative approaches including improving delivery systems, selecting new and improved strains, and preparing farmers and other stakeholders to deal with issues.

Keywords: Biopesticide, Promotion, Initiatives, Consumer awareness

Introduction

Pest and pathogen incidence is a natural occurrence that frequently goes unreported. However, they become a concern when their spectra expand and cause significant losses. Chemical pesticides are now commonly used in agricultural techniques to lessen the impact of such severe damages. Without a question, the use of chemical pesticides has put human health, ecological health, and sustainability at danger. Therefore, the use of biopesticides in pest management programmes has been recognised as a sustainable solution to free agriculture from the debt of disease occurrence and insect infestation. Increased organic farming areas and related efforts in India, such as SOM, NPOP, SMPMA, NMSA, PKVY, ZBNF, etc., are thought to support the market and use of biopesticides.

Government Initiatives

Due to the subsidy component/incentive on conventional pesticides, the current agro-industry is reticent to do research and produce biopesticides. However, due to restrictions on the broad use of chemical pesticides and the phasing out and banning of a few toxic substances, there has been an increased push in recent decades to develop biopesticides for commercial usage. The percentage share of biocontrol products is still considerably lower than that of chemicals, though. Policies such as entrepreneurial education, institutional finance availability, subsidies, insurance, and tax and duty exemption can all increase the production of biopesticides. Government support for the use of biopesticides and the designation of no-pesticide zones may help the situation for bioproducts. For example, the Sikkim Organic Mission (SOM), which converted about 75,000 hectares of agricultural land, is now India's first organic state as a result of more strictly enforcing the National Programme for Organic Production (NPOP) criteria connected to the organic mission. Examining the SOM model, it was discovered that in this situation, producers and authorities were urged to employ organic inputs while avoiding synthetic ones. The similar idea of becoming organic is also being tried in Kerala, Arunachal Pradesh, and Mizoram.

In order to advance the organic movement and lower chemical risk, the Ministry of Agriculture's Department of Agriculture & Cooperation introduced the Organic Farming Policy in 2005. The regulation recognised organic sources of nutrients such biofertilizers, organic manures, compost, and biocontrol agents as certified inputs for organic farming (biopesticides). The National Bank for Agriculture and Rural Development (NABARD) introduced the Strengthening and Modernizing Pest Management Approach in India (SMPMA) capital investment subsidy programme, which provided financial assistance for the establishment of bio-fertilizer/bio-pesticide units as a 25% subsidy up to a maximum of 4 million rupees.

The National Action Plan on Climate Change included the establishment of the National Mission for Sustainable Agriculture (NMSA), which dealt with "Sustainable Agriculture" issues (NAPCC). The third mission intervention of NMSA was related to pest management and aimed to promote biopesticide research, commercial manufacturing, and commercialization. The major objective was to develop new biopesticides and technology for disease prediction employing innovative botanical applications, sterile insect approaches, transgenic insects, semiochemicals, and endophytic microbial metabolites. Additionally, the "*Paramparagat Krishi Vikas Yojana*" (PKVY) and "Soil Health Management" (SHM) programmes have been launched to support organic farming by adopting organic communities through a cluster model and PGS certification (Reddy 2017). Farmers ensure their product is free of any synthetic chemicals, including fertilisers, pesticides, and hormones, under the self-regulatory PGS programme. A neighbourhood group of five or more organic farms supports the programme. The PGS Organic Council unifies the standards for production quality control and permits the use of its PGS label as a quality stamp on goods (https://www.pgsorganic.in).

In the last five years, the government has also taken the required steps to support the widespread use of biopesticides. The "Zero-Budget Farming" method, which has had considerable success in southern India, is already in use in a few other states across the country. Zero Budget Natural Farming (ZBNF), as the technique is known by the Food and Agriculture Organization of the United Nations (FAO), emphasises minimising the superfluous expenditure of agricultural inputs such the purchase of pricey seed, chemical fertilisers, and pesticides. Instead of such expensive machinery, it encourages the use of farmer-owned seeds, naturally occurring local fertilisers, and biopesticides for organic farming.

Consumer Awareness on Biopesticides

The inadequate usage of biopesticides is primarily the result of consumer and user ignorance (Arora et al. 2010). The word "biopesticides" is unfamiliar to many farmers, and others are unsure whether to use them instead of chemicals. Because of imprecise and unfavourable results, several people have stopped using biopesticides. However, since the production techniques utilised in such formulations do not meet the requirements specified by regulatory agencies, low-quality, non-registered products are particularly affected by this issue. It is imperative to stress that the proposed biopesticides should have reliable, repeatable, consistent, and focused activity (Mishra et al. 2015). The host range and circumstances under which the formulation will work should be clearly stated on the product. Knowing how farmers feel about biopesticides is important because they are the ultimate users of these products. This is because it helps determine the suggestions and needs for appropriate biological control measures in farming systems. However, there is a marked difference between small and large farmers in adopting biopesticides in practice.

Smaller farmers frequently ignore or disregard government initiatives and programmes pertaining to organic agriculture. Additionally, there are myths about biopesticide requirements such as higher costs, lesser yield, and other requirements. Furthermore, the illicit sale and usage of counterfeit goods is a serious problem that has caused farmers to lose faith in biopesticides; this calls for prompt government attention (FICCI 2015). By offering orientation and demonstration sessions where farmers may learn how to use quality products, private enterprises may also help to resolve this issue. For the same reason, farmer field schools (FFS), which offer field-based, location-specific instruction on biopesticides for the development of knowledge and confidence among the end-users, have been established in various states (Mohanty and Sahu 2019).

Despite being safer for the environment than synthetic pesticides, the use of biopesticides is far lower than that of synthetic pesticides due to lack of information, lack of trust, and unavailability in local markets. In order for governmental, non-governmental, and educational institutions to properly apply biopesticides, farmers must be educated about the advantages of doing so through on-farm training. The government should offer the biopesticides as a subsidy or for free when purchasing agricultural products to promote their use by farmers.

Conclusions

Although biopesticides have demonstrated their value in the sustainable management of pests and pathogens, they are currently a niche product in the crop protection market. The primary causes of the market for biopesticides still being in its infancy are farmers' lack of confidence as a result of their long-standing reliance on chemical pesticides for crop protection, their lack of awareness, poor government support, a lax regulatory system, inappropriate technologies, and a lack of knowledge. Governmental, non-governmental, corporate, and public institutions, as well as universities, must all take a holistic approach to meet these difficulties.

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Growth of Biopesticides Driving Force and Set-Back

Abstract

The toxicity and non-biodegradability of chemical pesticides have stoked the demand for more sustainable alternatives. In addition to this, the persistently rising demand for cost-effective pest control measures has increased the utilization of biopesticides across several countries. However, their use has remained low in certain under-developed nations but are expected to witness better growth in the coming years. India offers a wide range of options in terms of supplies for natural biological control organisms as well as natural plant-based insecticides because of its great biodiversity. The widely diverse indigenous tribes in India's rich traditional knowledge base may hold important hints for the development of more advanced and efficient biopesticide. The National Farmer Policy of 2007 aggressively encouraged the adoption of biopesticides to boost agricultural output while maintaining farmer and environmental health. Additionally, it states that biopesticides will receive the same funding and promotion as chemical pesticides. Biopesticides are yet to take off in a major way in India because of mixed constraints, despite their enormous market potential and the national and state initiatives to promote them as alternatives to chemical pesticides. This chapter seeks to examine the factors enabling growth in the market as well as those restraining its trajectory.

Keywords: Biopesticide, Adaptation, Driving force, Demerits

Introduction

Regulations should make it easier to utilise creative, long-lasting solutions, allowing for the selection of the most environmentally friendly pest management method. This can be accomplished by using expedited registration, priority registration, and a combination of comparative evaluation of pest control techniques and the substitution principle, which allows a natural pest control

technique to take the place of a synthetic pesticide. More microbial biological control agents will be registered more quickly as a result of the modifications in registration procedures, which will logically lead to lower product costs (EC 2009; van Lenteren et al. 2018).

Another significant step toward making biological control more appealing and available to farmers is the development of a standardised process for the registration of microbial biological control agents that may be used locally or globally. Use of biopesticides is prompted by the removal of pesticides from the market as a result of observed health, non-target, and environmental effects, the emergence of new pests for which no pesticides are available, the development of resistance that reduces the effectiveness of pesticides, and all stimulate use of biopesticides (Urbaneja et al. 2012). Non-governmental organisations (NGOs) have had success switching from chemical to biological control in a number of instances by providing information on the impacts of pesticides on the environment and their illegitimate usage (Calvo et al. 2012).

A rise in the use of biological control has also been attributed to the development of new and improved biological control methods, improved and more stable formulations for microbial biological control agents and their use as seed treatments, more practical application techniques for invertebrate biological control agents (equipment to release biological control agents in crops, use of drones, etc.), and steadily more stable formulations of microbial biological control agents. It's interesting to note that growers quickly adopted the additional information and techniques needed to make biological control effective, and in many cases they developed new ideas and technology to enhance the release and establishment of invertebrate biological control agents. Additionally, they inspired scientists and the biological control sector to develop fresh invertebrate biological control techniques for newly emergent pests. When farmer organisations recognise the various benefits of biopesticides, including their economics, crop protection will undergo a new renaissance. They should take a far more proactive stance and demand expedited registration of cutting-edge sustainable control technologies in order to protect their own interests.

The market for biological control would significantly expand if the "real cost" theory were applied to chemical pesticides. Governments support the use of pesticides since the industry is not held accountable for human illnesses and deaths brought on by prolonged exposure to pesticides, nor is it required to pay for the cost of repairing environmental harm. As a result, costs associated with pesticides that have negative effects on human health and the environment are externalised and paid for by society, which is unethical and unscrupulous

because the pesticide industry only benefits financially from these costs while bearing none of the responsibility. In the past, pesticides' profitability was in fact overstated. Chemical pesticide costs would increase significantly with realistic pricing that took into account true costs, and non-chemical alternative controls would face fairer competition. Despite the fact that there have been known hidden costs associated with pesticides since the 1980s, prices of pesticides have rarely increased as a result. Applying levies on synthetic pesticides would be a first step toward true cost pricing because it would result in higher, more accurate costs for these products as well as more competitive pricing for the biological control agents employed in IPM programmes. (Pimentel and Burgess 2014; Bourguet and Guillemaud 2016; https://www. fortunebusinessinsights.com/thoughtleadership/biopesticides-trend-9099).

Supporting Points for Biopesticide Growth

- 1. Because of the rigorous battery of tests required for commercialization, some promising strains created by publicly supported research organisations in India are essentially confined to the shelf
- 2. DNA bar-coding for precise identification of the species to be included in the creation of biopesticides before their field applications.
- 3. For the licensing and marketing of biopesticides in India, a comprehensive federal action plan, realistic budget, and efficient administrative procedures are required.
- 4. Farmers should receive sufficient training on using biopesticides in order to reap the greatest benefits.
- 5. The main drivers of the market expansion for biopesticides are the rise in demand for high-quality food, average entry hurdles, ecological imbalances, changing markets in developing nations, and ecological and health concerns for people, animals, and plants.
- 6. Environmental safety awareness raising, increased demand for chemical-free and environmentally friendly farming products, new product introductions, enhanced scientific validity of biopesticides, strict regulatory pressure, increased demand for organic products, and higher user confidence
- 7. Due to the efforts of Government of India programmes like *Paramparagat Krishi Vikas Yojana* (PKVY), Mission Organic Value Chain Development for North Eastern Region (MOVCDNER), and National Programme of Organic Production (NPOP), the area under organic cultivation increased from 1.5 million ha in 2016 to 1.9 million ha in 2018. A total of 1.35 million metric tonnes of organic food

were produced in the nation in 2016 by 0.65 million organic farmers. The biopesticide market in India is being driven by a sizable organic farming sector, and this trend is expected to continue (https://www.researchandmarkets.com/reports/5175605/india-biopesticides-market-growth-trends-and#rela2-5214644).

- 8. The 'Green Revolution' is progressively giving way to the 'Ever-green Revolution', especially in countries with strong agricultural foundations. Due to their eco-friendly, economical, farmer- and consumer-friendly qualities, the role of agri-bio inputs will thus be vital in fostering this shift. Additionally, it is anticipated that growing consumer demand for foods (organic food products) free of synthetic pesticides would further fuel market expansion.
- 9. Growers are becoming more and more eager to try biological solutions when conventional synthetic alternatives lose their effectiveness as a result of biotic stressors acquiring resistance.
- 10. The prohibitively high costs of developing synthetic crop protection chemistries are another factor driving the development of biopesticides. According to tech developers, the development and approval of a novel synthetic pesticide typically takes \$250 million and nine years, whereas a biopesticide requires less than \$10 million and four years.
- 11. The proliferation of start-ups in the biopesticide industry has produced a highly competitive and cutting-edge environment for advancements. Synthetic pesticides have not seen as much recent release as innovative biopesticide components.
- 12. The Central Insecticide Board and Registration Committee, GOI, provided straight forward and quick regulatory approval for the registration of biopesticides.
- 13. The government of India banned 18 active chemical compounds in response to growing environmental concerns and household awareness of food safety.
- 14. A lot of Indian export goods don't match the required minimum residue level
- 15. Biopesticide usage is scaled up owing to government support and increasing awareness about the use of non-toxic, environment-friendly pesticides.
- 16. A long-term collaboration agreement for the sale of biological products from Bioworks Inc., including biopesticides, in India and South Asia.

- 17. The market is primarily driven by the issue of chemical pesticide residues, the appeal of organic farming, environmental concerns, and the simple registration procedure.
- 18. New applications for biocontrol, such as nanotechnology, RNAi, etc. that are not achievable with synthetic crop protection are what are causing the expansion of biopesticides.
- 19. Concrete proof of biopesticides' effectiveness in reducing crop damage and the resulting rise in crop yield
- 20. Affordable, high-quality items are readily available.
- 21. Supply chain management needs to be improved in order to use biopesticides more frequently. An effective distribution mechanism for biopesticides from the plant where they are produced to the farm where they are used is crucial in this regard.

Factors Restrain Biopesticide Market

- 1. Due to federal and state initiatives, the demand for biopesticides has increased, which has resulted in "driving the marketing of fake biopesticides."
- 2. The limited production of biopesticides with biopesticidal formulations, registered under the 1968 Insecticide Act
- 3. The expense and lengthy licensing process for biopesticides in India discourage businesses from investing in the study and development of biological pesticides.
- 4. Before registering and propagating biopesticides, it is necessary to confirm the microorganism's bio-safety. In order to do rigorous safety and allergy tests, many universities and research institutes who conduct the original research and create biopesticides are unable to cover the additional costs. For instance, immune-compromised people have reported allergies to various fungi, such as *Trichoderma, Metarhizium, Anisopliae*, and *Beauveria*.
- 5. They cannot be employed against a variety of pests since they are targetspecific, which is a limitation.
- 6. Biopesticides' effectiveness varies from climate to climate and is also extremely dose-dependent.
- 7. Due to the wide variations in the active and related substances of the parent plants in different agro-climatic zones, it is frequently challenging to make pure botanical pesticides, in contrast to synthetic pesticides, which can be made in desired purity and yield. Their physical and

chemical characteristics, as well as toxicological and other relevant features, change as a result. Their contamination by physical, chemical, or microorganisms also makes things more difficult.

- 8. The main market restrictions are farmers' lack of awareness and biopesticides' expensive price.
- 9. The market's expansion may be constrained by the lower acceptance rate of biopesticides than that of chemical agri-inputs.
- 10. Key heavyweights are strongly represented in the traditional and conventional agri-inputs sector, which is well-structured globally. But the biopesticides business is characterised by a number of start-ups that are having trouble getting enough money, building the right infrastructure, and getting traction with customers.
- 11. Many smaller developers may find it difficult to compete with established, potent synthetic pesticides, both in terms of proving the effectiveness of biopesticides and, more crucially, in persuading producers to switch from their tried-and-true ways to new and somewhat unproven products.
- 12. In addition, several types of biopesticides, particularly those produced from genetic materials or crop diseases, face unknown regulatory approval paths; this increases the difficulty of licensing and commercialization and impedes the innovation and development of biopesticides.
- 13. The bulk of biocontrol strategies necessitate repeated, frequent treatments for best results. These applications require more work and money, which sometimes acts as a barrier and stunts the development of biopesticides.
- 14. The research and development (R&D) of biopesticides carried out by small businesses consistently fails to understand the demand dynamics of a given location, which can further impede market growth.
- 15. The development of biopesticides is a high-risk business since it requires an initial large capital outlay to choose prospective strains for sales, as well as packaging, storage, and distribution.
- 16. Aside from the aforementioned problems, the single biggest barrier to the development and growth of biopesticides is the widespread selling of substandard (low CFU count), fake (no CFU count products) (Alam 1995). and misbranded (pesticide-laced bioproducts-pseudo-biopesticides) biopesticides (Keswani et al. 2016). APEDA (Ministry of Commerce)certified organic bio-inputs supplied under the pretence of uncontrolled organic bio-inputs (not permitted by CIBRC) also constitute a severe threat to high-quality biopesticides. The organic bio-input products are not put through any bio-efficacy/safety experiments as required by

CIBRC. These categories account for over 65% of overall biopesticide sales (Singh and Arora 2016). To make biopesticides an effective tool for IPM/Sustainable Agriculture, the agriculture departments should strictly enforce the licensing requirements and quality controls for them.

- 17. The high prices for biopesticide registration (http://ppqs.gov.in/ divisions/cib-rc/guidelines) are another barrier to advancing research, development, and usage of biopesticides in IPM/Sustainable Agriculture. Relaxed rules for CIBRC registration should be drafted in order to enable the registration of many biopesticides (GHS). The main barrier to promoting the use of biopesticides, biofertilizers, and botanicals is the imposition of a 12 percent Goods and Services Tax (GST) on microbial goods as well as on the botanical product (neem) (the same GST rate as for toxic/hazardous conventional chemical pesticides).
- 18. Farmers have a lot of concerns about the short shelf life of biopesticides. Because live bacteria make up the majority of biopesticides, changes in temperature, humidity, or even exposure to ultraviolet radiation reduce their effectiveness (Arora et al. 2016). Additionally, contamination may significantly lower the product's microbial count, greatly decreasing its efficacy in real-world settings (Alam 2000; Evans et al. 1993). Due to a shortage of money for the next steps, everything stalls before: adherence to regulatory requirements, scalability for application and delivery, marketing, and commercialization.
- 19. The rapid emergence of the corona virus pandemic has had an impact on the world market as countries have implemented lockdown measures and restricted public movement. These activities are having a substantial impact on the manufacturing of biopesticides, as firms are experiencing supply chain disruptions, a shortage of raw materials, forced plant closures, and a lack of staff.

Conclusions

Long-lasting shelf of biopesticide formulations, DNA-bar-coding for precise identification of organism, comprehensive federal action plan, training and awareness, rise in demand for high quality food, strict regulatory pressure, increase of area under organic farming, Green revolution to Ever-green revolution, proliferation of start-ups of biopesticide units, quick regulatory approval for registration by CIB&RC, New Delhi, India, withdrawal of dangerous chemical pesticides, negative consequences of Green revolution, failure of novel applications such as nanotechnology, RNAi etc., in development synthetic plant protection solutions etc., are considered as few driving forces to boost the growth of biopesticide in Indian agriculture. The USEPA has implemented a number of measures for the quick expansion of the biopesticide market, including appropriate changes to the registration process for the quick commercialization of biopesticide formulations, real cost policy, stakeholder perspectives, manufacturer and dealer of biopesticides, etc. Sustainable agriculture is referred to as "Conscious Agriculture" and is positioned between contemporary agriculture and conventional agriculture. All parties involved in the production and consumption processes must participate in conscious agriculture, which also protects the environment and the availability of resources for future generations. According to a position paper Fresco and Poppe recently issued, conscious agriculture is smoothly integrated into a "shared agricultural and food policy" (2016).

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Microbial Biopesticides in India

According to estimates, plant pests and diseases cause at least 10% of the world's food production to be lost. Additional pest and disease losses occur after harvest, where it is estimated that up to 13% of the total calories generated are lost after leaving the farm gate, in addition to in-field losses. According to Indian viewpoints, biotic stressors cause a 20-26% annual loss of food commodities. The effects of the "Green Revolution" (GV) multiplied crop production and productivity, which ultimately affected the livelihoods of Indian civilians. However, the introduction of numerous inorganic plant protection inputs resulted in irreversible harm to the environment, human lives, and material wealth. The focus on environmentally friendly plant protection measures, particularly microbial biopesticides, has expanded as a result of current conditions and Indian government regulations on chemical pesticides that take into account their negative effects. In the recent past, India's biopesticide industries, demand, consumption, market, etc., have grown tremendously. It is expected that between 2040 and 2050, the biopesticide industry would either match or surpass those of chemical pesticides. Global population growth is projected to reach 9 billion people by 2050, and the duty to increase food production by 70% of current levels in order to feed this population has led to a ruthless shift toward the use of artificial agricultural inputs. Nevertheless, there are ways to enhance food production to the anticipated level by using Integrated Pest Management, which enables the prudent application of inorganic inputs in conjunction with environmentally benign tactics like microbial biopesticides.

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