Aflatoxins in Cereals and Cereal-Based Products: Occurrence, Toxicity, Impact on Human Health, and Their Detoxification and Management Strategies

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Abstract: Cereals and cereal-based products are primary sources of nutrition across the world. However, contamination of these foods with aflatoxins (AFs), secondary metabolites produced by several fungal species, has raised serious concerns. AF generation in innate substrates is influenced by several parameters, including the substrate type, fungus species, moisture content, minerals, humidity, temperature, and physical injury to the kernels. Consumption of AF-contaminated cereals and cereal-based products can lead to both acute and chronic health issues related to physical and mental maturity, reproduction, and the nervous system. Therefore, the precise detection methods, detoxification, and management strategies of AFs in cereal and cereal-based products are crucial for food safety as well as consumer health. Hence, this review provides a brief overview of the occurrence, chemical characteristics, biosynthetic processes, health hazards, and detection techniques of AFs, along with a focus on detoxification and management strategies that could be implemented for food safety and security.
Keywords: aflatoxins; food contamination; health issues; detection technique; conventional and novel management strategies

Key Contribution: Aflatoxin contamination of cereals and cereal-based products poses a serious concern for human health worldwide. This review highlights the occurrence; chemistry and biosynthesis; and health effects of aflatoxins, with a major focus on detection and mitigation strategies to ensure food safety and security.

1. Introduction

Crops, mainly cereal grains, serve as a major source of energy and nutrition in the human diet worldwide. Cereals are generally consumed as raw or cooked grains or in the form of processed products such as flour, semolina, bread, and cookies. Often, cereal crops and their byproducts are also used as animal feed for livestock and poultry, which are eventually rendered as sources of dairy, poultry, and meat products for human consumption. According to the Food and Agriculture Organization (FAO), one-fourth of the world’s total cereal crop production is contaminated with mycotoxins [1,2]. In the last few decades, the number of cases of mycotoxicosis in humans has increased due to the consumption of food contaminated with one or more mycotoxins, which has ultimately also affected the sustainability of agribusinesses [3–5]. The acute toxicity and carcinogenic effects of mycotoxins and their contamination in cereals and cereal-based products pose a serious food safety and security concern for humans, along with huge economic losses [6].

Mycotoxins, such as fumonisins, ochratoxin A, trichothecenes (type A and B), aflatoxins, and patulin, are secondary metabolites secreted by several fungal species, mainly Aspergillus, Fusarium, Penicillium, and Alternaria, which often contaminate cereal crops at the farm level during handling, transportation, and/or storage [6–14]. Most mycotoxins are resistant to food processing techniques due to their thermostable nature; hence, their contamination in processed products must be checked before consumption. According to Khaneghah et al. [15], total aflatoxin (AF) is the most prevalent mycotoxin contamination worldwide after ochratoxin, zearalenone, and deoxynivalenol, found in cereals and cereal products such as whole grains, bread, cornflakes, breakfast cereals, and pasta products. As secondary metabolites of fungal strains (Aspergillus parasiticus, Aspergillus flavus, and Aspergillus nomius), AFs (B1, B2, G1, and G2) develop on a variety of food and feed materials during growing, harvesting, storing, and shipping processes [16,17]. Out of 20 known AFs, 4 (B1, B2, G1, and G2) have been identified as major contaminants in cereals, such as peanuts, maize, rice, barley, and sorghum, and in their products [18]. AFB1 has the highest prevalence in cereal products of all AFs [19]. Under UV light (365 nm), aflatoxins emit blue (B1 and B2) or green (G1) and green-blue (G2) fluorescence [20]. These different groups of aflatoxins differ in their structures at the molecular level; for instance, the cyclopentane ring is identical in group B aflatoxins (B1 and B2) and group M aflatoxins (M1 and M2), while group G aflatoxins (G1 and G2) contain a lactone ring (Figure 1). Based on toxicity, aflatoxin types can be arranged as B1 > G1 > B2 > G2 [21]. Andrade and Caldas [22] reported that 37.6% of 18,097 tested cereal samples worldwide were contaminated with at least one AF, according to 89 publications.
The World Health Organization (WHO) has recognized AFs as a global food safety concern, and rural populations in developing countries are at an especially high risk of AF exposure [23]. The International Agency for Research on Cancer (IARC) classified AFs as group 1 carcinogens due to their toxic, carcinogenic, mutagenic, teratogenic, and immunotoxic nature [24,25]. Due to serious health complications in humans and animals, several countries have implemented strict regulations to prevent AF contamination in food and feed. According to the European Commission Regulation, the maximum permissible limit for total AFs and AFB1 in all cereals and their derived products intended for direct consumption should not be more than 4 µg/kg and 2 µg/kg, respectively [26,27]. However, 20 µg/kg is the maximum acceptable limit for AFs in the United States [28]. In some countries (such as in the EU), there are limits for raw cereals and processed products. Apart from limiting the maximum acceptable limit, several innovative techniques and management practices are also adopted at pre- and post-harvest processing levels to control and/or prevent aflatoxin contamination in cereals and their derived products. Though several reports and publications are available on control and management strategies for aflatoxin contamination in food and feed, a gap exists in the literature with a focus on novel and environmentally friendly approaches. Hence, the present review aims to provide an overview of AF contamination in cereals and cereal-based products and suggests the best environmentally friendly practices that could be implemented to ensure food safety as well as prevent possible AF outbreaks.

2. Major Source and Occurrence of Aflatoxins

AFs are one of the major mycotoxins produced in cereals by several species of *Aspergillus*, mainly *A. flavus*, *A. nomius*, *A. parasiticus*, and *A. astellatus*. Other AF-producing species, such as *A. bombycis*, *A. ochraceoroseus*, and *A. pseudotamarii*, have been identified using advanced genome sequencing techniques [29]. The warm and humid environment of tropical and subtropical regions is favorable for the growth of these fungal species [30]. Out of different AF types, B1, B2, G1, and G2 are found in plant-based foods, including cereal grains, while the metabolites of type AFB1, i.e., AF M1 and M2, are especially found in foods of animal origin [31]. *Aspergillus* species such as *A. flavus* and *A. pseudotamarii* are mainly responsible for the production of type B AFs only, as they cannot produce type G aflatoxins due to the absence of 0.8 to 1.5 kb in the 28-gene cluster responsible for AF biosynthesis. However, other *Aspergillus* species, such as *A. parasiticus*, *A. nomius*, and *A. bombycis*, are capable of producing all four major AFs. AFs M1 and M2 are the hydrated metabolites of AFs B1 and B2, respectively, and their contamination is usually observed in products derived from animals when exposed to feed contaminated with AF B1 and/or...
B2. The presence of AF M1 at a higher concentration has been reported in human breast milk from countries such as Australia and Thailand, which shows the risk of aflatoxicosis in infants [32].

Among cereals, AF contamination is frequently observed in crops such as rice and corn compared to other cereals [33]. AFB1 contamination in rice has been reported in several countries, including China, Egypt, India, Iran, Malaysia, Nepal, Pakistan, the Philippines, the United Kingdom, and the United States [34]. The improper drying of rice grains, when the moisture content is >14%, is mainly responsible for fungal growth, which later causes the discoloration of grains and/or husks, and the production of toxic secondary metabolites, such as AF, and ultimately leads to the complete deterioration of edible-grain quality [2]. Climate changes, including temperature, moisture content, water activity (aw), type of soil, and storage conditions, are major factors influencing fungal growth and their ability to produce AFs in cereals crops [35,36]. Lv et al. [37] reported that the maximum AFB1 production occurs at a temperature of 33 °C and water activity (aw) of 0.96, whereas Gizachew et al. [38] reported that temperatures ranging from 28–37 °C at 0.92–0.96 aw led to the optimal growth of fungi (A. flavus and A. parasiticus) on polished rice. According to Battilani et al. [39], every 2 °C increase in temperature as a result of climate change could increase the emergence of AFB1 in various regions of Europe, such as Albania, Bulgaria, Cyprus, Greece, Italy Spain, Portugal, and Turkey. Furthermore, in the next 30 years, the risk of AF contamination in maize crops is expected to rise in Europe due to changing climatic conditions that are favorable for AF-producing fungi such as A. flavus [40]. Further, the type of AFs detected in various food sources around the world between 2010 and 2022 and the method of detection using various techniques are presented in Table 1. Aflatoxins have been found in a variety of cereals and their products, including barley-based products, corn, corn bran, corn flour, corn ingredients, corn-based opaque beers, multigrain-cereal baby foods, pearl millet, rice, rice-based baby foods, rice flour, sorghum, sorghum beer, sorghum malt, sorghum-based products, wheat and wheat-based baby foods, wheat bran, wheat flour, and wheat-based products. Various research studies on aflatoxins in cereals and their byproducts have been conducted, and they were detected in nearly every country, such as Africa, Bangladesh, Brazil, Burkina Faso and Mozambique, China, Colombia, Costa Rica, Egypt, Ethiopia, Ghana, India, Iran, Kenya, Mediterranean area, Namibia, Niger, Pakistan, Peru, Serbia, South Africa, South Korea, Spain, Tanzania, Tanzania, Thailand, Togo, Tunisia, Turkey, Uganda, Vietnam, and Zimbabwe.
Table 1. Occurrence of aflatoxins in cereals and cereal-based products around the world.

| Food Matrix               | Country                  | No. of Samples | Aflatoxin | Range (µg/kg) | Limit of Detection (LOD, µg/kg) | Detection Technique | References |
|---------------------------|--------------------------|----------------|-----------|---------------|-------------------------------|---------------------|------------|
| Barley-based products     | Mediterranean area       | 1/4            | AFB1      | 24            | 0.25                          | LC-MS/MS            | [41]       |
| Burkina Faso and Mozambique | Burkina Faso and Mozambique | 13/26         | AFB1      | 3.4–636       | 3.0                           | LC-MS/MS            | [42]       |
|                           | India                    | 28/150         | AFB1      | 48–383        | 3.9                           | HPLC                | [43]       |
|                           | Tanzania                  | 60             | AFB1      | 3–1081        | 0.6                           | UPLC/TOFMS          | [44]       |
|                           | Tanzania                  | 60             | AFB2      | 12–177        | 0.4                           | UPLC/TOFMS          | [44]       |
|                           | Zimbabwe                  | 95             | AFB1      | 0–11          | 3.75                          | LC-MS/MS            | [45]       |
|                           | Zimbabwe                  | 95             | AFB1      | 0–3           | 1.75                          | LC-MS/MS            | [45]       |
|                           | Zimbabwe                  | 80/388         | AFB1      | 0.57–26.6     | 0.005                         | HPLC                | [46]       |
|                           | China                     | 108            | AFB1      | 0.4–136.8     | 0.1                           | HPLC                | [47]       |
| Corn                      | Costa Rica                | 108/970        | Total AFs | 0–290.4       | 0.01 & 3                      | ELISA and HPLC      | [48]       |
|                           | Iran                      | 30             | AFB1      | 6.25–1060     | 2                             | UHPLC-MS/MS         | [49]       |
| Corn ingredients          | South Africa              | 2/32           | AFB1      | 0–7           | 2.5                           | LC-MS               | [50]       |
|                           | Africa                    | 233            | AFB1      | 19.2–1137.4   | NA                            | ELISA               | [51]       |
|                           | Colombia                  | 3/20           | Total AFs | 8.2–588.9     | 5                             | LC-MS/MS            | [52]       |
|                           | Turkey                    | 38/1055        | Total AFs | 7.96–163.62   | 5                             | LC-MS/MS            | [53]       |
|                           | Ethiopia                  | NA             | Total AFs | 20–91.04      | NA                            | HPLC                | [54]       |
|                           | Brazil                    | 38/148         | Total AFs | 0.4–49.9      | NA                            | LC-MS/MS            | [55]       |
|                           | South Korea               | 507            | AFB1      | 1–5.2         | 0.1                           | LC/MS/MS            | [56]       |
|                           | Vietnam                   | 1486/2370      | AFB1      | 2–5           | 34.8                          | ELISA               | [57]       |
| Corn                      | Niger and Benin           | 112            | Total AFs | 0–3000        | NA                            | ELISA               | [58]       |
|                           | China                     | 44             | AFB1      | 0–148.4       | 1                             | HPLC                | [59]       |
|                           | Pakistan                  | 72             | Total AFs | 0.5–10        | 0.5                           | HPLC                | [60]       |
|                           | Ghana                     | 326            | Total AFs | 0–341         | 0.1                           | TLC                 | [61]       |
|                           | Peru                      | 82             | Total AFs | 1–17          | 0.4                           | LC-MS/MS            | [62]       |
|                           | Uganda                    | 256            | Total AFs | 0–3760        | NA                            | HPLC                | [63]       |
|                           | Togo                      | 70             | AFB1      | 1.1–75.9      | 0.08                          | HPLC                | [64]       |
|                           | Ghana                     | 70/90          | AFB1      | 0.78–339.3    | 0.13                          | HPLC                | [65]       |
| Corn flour                | Serbia                    | 27/56          | Total AFs | 1–9.14        | 0.4                           | HPLC-FD             | [66]       |
| Corn flour                | Turkey                    | 24             | AFB1      | 0.041–1.12    | 0.026                         | HPLC                | [67]       |
| Corn bran                 | Uganda                    | 40             | Total AFs | 7.5–393.5     | 1                             | HPLC                | [68]       |
| Corn bran                 | Tanzania                  | 340            | Total AFs | 9.4–16.8      | NA                            | ELISA               | [69]       |
| Pearl millet              | South Korea               | 507            | AFB1      | 1–1.1         | 0.1                           | LC/MS/MS            | [56]       |
| Pearl millet              | Kenya                     | 86             | AFB1      | 0.4–5.6       | NA                            | ELISA               | [51]       |
| Pearl millet              | Tunisia                   | 220            | AFB1      | 117–1046      | 0.24                          | LC-MS/MS            | [70]       |
| Food Matrix        | Country          | No. of Samples | Aflatoxin | Range (µg/kg) | Limit of Detection (LOD, µg/kg) | Detection Technique | References |
|-------------------|------------------|----------------|-----------|---------------|----------------------------------|---------------------|------------|
| Pearl millet      | Tunisia          | 220            | AFB2      | 0–96.1        | 0.4                              | LC-MS/MS            | [70]       |
|                   | China            | 29             | AFB1      | 0.1–1.4       | 0.1                              | HPLC                | [47]       |
|                   | Pakistan 88/120  | Total AFs      | 1.18–11.46| 0.4           | TLC                             | [71]               |
|                   | Pakistan 100/120 | Total AFs      | 0.21–11.89| 0.142         | HPLC                            | [71]               |
|                   | Pakistan 104/120 | Total AFs      | 0.10–12.39| 0.092         | LC–MS/MS                        | [71]               |
|                   | Pakistan 88/120  | Total AFs      | 1.24–11.68| 1.0           | Detection Technique              |                     |
|                   | Nigeria 38       | AFB1           | 3.7–20.2  | 0.15          | LC-MS/MS                        | [72]               |
|                   | Nigeria 38       | AFB2           | 1.32–6.11 | 0.2           | Yellowish-greenish               | [72]               |
|                   | Nigeria 38       | AFG1           | 3.76–7.21 | 0.2           | LC-MS/MS                        | [72]               |
|                   | Iran 40          | AFB1           | 0.29–2.92 | NA            | ELISA                           | [73]               |
|                   | China 235/370    | AFB1           | 0.03–20   | 0.65          | HPLC                            | [74]               |
|                   | China 235/370    | AFB1           | 0.0–1.6   | 0.15          | HPLC                            | [74]               |
|                   | Pakistan 2047    | AFB1           | 1.17–6.91 | 1             | TLC                             | [75]               |
|                   | Bangladesh 227   | AFB1           | 0–0.9     | 0.2           | HPLC                            | [76]               |
|                   | Thailand 240     | AFB1           | 1.43–26.61| 0.093         | HPLC-FD                         | [77]               |
|                   | India 2/87       | Total AFs      | 21.58–22.98| NA            | TLC                             | [78]               |
|                   | Egypt            | 51             | AFB1      | 100–200       | NA                              | TLC                 | [79]       |
|                   | Colombia 3/24    | Total AFs      | 0.2–23.9 | 5             | LC-MS/MS                        | [52]               |
|                   | South Korea 507  | AFB1           | 1–1.1     | 0.1           | LC/MS/MS                        | [56]               |
|                   | Mediterranean area| AFB1         | 26.0–33.0 | 0.25         | LC-MS/MS                        | [41]               |
|                   | Mediterranean area| AFB2         | 7.5       | 1.5           | LC-MS/MS                        | [41]               |
| Rice flour        | Serbia 2/6       | Total AFs      | 1.59–4.76 | 0.4           | HPLC-FD                         | [66]               |
| Rice flour        | Turkey 16        | AFB1           | 0–0.029   | 0.026         | HPLC                            | [67]               |
| Rice-based baby foods | Iran 27/30  | AFB1           | 0–15.15   | 0.025         | HPLC-FD                         | [80]               |
| Sorghum malt (Omalodu) | Ethiopia 90 | AFB1          | 0–33.10   | 0.01–0.03     | ELISA                           | [81]               |
|                   | Nigeria 19/35    | Total AFs      | 0.96–21.74| 1             | TLC                             | [82]               |
|                   | India 15/21      | AFB1           | 0.005–0.02| NA            | TLC                             | [83]               |
|                   | India 3/21       | AFB2           | 0–0.005   | NA            | TLC                             | [83]               |
|                   | South Korea 507  | AFB1           | 0.7–1.7   | 0.1           | LC/MS/MS                        | [56]               |
|                   | Namibia 45       | AFB1           | 0.61–28.3 | 0.17          | LC/MS/MS                        | [84]               |
|                   | Namibia 45       | AFB2           | 0.14–2.35 | 0.04          | LC/MS/MS                        | [84]               |
|                   | Namibia 45       | AFG1           | 0.39–6.95 | 0.1           | LC/MS/MS                        | [84]               |
|                   | Burkina Faso 20  | AFB1           | 46.33–254.73| 0.2          | HPLC                            | [85]               |
Table 1. Cont.

| Food Matrix                  | Country         | No. of Samples | Aflatoxin | Range (µg/kg) | Limit of Detection (LOD, µg/kg) | Detection Technique | References |
|------------------------------|-----------------|----------------|-----------|---------------|-------------------------------|--------------------|------------|
| Sorghum malt (Otambo)        | Namibia         | 45             | AFB1      | 0.56–54.2     | 0.17                          | LC/MS/MS           | [84]       |
| Sorghum beer                 | Namibia         | 45             | AFB2      | 0.5–4.48      | 0.04                          | LC/MS/MS           | [84]       |
| Sorghum-based products       | Mediterranean area | 1/4           | AFB1      | 0–6.4         | 0.25                          | LC/MS/MS           | [41]       |
| Wheat                        | Brazil          | 35             | Total AFs | 0–6.2         | 5.0                           | HPLC-FD            | [86]       |
| Wheat                        | Spain           | 14/60          | AFB1      | 1.03–9.50     | 0.08                          | LC/MS/MS           | [87]       |
| Wheat                        | Spain           | 19/60          | AFB2      | 0.34–0.67     | 0.08                          | LC/MS/MS           | [87]       |
| Wheat                        | Spain           | 6/60           | AFG1      | 0.53–1.05     | 0.16                          | LC/MS/MS           | [87]       |
| Wheat                        | China           | 21/32          | AFB1      | 0.03–0.12     | 0.03-0.2                      | LC/MS/MS           | [88]       |
| Wheat                        | Egypt           | 36             | AFB1      | 0.13–49.79    | 0.04                          | HPLC               | [89]       |
| Wheat                        | Egypt           | 36             | AFB2      | 0.09–2.96     | 0.12                          | HPLC               | [89]       |
| Wheat                        | Egypt           | 36             | AFB1      | 0–1.8         | 3                             | HPLC-FD            | [80]       |
| Wheat                        | Bangladesh      | 227            | AFB1      | 0.9–1.6       | 0.2                           | HPLC               | [76]       |
| Wheat                        | Mediterranean area | 3/21      | AFB2      | 6.7–26.0      | 1.50                          | LC/MS/MS           | [41]       |
| Wheat                        | Mediterranean area | 10/65     | AFB1      | 5.5–66.7      | 0.25                          | LC/MS/MS           | [41]       |
| Wheat                        | Mediterranean area | 2/65       | AFB2      | 5.6–7.6       | 1.5                           | LC/MS/MS           | [41]       |
| Wheat-based baby foods       | Iran            | 4/16           | AFB1      | 0–1.8         | 3                             | HPLC-FD            | [80]       |
| Wheat bran                   | Iran            | 2/2            | AFB1      | 1.03–2.50     | 3                             | HPLC-FD            | [80]       |
| Wheat flour                  | Brazil          | 32             | Total AFs | 4.8           | 5.0                           | HPLC-FD            | [86]       |
| Wheat flour                  | Iran            | 144/180        | Total AFs | 0.01–0.5      | 0.003                         | HPLC               | [91]       |
| Wheat flour                  | Turkey          | 60             | AFB1      | 0–0.04        | 0.026                         | HPLC               | [67]       |
| Wheat flour                  | China           | 108            | AFB1      | 0.1–0.9       | 0.1                           | HPLC               | [47]       |
| Wheat flour (whole)          | Brazil          | 16             | Total AFs | 3.4           | 5.0                           | HPLC-FD            | [86]       |
| Wheat flour (refined)        | Brazil          | 15             | Total AFs | 1.2           | 5.0                           | HPLC-FD            | [86]       |
| Wheat bran                   | Iran            | 54/60          | Total AFs | 0.06–0.99     | 0.01                          | HPLC               | [91]       |

HPLC-FLD: high-performance liquid chromatography–fluorescence detector; DLLME-HPLC: dispersive liquid–liquid microextraction coupled to high-performance liquid chromatography with fluorescence detection; TLC: thin-layer chromatography; UHPLC/TOFMS: ultrahigh-performance liquid chromatography/time-of-flight mass spectrometry; ELISA: enzyme-linked immunosorbent assay; LC-MS/MS: liquid chromatography–tandem mass spectrometry; AFB1: aflatoxin B1; AFB2: aflatoxin B2; Total AFs: aflatoxins (B1 + B2 + G1 + G2).
3. Chemistry and Biosynthesis of Aflatoxins

All AFs are heterocyclic compounds with a basic benzene ring, with minor differences in the occurrence of double bonds and ketonic groups, as well as the presence of hydroxy groups in the metabolites, with hydroxylation sites varying from one molecule to another. These structures imply a low water solubility and an easy epoxidation reaction, which are expected to impact both elimination and lethality. The most common and potent human health concern in the world, AFB1, contains a unique double bond in the cyclic ring, which is also seen in G1 and M1. AFB1 must be epoxidized to AFB1 2,3-epoxide in order to be functional. The toxin is biotransformed into the less lethal AFs M1 and G1 by microsomal cytochrome P450 (CYP450) monoxygenases [92,93]. A double bond at carbons 8 and 9 in AFs B1 and G1 facilitates the synthesis of epoxide, a more lethal version of AFs B1 and G1, but not in AFs B2 and G2. The dihydroxy derivatives of B1 and G1 were identified as AFs B2 and G2, respectively. AF M1 is a 4-hydroxy AFB1, whereas AF M2 is a 4-dihydroxy AF B2. B1 and G1 are hydrogenated to produce B2 and G2, respectively [92,94].

The primary substrate of hexanoyl is transformed into a polyketide by a polyketide synthase and two fatty acid synthases during the biosynthesis of aflatoxins in crops by *Aspergillus flavus* and *Aspergillus parasiticus* [95,96], followed by the conversion of the polyketide to norsolorinic acid anthrone by polyketide synthase; thereafter, norsolorinic acid anthrone is converted to norsolorinic acid (NOR), which is the first stable forerunner of aflatoxins (Figure 2) [97,98]. Then, the reductase enzyme converts NOR to averantin [99], and 5′-hydroxyaverantin (HAVN) is created from averantin using the monoxygenase enzyme [100]. Further, dehydrogenase converts HAVN to 5′-oxoaverantin (OAVN), which forms averufin (AVF) using cyclase [101–103], followed by the Baeyer–Villiger reaction, forming hydroxyversicolorone (HVN) from AVF [104]. Next, HVN is oxidized to versicolonal hemiacetal acetate (VHA), which is further converted to versiconal acetate (VOAc) and then to versiconol (VOH) [105]. Using esterase, VOH forms versiconal, which is then transformed into versicolorin B by cyclase [106]. Furthermore, versicolorin B is converted to versicolorin A and dimethyl-dihydro-sterigmatocystin (DMDHST). Next, versicolorin A forms dimethyl-sterigmatocystin (DMST), and DMDHST forms dihydro-sterigmatocystin (DST) [107,108]. Thereafter, O-methyltransferases transform the intermediates of DMST and DHST to sterigmatocystin (ST) and dihydro-O-methylsterigmatocystin (DHOMST), respectively, playing a crucial role in the biosynthesis of AFs [109]. Next, ST produces O-methylsterigmatocystin (OMST), which, along with DHOMST, finally produces AFs [110,111].
Figure 2. Biosynthesis pathway of aflatoxins. Reprinted with permission from Nazhand et al. [112].

In Aspergilli, DNA information is structured into eight chromosomes, with genes relevant for AF production found in the 54th cluster, 80 kb from chromosome 3’s telomere [113]. This cluster contains 30 genes, and aflR and aflS are the major regulators of its activation [114,115] (Figure 3). In Aspergillus flavus and Aspergillus parasiticus, the AF gene cluster has been extensively explored. The clustered genes of the two fungal species are 90–99% homologous [109]. The ability to create B- and G-type AFs is one of the key differences between the two species. Aspergillus flavus produces mostly B-type AFs (B1 and B2), while Aspergillus parasiticus produces both B- and G-type AFs (B1, B2, G1, and G2). The functional genes aflU, aflF, and nadA, which code for a cytochrome P-450 monoxygenase,
an aryl alcohol dehydrogenase, and an oxidase, respectively, are involved in the formation of G-type AFs [116,117]. Experiments on the potential of *Aspergillus nidulans* to make sterigmatocystin have also aided in the understanding of the aflatoxin cluster. Indeed, there is 55–75% similarity between *Aspergillus parasiticus* and *Aspergillus nidulans* clusters [109].

![Figure 3](image_url). Aflatoxin gene cluster organization, including old and new cluster gene nomenclatures (red dotted lines represent the binding sites of AflR in the above pathway). Figure reprinted from Caceres et al. [96].

4. Health Effects and Mechanism of Toxicity

Human exposure to AFs can occur at any stage of life, either directly by the ingestion of AF-contaminated food or indirectly due to the intake of foods (milk, egg, meat, etc.) derived from livestock previously exposed to AF-contaminated feed [118,119]. When ingested, AF passes through the metabolic process in mammals and remains unaltered, and it later accumulates in the tissues [21]. It is now well established that, apart from cancer, AFs also cause acute and severe chronic diseases. Initially, the carcinogenicity of AFs was identified and associated with the liver, which first metabolizes them and produces reactive intermediary metabolites. However, subsequent epidemiological and animal studies revealed their carcinogenic effects on other organs, including the kidney, pancreas, bladder, bone, viscera, and central nervous system [120]. Evidence has shown that AFB1-mediated oxidative stress is equally or even more responsible for AF-induced genotoxicity. The second-most documented toxicological effect of AFs is probably immunotoxicity, and its mechanisms of action (immunosuppressive and immunostimulatory actions) have been extensively illustrated [121]. Apart from the above, malnutrition, disease, impaired child growth, retardation of physical and mental maturity, reproduction, nervous system diseases, etc., are some other AF-induced acute and chronic health issues reported in mammals. However, further studies are required to demonstrate their precise mechanisms of action [122].

Different mechanisms of action are responsible for the various toxicological effects of AFs, but most of them are not fully understood yet. Since the AF discovery, AFB1 has been a major focus, as it is responsible for forming the intermediate metabolite AFB1-exo-8,9 epoxide (AFBO) [123]. This intermediate metabolite is a highly unstable molecule that reacts with different cellular macromolecules, including nucleic acids, proteins, and phospholipids, and thereby induces various disruptions at the genetic, metabolic, signaling, and cellular structure levels [124,125]. However, several studies have also evidenced that AFB1 has equivalent or even more effects on the integrity of cell function due to induced oxidative stress (OS) [122,126,127].

5. Effects of Environmental Factors on Aflatoxin Production

Environmental factors, such as water activity ($a_w$), temperature, and pressure, are critical factors for *A. flavus* growth and AF accumulation. In addition, pH, CO$_2$ levels, and light exposure have also been shown to significantly impact fungal growth and AF production [128–130]. Lowering $a_w$ in foods inhibits microbiological proliferation and biochemical processes, extending the food product shelf life [131]. The proportions of AF-producing microbial communities that develop during the pre-harvest step have a significant impact...
on the post-harvest step, and the impact of prolonged harvesting on contamination is especially critical when rain damages crops before or during harvesting [132]. Variables for germination, proliferation, and AF production by *A. flavus* and *A. parasiticus* reveal that proliferation happens on a broader spectrum than production, with AF production occurring over an even smaller range than growth. The ideal conditions for AF production in these two microbial species (*A. flavus* and *A. parasiticus*) are 33 °C and 0.99 a_w; on the contrary, the optimal conditions for production are 35 °C and 0.95 a_w [133]. Pitt and Miscamble [134] reported that the influence of environmental conditions on the development of *A. flavus*, *A. parasiticus*, and *A. oryzae* was comparable, exhibiting minima at 0.82 a_w/25 °C and 0.81 a_w/30 and 37 °C. However, there was no assessment of AF production comparing *A. flavus* and *A. parasiticus* in the investigative study. Giorni et al. [135] reported that moist maize supplemented with 25% CO_2 is adequate for the inhibition of *A. flavus* germination, while about 50% CO_2 was necessary to substantially reduce AF formation. Managing hydrogen peroxide [136] and carbon dioxide [137] concentration through food processing and storage is, thus, an appropriate management method for avoiding *A. flavus* infestation and consequent AF production.

6. Detection Techniques

Various chromatographic, spectrometric, and sensor-based techniques have become prevalent for the sensitive detection and quantification of AFs in cereals and cereal-based products. Among chromatographic techniques, liquid chromatography (LC), thin-layer chromatography (TLC), and high-performance liquid chromatography (HPLC), along with numerous detectors, have been utilized for cereal-based food and feed products. HPLC is the most sensitive and accurate technique. Samples for HPLC are prepared through dry, wet, or cryogenic grinding, solvent extraction, solid-phase extraction, immunoaffinity column clean-up, and supercritical fluid extraction [138]. Namjoo, et al. [139] proposed the use of an HPLC column with a mobile phase of water–methanol–acetonitrile (60:30:15, v/v/v) coupled with a fluorescence detector to quantify the levels of AF B1, B2, G1, and G2, in wheat silos in Golestan Province of Iran. AF presence was found in ten out of thirty-four wheat samples, all of which were below the permitted levels in Iran (15 µg/kg). The highest levels found in samples for total AFs, AFB1, AFB2, AFG1, and AFG2 were 7.08, 6.91, 0.29, 1.37, and 0.23 µg/kg, respectively. Likewise, AFB1 was analyzed in wheat and rice flour samples with immunoaffinity column clean-up using post-column photochemical derivatization and HPLC-FLD, where the limits of determination of AFB1 were observed to be 0.015 and 0.05 µg/kg, respectively. Spectrophotometric detection methods are preferred over these chromatographic methods, as they help to directly screen large lots of cereals to segregate putatively contaminated kernels. Near-infrared spectroscopy, fluorescent spectroscopy, and multispectral and hyperspectral imaging are some commonly used techniques that allow the inexpensive and rapid detection of AFs in cereal grains such as wheat, maize, corn, sorghum, rice, etc. [140–142]. For instance, Liu et al. [143] applied surface-enhanced Raman spectroscopy (SERS) combined with the QuEChERS sample pretreatment technique for the trace-level detection of aflatoxin B1 in wheat and corn with high sensitivity and strong specificity. The method showed a good linear-log relationship between the Raman signal intensity of AFB1 in the 1–1000 µg/L concentration range with a limit of detection of 0.85 µg/kg. SERS is a rapid and sensitive technique for detecting trace mycotoxins. However, the sample preparation of SERS substrates for the quantitative analysis of foods is relatively challenging. Collectively, chromatographic–spectrometric techniques offer the advantage of a “dilute and shoot” approach, where sample extracts are analyzed without clean-up, and with the added advantage of multi-mycotoxin analysis, whereby a range of mycotoxins can be analyzed in the same sample analysis [144].

Lately, nanomaterial-based sensors with versatile properties have been observed to be highly selective and specific in terms of the trace detection of AFs in food samples. These also cater to the limitations posed by other conventionally used detection methods, as mentioned previously. Many nanomaterials have been used for the immobilization
of biomolecules as signal generators or fluorescent quenchers or for signal amplification in AFB1 detection [145,146]. Recently, graphene oxide and gold nanowires were used as novel nanomaterials to develop fluorescence resonance energy transfer (FRET) and electrochemical-based sensors for the detection of AFs [147]. Further, electrochemical aptasensors based on carbon-based nanocomposites have also demonstrated an exceptional ability to detect and quantify AF concentrations in cereal-based foods [148–150].

7. Detoxification and Management Strategies

7.1. Conventional Agricultural Practices

AF production is stimulated by drought, stress, and high temperatures due to improper cultivation practices and post-harvest management [31]. The execution of modern agricultural practices, such as good agricultural practices (GAPs), good manufacturing practices (GMPs), good storage practices (GSPs), and preharvest management strategies [2,151], might be helpful for the prevention of AF contamination in grains. Various breeding programs have experimented on maize for the development of AF-resistant lines using inbred lines resistant to AF-resistant germplasm and the associated resistance proteins for host resistance, proteomic and molecular breeding, and genetic engineering, employing potential biochemical markers and genes for resistance [152–156]. These approaches could help manage AFs in pre- and post-harvest conditions. The AF content and the chances of contamination under field conditions also depend on the sowing time and cultivation region. Higher contamination was observed in late-cultivated compared to early-cultivated maize crops [157,158]. Further, several physical, chemical, and biological methods are being applied for the decontamination of AFs in cereal and cereal-based products.

7.2. Physical Methods

Physical methods can be applied through washing, cleaning, density segregation, sieving, dehulling, adsorption, thermal treatments, and the use of gamma, UV, and visible light radiation [159]. Generally, agricultural produce goes through a common practice of sorting. This practice is supported by the fact that mycotoxin contamination tends to have a skewed distribution, with the majority of toxins found in a small number of grains or kernels [160]. Matumba et al. [161] performed experiments to evaluate the effectiveness of various physical methods in mycotoxin-contaminated white maize. Out of the experimented methods, hand sorting was found to be the most effective in reducing AFs, which removed about 95% of the AF content in white maize. However, sorting practices have been improved, and better technologies are being used for better accuracy and speed. Pearson et al. [162] used a high-speed dual-wavelength method for sorting yellow corn using absorbances at 750 and 1200 nm. It correctly identified >99% of AF-contaminated kernels, which reduced about 81% of AFs.

Irradiation is a method in which a product is penetrated with ionizing radiation [163]. Silva et al. [164] investigated gamma irradiation ($^{60}$Co) as a rice storage option. They found that it effectively controlled Penicillium spp. and Aspergillus spp., resulting in a reduction in AF levels in stored grains. The application of low-temperature radiofrequency plasma was used to degrade AFs, and the quantities of AFB1 (2–50 g/L) in the solution were reduced by 88% [165]. In addition, aflatoxigenic molds were successfully managed at different stages of germination, sporulation, and growth by 5 kGy gamma irradiation. The inhibition efficacy was observed to be dose-dependent in the feed sample. A 10 kGy dose reduced about 90% of AFB1, while a 5 kGy dose reduced 70% of AFB1 in artificially contaminated maize samples [166]. Besides this, ultraviolet (UV) irradiation for 1–3 h followed by 6 months of storage revealed 22–79% fewer fungal colonies as well as lower AF content in brown, black, and red rice grains [167]. Furthermore, pulse light (PL) is a non-thermal approach used for the inhibition of AFs. A. flavus-contaminated rough rice and rice bran were treated with PL at 0.52 J/cm²/pulse for varying time durations. The treatment removed 75.0% of AFB1 and 39.2% of AFB2 in rough rice and 90.3% of AFB1 and 86.7% of AFB2 in rice bran when treated for 80 s and 15 s, respectively [168].
Besides this, extrusion and baking are two of the most utilized heat-based procedures in the food sector for minimizing mycotoxins. Extrusion is a high-temperature, short-time procedure involving heat, humidity, and mechanical force. It alters raw materials in such a way that they take on new shapes and structures with distinct functional and nutritional properties. In the food industry, this heat treatment is used to produce biscuits, pasta, ready-to-eat cereals, snacks, pellets, etc. The presence or absence of additives, moisture content, and temperature all appear to alter the effect of extrusion on AF levels. Extrusion alone was demonstrated to reduce AF levels by 50–80%, and adding ammonia, either as hydroxide (0.7–1.0%) or as bicarbonate (0.4%), increased the AF reduction to more than 95% in cereals [169]. Baking is another crucial process in the preparation of cereal-based meals, such as cookies, loaves, and cakes. Milani, et al. [170] reported that employing active dried or compressed yeast during the bread-making procedure lowered the amounts of AFs in bread by 32–64%. Further, in one of the studies involving muffins prepared from corn originally contaminated with AFB1, the toxin was reduced from 87 to ± 4% due to the frying process [171]. Another approach for lowering AF concentrations is nixtamalization (alkalization at higher temperatures), which is employed in manufacturing corn chips, tortilla chips, and corn tortillas. This treatment resulted in a 51–78% reduction in AF concentrations for various AF subtypes [172].

Plasma is the fourth state of matter and an emerging technology that shows great potential in various applications. Plasma is a quasi-neutral ionized gas that is primarily composed of free electrons, photons, and ions. Based on its generation, it can be classified as thermal and non-thermal plasma. Treatment with cold atmospheric plasma significantly reduces the microbial load without greatly affecting the quality of food. For example, plasma application at 40 W for 20 min significantly inhibited the growth of A. flavus for up to 20 days during storage at 25 °C [173]. Shi et al. [174] treated AF-contaminated corn with high-voltage cold atmospheric plasma under different conditions of the gas type, relative humidity, and treatment time. The results showed that the combination of MA65 gas, higher relative humidity (80%), and longer time (10 min) degraded the AF content up to 82%. However, the loss of dietary material and/or nutrients should be considered when lowering mycotoxin levels. Mycotoxin reduction treatments may also release and increase the bioavailability of masked mycotoxins in treated products [175] or alter their chemical structures into forms that are undetectable by standard analytical methods while preserving their toxicity [176]. Hence, specific analytical tools are required to discover structural alterations and links to the food matrix to solve these issues.

### 7.3. Chemical Methods

Chemical methods are generally easier to apply and quicker in their responses post-treatment. Mostly plant protection chemicals such as fungicides and insecticides are used [177]. Some food additives are used for the disinfection of aflatoxin in food materials. Shi et al. [178] used four food additives, viz., sodium bisulfite, sodium hypochlorite, citric acid, and ammonium persulfate, to reduce aflatoxin content from distillers wet grains (DWG) and condensed distillers soluble (CDS). These DWG and CDS are nutrient-rich byproducts of shelled maize. Out of four food additives, citric acid showed the most significant result. It removed about 65 and 80% AF content in DWG and CD, respectively, at a 2.5% concentration when heated at 90 °C for 60 min.

Chlorine is a strong oxidizing agent that is used as a sanitizing and cleaning agent for water, cereals, and horticultural commodities. This effectively degrades organic compounds via electron transfer. Samples contaminated with aflatoxin were collected from China and treated with chlorine gas. AFB1 was broken down into four compounds, and it was found to be non-toxic when tested on human embryo hepatocytes. The highest degradation achieved was up to 90%; however, the optimal degradation efficiency was about 79.6%. So, the application of chlorine was found to be a very effective and economically viable approach to detoxifying AFB1 in grains [179]. Besides this, ozone (O3) treatment was also found to be beneficial in reducing AFB1 in maize. The effect of treatment was noted to
occur in a time- and concentration-dependent manner; however, the efficacy was higher in low-moisture maize. Ozone treatment (90 mg/L) successfully removed about 78.16 to 88.07% of AFB1 with 20 and 40 min of treatment [180]. Similarly, the application of O₃ (60 µmol/mol) for 180 min completely inhibited the growth of *A. flavus*, and O₃ also degraded up to 95% AF in wheat [181].

Besides these practices, synthetic fungicides have also been used to manage AFs. Several synthetic fungicides have been proven to control AFs produced by *Aspergillus* sp. In a study, Aleksić et al. [182] investigated the use of pyrimethanil to control the AFs produced by *Aspergillus niger*. They reported that pyrimethanil effectively retarded the appearance of fungal rot [182]. Lagogianni and Tsitsigiannis [177] studied the efficacy of several synthetic fungicides (azoxyystrobin, boscalid, cyprodinil, fludioxonil, and cyprodinil + fludioxonil) on AFs produced by *A. flavus* in maize. They reported that cyprodinil was the most effective fungicide (EC₅₀ < 0.05 µg/mL), followed by fludioxonil (EC₅₀ < 0.11 µg/mL). The least effective was boscalid (EC₅₀ 4.35–4.50 µg/mL). Moreover, they reported that in a 2-year field study, cyprodinil + fludioxonil showed an 83% reduction in AF contamination by *A. flavus* in maize [177]. Prochloraz, an azole fungicide, is more effective than tebuconazole in controlling the growth and production of aflatoxin according to Mateo et al. [183], who assessed the efficacy of azole-based fungicides (prochloraz, tebuconazole, and prochloraz + tebuconazole [2:1 w/w]) to manage the AFs produced by *A. flavus* in maize kernels. They reported that prochloraz was more effective, followed by prochloraz + tebuconazole (2:1) and tebuconazole. Ferrigo et al. [184] reported the higher efficacy of prothioconazole + tebuconazole in controlling the growth of AFs produced by *A. flavus*. In addition, Masiello et al. [185] investigated the efficacy of the succinate dehydrogenase inhibitor fungicides boscalid and isopyrazam in controlling AFs produced by *A. flavus*. Further, Magnoli et al. [186] studied the use of chlorpyrifos to control the growth of aflatoxin B₁ produced by the *Aspergillus* section *Flavi* strain. These approaches, however, have numerous disadvantages. These fungicides are harmful and pollute the environment. The usage of synthetic fungicides might pose health concerns. These synthetic fungicides are carcinogenic to humans.

7.4. Biological Methods

Biological detoxification involves using fungi, bacteria, and actinomycetes to reduce or completely remove AFs in food products through either adsorption or enzymatic degradation [89]. Some strains of microorganisms and volatiles have been reported to have a positive response in the inhibition of AFs. The volatiles produced by *Bacillus megaterium* KU143 and *Pseudomonas protegens* AS15 noticeably repressed the growth, sporulation, and conidial germination of *A. flavus* under in vitro conditions, as well as fungal populations in rice grains during storage [187]. Further, Shetty et al. [188] observed the AFB1-binding ability of *Saccharomyces cerevisiae*. The highest binding efficacy was observed to be up to 53% during the exponential growth phase, and the efficacy was reduced when reaching the stationary phase.

In addition, the effect of six biopesticides/biostimulants, viz., Botector®, Mycostop®, Serenade Max®, Trianum®, Vacciplant®, and zeolite, was evaluated by Lagogianni and Tsitsigiannis [189] under in vitro and field conditions. Mycostop®, Serenade Max®, Vacciplant®, and zeolite significantly inhibited the growth of *A. flavus* conidia production by 38.8–63.1% under lab conditions. However, Mycostop® and Botector® treatments exhibited significant decreases in disease incidences of 16.5 and 21.9%, respectively. Further, they also reduced AFB1 concentrations in maize kernels by 43.05 and 43.09%, respectively, when applied twice during the silk stage. The biological agents contained in Mycostop® and Botector® are *Streptomyces griseoviridis* and *Aureobasidium pullulans*, respectively [189]. Additionally, a pre-harvest spray treatment of *Trichoderma harzianum* T77 on the silks of sweetcorn plants under greenhouse and field conditions exhibited the inhibition of *A. flavus* infection. The author suggested that the combination of proper post-harvest management could reduce AF content [190]. Hruska et al. [191] applied a non-aflatoxigenic *A. flavus* biocontrol
strain on maize, which suppressed the growth of aflatoxicogenic *A. flavus* by up to 82%, and consequently, a smaller population, with up to 73% of suppression, was observed in AF content.

7.5. Use of Phytochemicals

Due to the harmful effects and risks associated with synthetic chemicals, the demand for natural, safer, and eco-friendly methods has increased. Several plant extracts, including essential oils (EOs), are suitable alternatives [192]. A screening study of plants such as *Psorospermum febrifugum*, *Prospis Africana*, and *Curcuma longa* showed the presence of phytochemicals, including steroids, terpenoids, glycosides, phenols, tannins, saponins, and flavonoids, with anti-aflatoxicogenic activity [193]. Kavitha et al. [194] reported that plants such as *Zingiber officinalis* (Zinger), *Oxalis corniculate* (Indian Sorrel), *Trigonella foenum-graecum* (Fenugreek), *Stevia rebaudiana* (Sugar Leaf), and *Equisetum arvense* (field horsetail) have anti-aflatoxicogenic and anti-fungal activities. The aqueous leaf extract of *Salvia farinacea* and *Azadirachta indica* at 2 mg/mL has been reported to have antifungal activity against *Aspergillus parasiticus* in different food samples. Phytochemicals extracted from *Thymus* spp. have antifungal activity against *A. flavus* and *A. parasiticus*. Thymol from *Thymus kotschyanus* was used against *A. flavus* at 0.5 µg/mL [195]. Similarly, curcumin has the potential to inhibit *A. flavus* by inhibiting the Cyt450 isoenzyme CYP2A6 and reducing the formation of AFB1-8, 9 epoxide [196]. Several phytochemicals against AF-producing fungi are presented in Table 2.

| Phytochemical Source | Phytochemical Form | Target Fungi | Food Commodity | Outcomes | References |
|----------------------|--------------------|--------------|----------------|----------|------------|
| Schinus mole (Pepperina) | Nanoparticles | *A. parasiticus* | Maize | 59% control of aflatoxin production; fungal contamination and production of AFB1 and AFB2 inhibited at 250 µL/mL | [197] |
| Rosmarinus officinalis (Rosemary) | EOs | *Aspergillus flavus* | Not available (NA) | Significant prevention of fungal contamination and aflatoxin production | [198] |
| Pelargonium graveolens (Sweet scented or rose Scented Geranium) | Nanogel | *A. flavus* | Maize | 77.96% prevention at 1.0 µL/mL of nanogel | [199] |
| Carum carvi (caraway), Juniperus communis (juniper), Satureja Montana (winter savory), Origanum virens (Oregano) | EOs | *A. flavus*, *A. parasiticus* | Maize flour | Significant prevention of fungal contamination and aflatoxin production | [200] |
| Zataria multiflora (Satar) | Niosome | *A. flavus* | Maize | Reduction in fungal growth and aflatoxin accumulation | [201] |
| Satureja montana (winter savory), Origanum virens (Oregano), Thymus vulgaris (garden thyme), Melaleuca alternifolia (tea tree), Mentha piperita (Peppermint), Myristica fragrans (Nutmeg), Clove & Quercetin from Syzygium aromaticum, Cyanidin from Solanum lycopersicum, Curcumin from Curcuma longa L. (Turmeric) | Solid lipid nanoparticles and EOs | *A. flavus* | NA | Enhanced antifungal activity observed | [202] |
| | EOs | *A. flavus* | NA | Significant reduction at 0.96aw | [203] |
| | Nanocomposite films | *A. flavus*, *A. parasiticus* | Rice | 51–77% reduction in fungal growth during storage | [204] |
| | Nanoemulsion | *A. flavus* | Rice | Significant inhibition of AFB1 production | [205] |
| | Phytochemical | *A. flavus*, *A. parasiticus* | NA | Inhibited AF production | [196] |
| | Phytochemical | *A. flavus*, *A. parasiticus* | NA | Inhibition of AFB1 production | [206] |
| | Phytochemical | *A. flavus* | NA | Prevention of hyphae production | [207] |

Table 2. Phytochemicals from various plant sources effective against aflatoxin-producing fungi.
Further, EOs have the ability to cross the plasma membrane and cause lipid partition in the cell membrane of fungi and the subsequent leakage of the cell contents. EOs can decrease the biosynthesis of compounds such as ergosterol and sterol [212]. EOs from six plant species (Rosmarinus officinalis, Thymus vulgaris, Satureja montana, Origanum virens, O. majoricum, and O. vulgare) were tested in in vitro conditions against A. flavus and AF contents. At the higher concentration, all six EOs showed great efficiency and completely inhibited fungal growth as well as inhibited about 100% of AFB1 production at 500 µg/mL [201].

EOs from anise (Pimpinella anisum) and boldus (Peumus boldus) were effective against Aspergillus parasiticus and Aspergillus flavus in stored maize [212,213]. EOs from bergamot, bitter orange, and lemon at 2% showed the inhibition of mycelial growth as well as AFB1 production by A. flavus. These EOs can be recovered and reused from citrus waste and are eco-friendly fungicides that could also be profitable for the agriculture sector [214]. The EOs extracted from Cananga odorata showed antifungal activity against A. parasiticus and A. flavus at 4 mg/mL [215]. The shelf life of brown rice was enhanced by Michelia alba oil in combination with linalool and caryophyllene at different ratios, while a 1:10 ratio showed the strongest antifungal activity against A. flavus [216]. EOs from the seeds of Anethum graveolens were reported to have inhibitory potential against A. flavus, along with other fungi at a MIC (minimum inhibitory concentration) of 120 µL/mL [217]. A. flavus sporulation on infected maize seeds was significantly suppressed by black Piper nigrum essential oil (BPEO) fumigation, with 100% inhibition being the highest result at 50 and 100 µL/mL of BPEO [218]. The nanoencapsulated Pimpinella anisum essential oil (PAEO) was found to preserve the stored rice samples against fungal and AFB1 contamination [219]. The nanoencapsulated EO of Cananga odorata with chitosan has been reported to have in vitro and in vivo preservative actions against A. flavus contamination and AFB1 production [220].

Furthermore, a phytochemical, terpinen-4-ol (28.92%) from Origanum majorana EO (nanoencapsulated), was reported to cause the in situ inhibition of lipid peroxidation and AFB1 production in maize [221]. In addition, ginger essential oils (GEO) containing 23.85% α-zingiberene and 14.16% geranial inhibited A. flavus growth and AFB1/AFB2 production, respectively, at 25 and 50 µg/g GEO in stored maize grains [222]. However, the efficacies of plant extracts and their phytochemicals in the management of toxigenic fungi and their toxins inherit limitations to solely be used as biofungicides and nutraceuticals [196]. Hence, these can be used in combination with the other detoxification methods discussed above for the better control and management of AFs in cereals and cereal-based products.

8. Conclusions

Aflatoxin (AF) contamination of cereals and cereal-based products has caused severe health concerns for humans, in addition to substantial economic losses. AF formation is influenced by various parameters in field conditions and/or during storage. Further, their detoxification using physical, chemical, and biological methods can be controlled and
managed to some extent. The growing awareness of the adverse effects of synthetic chemicals used in conventional practices on human health and the environment has focused the interest of researchers on AF management using phytochemicals and their essential oils (EOs). In addition, the use of phytochemicals has proven to be a natural, safe, and eco-friendly method for AF detoxification, benefiting both the environment and the consumer. These practices are sustainable, as they are natural and environmentally friendly. Thus, they pose no risk to the environment or consumers. These phytochemicals release some EOs that are effective against AFs. These EOs may be reused and recovered, making them more environmentally friendly and sustainable. Thus, this practice for AF detoxification ensures food safety, as these agents are free from any toxic residues that are generated due to employing conventional practices. Moreover, combining these phytochemicals with encapsulation techniques can detoxify AFs with little consequences. This approach further enhances the bioavailability of phytochemicals with high efficacy at low concentrations with strong AF-inhibiting properties. Furthermore, these phytochemicals play dual pharmacological and nutraceutical roles as biofungicides and detoxifying agents in mitigating the effects of AFs. Thus, phytochemicals can be used as an alternative AF detoxification practice to ensure food safety. To ensure food safety, there is a strict need for effective and safer management practices that can delay the growth of AFs in food without altering their sensorial attributes. Moreover, phytochemicals and their EOs have been regarded as GRAS (Generally Recognized as Safe). Using phytochemicals and their EOs strengthens their application in achieving “green consumerism” in the agriculture and food sector. However, further study is needed to examine the usefulness of phytochemicals in managing AFs. In-depth research on the interaction processes between encapsulated phytochemicals and food products and their impact on human and animal health should be conducted. Inadequate information in these areas highlights the need for more in-depth research into their chemical characteristics, biosynthetic pathways, and diverse practices for detoxification and management strategies to ensure food safety and security.

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