







HPPD-resistant cotton response to isoxaflutole applied preemergence and postemergence

Joshua D. Joyner¹ , Charles W. Cahoon² , Wesley J. Everman³ ,
Guy D. Collins³ , Zachary R. Taylor⁴  and Andrew C. Blythe¹ 

Research Article

Cite this article: Joyner JD, Cahoon CW, Everman WJ, Collins GD, Taylor ZR, Blythe AC (2022) HPPD-resistant cotton response to isoxaflutole applied preemergence and postemergence. *Weed Technol.* **36**: 238–244. doi: [10.1017/wet.2022.6](https://doi.org/10.1017/wet.2022.6)

Received: 28 September 2021
Revised: 23 December 2021
Accepted: 19 January 2022
First published online: 10 February 2022

Associate Editor:

Daniel Stephenson, Louisiana State University Agricultural Center

Keywords:

cotton tolerance; cotton injury

Nomenclature:

acetochlor; dimethenamid-*P*; diuron; fluometuron; fluridone; fomesafen; glufosinate; glyphosate; isoxaflutole; pendimethalin; pyriithiobac; cotton; *Gossypium hirsutum* L.

Author for correspondence:

Charles W. Cahoon Jr., Department of Crop and Soil Sciences, North Carolina State University, Campus Box 7620, Raleigh, NC 27695
Email: cwcahoon@ncsu.edu

¹Graduate Research Assistant, Department of Crop and Soil Sciences, North Carolina State University, Raleigh, NC, USA; ²Assistant Professor, Department of Crop and Soil Sciences, North Carolina State University, Raleigh, NC, USA; ³Associate Professor, Department of Crop and Soil Sciences, North Carolina State University, Raleigh, NC, USA and ⁴Research Specialist, Department of Crop and Soil Sciences, North Carolina State University, Raleigh, NC, USA

Abstract

Studies were conducted in 2019 and 2020 in Lewiston, NC, to determine the crop response of 4-hydroxyphenylpyruvate dioxygenase (HPPD)-resistant cotton to isoxaflutole (IFT) and other cotton herbicides as part of a cotton weed management program that included herbicides applied preemergence, early postemergence (EPOST), and mid-postemergence (MPOST). IFT was applied PRE at 105 g ha⁻¹ alone and in various combinations with acetochlor, diuron, fluometuron, fluridone, fomesafen, pendimethalin, and pyriithiobac. EPOST treatments included IFT at 53 or 105 g ha⁻¹ alone or in combination with glyphosate or glufosinate, or dimethenamid-*P* + glufosinate. Glyphosate + glufosinate was applied MPOST to all treatments except the nontreated control. Cotton injury from IFT applied PRE was minimal (0% to 3%). Injury following EPOST application of dimethenamid-*P* + glufosinate ranged from 3% to 5% and 6% to 9% in 2019 and 2020, respectively. In both years, injury from IFT applied PRE followed by IFT applied EPOST never exceeded injury from IFT applied PRE followed by dimethenamid-*P* + glufosinate. Isoxaflutole applied PRE followed by IFT applied EPOST at 105 g ha⁻¹ resulted in 0% to 2% cotton injury, indicating that IFT can be applied either PRE or EPOST with minimal risk to cotton. Late-season cotton height and cotton lint yield were not affected by any herbicide treatment. The experimental HPPD-resistant cotton cultivar was minimally injured by IFT applied PRE and EPOST, it tolerated standard cotton herbicides, and yield loss was not observed. Given these results, HPPD-resistant cotton and IFT may be integrated into cotton weed management systems with minimal risk for cotton injury and provide an additional effective mechanism of action for managing troublesome weeds in cotton.

Introduction

In 2020, 96% of U.S. cotton acreage included genetically engineered resistance to one or more herbicides (USDA-NASS 2020). This trend began with rapid adoption of glyphosate-resistant (GR) cotton, which allowed cotton growers to control troublesome weeds postemergence (POST) with minimal risk for crop injury (Gianessi 2008). Extensive use of glyphosate hastened the evolution of GR weeds, thus creating need for a return to integrated weed control systems and the establishment of additional management tools (Duke and Heap 2017; Kniss 2018).

Before GR cotton was widely adopted, producers used several soil-residual herbicides with multiple effective mechanisms of action (MOAs). A typical recommendation of the time would have included application of a preplant-incorporated (PPI) herbicide, such as pendimethalin or trifluralin, followed by a photosystem II-inhibiting herbicide applied preemergence (PRE), such as diuron or fluometuron. In some instances, application of a postemergence (POST)-directed herbicide would follow to provide additional late-season control (Wilcut et al. 1995). Like the aforementioned strategy, which uses multiple effective MOAs and layered soil-residual herbicides, similar programs are once again encouraged by extension weed scientists to control GR Palmer amaranth (*Amaranthus palmeri* S. Watson; Cahoon and York 2020; Culpepper 2019). Soil-residual herbicides are a critical component of an integrated approach to weed management (Culpepper et al. 2010; Norsworthy et al. 2014; Whitaker et al. 2011; Wiggins et al. 2016) and can delay further evolution of herbicide-resistant weed biotypes (Busi et al. 2020; Neve et al. 2011). In recent years, fluridone, an inhibitor of phytoene desaturase and a new MOA to cotton (Bartels and Watson 1978; Cahoon et al. 2015b; Chamovitz et al. 1993; Kowalczyk-Schroder and Sandmann 1992), was commercialized specifically for controlling herbicide-resistant Palmer amaranth (Anonymous 2019a; York 2014; York 2016). Protoporphyrinogen oxidase (PPO) inhibitors applied preplant and/or PRE, and very long chain fatty acid (VLCFA) inhibitors applied PRE and/or early POST (EPOST) also control GR Palmer amaranth (Bond et al. 2006; Cahoon et al. 2015a; Cahoon and York 2020; Culpepper 2019; Whitaker et al. 2010). However, Palmer amaranth biotypes resistant to PPO and VLCFA inhibitors have been

© The Author(s), 2022. Published by Cambridge University Press on behalf of the Weed Science Society of America. This is an Open Access article, distributed under the terms of the Creative Commons Attribution licence (<http://creativecommons.org/licenses/by/4.0/>), which permits unrestricted re-use, distribution and reproduction, provided the original article is properly cited.



discovered in Tennessee and Arkansas, respectively, bringing into question the longevity of these important MOAs as tools for managing Palmer amaranth (Brabham et al. 2019; Copeland et al. 2018; Copeland et al. 2019; Varanasi et al. 2018; Ward et al. 2013). With the looming threat of additional herbicide resistance and the pace of herbicide discovery at a near standstill (Beckie and Harker 2017; Dayan 2019; Duke 2012), a great need exists to incorporate new effective weed control strategies into cotton production.

Research is underway in anticipation of the commercialization of cotton resistant to 4-hydroxyphenylpyruvate dioxygenase (HPPD)-inhibiting herbicides, for which commercial launch is projected by 2023 (G Baldwin, BASF Corporation, personal communication). In addition to tolerance to the HPPD-inhibiting herbicide isoxaflutole (IFT), these cultivars are anticipated to have tolerance to glyphosate, glufosinate, and dicamba. Isoxaflutole is being evaluated for use both PRE and EPOST. Cotton tolerance to HPPD-inhibiting herbicides was achieved by insertion of an HPPD protein from *Pseudomonas fluorescens*. A single substitution of glycine for tryptophan at position 336 allows for reduced sensitivity to IFT (Boudec et al. 2001; Matrigne et al. 2005; USDA-APHIS 2017).

Isoxaflutole is presently labeled for use in corn (*Zea mays* L.) for control of broadleaf and grass weeds and can be used PPI, PRE, or EPOST (Anonymous 2019b; Anonymous 2019c) and in transgenic HPPD-resistant soybean [*Glycine max* (L.) Merr.] PPI or PRE (Anonymous 2019d). Considerable corn injury has been documented following applications of IFT (Wicks et al. 2007; Wilson et al. 1999), which in some cases, have been attributed to coarse soil texture, low organic matter content, and elevated soil pH (Vrabel 1996; Wicks et al. 2007). The propensity of IFT to injure corn in coarse soils with low organic matter raises concern for future use in cotton, because these soil characteristics are common to cotton production in the southeastern United States (Garcia et al. 2011; NASA 2016; USDA-NASS 2019).

Isoxaflutole could be valuable for managing *Amaranthus* species in cotton, as previous research reports effective residual control (Knezevic et al. 1998; Starkey et al. 2016; Stephenson and Bond 2012; Zhao et al. 2017). Outside of *Amaranthus* species, IFT has also demonstrated activity on other weeds, some of which are troublesome in cotton. Smith (2019) reported that IFT controlled common lambsquarters (*Chenopodium album* L.), common ragweed (*Ambrosia artemisiifolia* L.), and velvetleaf (*Abutilon theophrasti* Medik.). Zhao et al. (2017) noted that IFT has the potential to control large-seeded weeds more effectively than VLCFA-inhibiting herbicides. Some of the most common and troublesome weeds in cotton include grasses, such as barnyardgrass (*Echinochloa crus-galli* L.), johnsongrass (*Sorghum halepense* L.), and goosegrass (*Eleusine indica* L.; Van Wychen 2019). Data suggest that each of these troublesome weeds may be controlled by IFT applied PRE (Bhowmik et al. 1999; Stephenson and Bond 2012; Takano et al. 2018).

Reports of IFT efficacy on weeds following POST application is limited. Starkey et al. (2016) evaluated IFT for POST control of Palmer amaranth and found that control was highly dependent upon weed size. Palmer amaranth ≤ 10 cm was controlled $\geq 94\%$. Young and Hart (1998) reported that methylated seed oil (MSO) applied in combination with IFT improved adsorption and translocation of the herbicide in giant foxtail (*Setaria faberi* Herrm.). Weed control by IFT applied POST could be a tool for managing emerged weeds in reduced-tillage cotton while still providing residual control. Similar assertions were made regarding potential use of IFT in no-till corn (Armel 2002; Vrabel et al. 1996).

Following commercialization of HPPD-resistant crops, careful stewardship of HPPD-inhibiting herbicides will be needed to avoid evolution of HPPD-resistant weed biotypes. Currently, resistance to HPPD-inhibiting herbicides has evolved in biotypes of Palmer amaranth and tall waterhemp (Heap 2021). Most notably, a recent North Carolina survey found that nearly 40% of screened Palmer amaranth populations contained survivors following mesotrione applied POST at 105 g ai ha⁻¹ (Mahoney et al. 2020). Confirmation of HPPD-resistant Palmer amaranth in the southeastern United States brings into question the longevity of this MOA.

The main objective of this research was to determine how IFT would integrate into cotton weed management systems in North Carolina. The goal was to evaluate cotton tolerance following PRE and POST applications of IFT alone and in combination with commonly used cotton herbicides in HPPD-resistant cotton.

Materials and Methods

A weed-free experiment was conducted at the Peanut Belt Research Station near Lewiston-Woodville, NC (36.14°N, 77.16°W) in 2019 and 2020. Soil in 2019 was a Goldsboro sandy loam (fine-loamy, siliceous, subactive, thermic Aquic Paleudults) with 1.1% humic matter and pH 6.0, whereas the soil in 2020 was a Lynchburg sandy loam (fine-loamy, siliceous, thermic Aeric Paleaquults) with 0.7% humic matter and pH 6.0 (Mehlich 1984).

Sites were prepared using conventional tillage and then bedded on 91-cm rows, with plots of four 7.6-m rows in 2019 and four 9-m rows in 2020. The experimental design was a randomized complete block with four replications. An experimental 'GLIXTP' cotton cultivar (BASF Corporation, Research Triangle Park, NC) with tolerance to dicamba, IFT, glufosinate, and glyphosate was planted on May 14, 2019, and May 13, 2020. Cotton was seeded at a rate of 107,500 and 123,500 seeds ha⁻¹ at a depth of 2 and 2.5 cm in 2019 and 2020, respectively. Fertilizers, insecticides, growth regulators, and harvest aids were applied in accordance with recommendations from North Carolina Cooperative Extension.

Treatments consisted of IFT alone or in combination, PRE, and EPOST compared to commercial standards. A nontreated control was included for comparison. All treatments, except the nontreated control, received glyphosate + glufosinate + ammonium sulfate (AMS) mid-POST (MPOST). Application dates and rainfall data are presented in Table 1; herbicides, adjuvants, and rates can be found in Table 2; and herbicide treatments can be found in Table 3.

PRE herbicides were applied immediately following planting with a CO₂-pressurized backpack sprayer equipped with flat-fan nozzles (TTI TeeJet® Turbo TeeJet Induction Flat Spray Tips; TeeJet Technologies, Wheaton, IL) delivering 140 L ha⁻¹ at 207 kPa. In 2019, EPOST treatments were applied 31 d after planting (DAP) when cotton was 3- to 4-leaf, whereas MPOST herbicides were applied to 10-leaf cotton 18 d after (DA) EPOST. In 2020, EPOST treatments were applied to 1- to 2-leaf cotton (26 DAP); MPOST herbicides were applied to 6-leaf cotton (15 DA EPOST). POST herbicides were applied using a CO₂-pressurized backpack sprayer delivering 140 L ha⁻¹ equipped with DG 11002 (TeeJet® Drift Guard flat spray nozzles; TeeJet Technologies) and AIXR 11002 (TeeJet® Air Induction Extended Range spray nozzles; TeeJet Technologies) nozzles in 2019 and 2020, respectively. To avoid weed interference influencing cotton development and yield, hand-weeding was employed as necessary.

Table 1. Planting and herbicide application dates and rainfall following preemergence herbicides applied at cotton planting.^a

Year	Planting	Herbicide application date			Rainfall	
	Date	PRE	EPOST	MPOST	0–7 DAP	8–14 DAP
2019	May 14	May 14	June 14	July 18	0.4	0
2020	May 13	May 13	June 8	July 23	1.1	1.6

^aAbbreviations: DAP, days after planting; EPOST, early postemergence; MPOST, mid-postemergence; PRE, preemergence.

Table 2. Herbicides and adjuvants used preemergence and postemergence.^a

Herbicides and adjuvants	Trade names	Formulation	Manufacturer
		concentration	
Acetochlor	Warrant®	360 g ai L ⁻¹	Bayer CropScience
Ammonium sulfate	Amaze Gold®	34%	Loveland Products, Inc.
Crop oil concentrate	Agri-Dex®	99%	Helena Chemical Co.
Dimethenamid-P	Outlook®	719 g ai L ⁻¹	BASF Corporation
Diuron	Direrx® 4L	480 g ai L ⁻¹	Makhteshim Agan of North America
Fluometuron	Cotoran® 4L	480 g ai L ⁻¹	Adama US
Fluridone	Brake®	144 g ai L ⁻¹	SePRO Corporation
Fomesafen sodium salt	Reflex®	240 g ai L ⁻¹	Syngenta Crop Protection
Glufosinate	Liberty® 280 SL	280 g ai L ⁻¹	BASF Corporation
Glyphosate	Roundup PowerMAX® II	660 g ae L ⁻¹	Bayer Crop Science
Isoxaflutole	Alite 27®	479 g ai L ⁻¹	BASF Corporation
Pendimethalin	Prowl® H2O	395 g ai L ⁻¹	BASF Corporation
Pyriithiobac sodium	Staple® LX	384 g ai L ⁻¹	Corteva Agriscience

^aSpecimen labels for each product and mailing addresses and website addresses of each manufacturer can be found at www.cdms.net.

Cotton stand was evaluated 21 DAP by counting all emerged cotton in the two center rows of each plot. Visual estimates of cotton injury (Frans et al. 1986) were collected 9 to 14 DAP, 21 DAP, 7 and 15 to 18 DA EPOST, and 13 to 14 and 27 to 28 DA MPOST. In addition, cotton height was collected 15 DA EPOST and just prior to harvest by measuring the height of 10 plants from the two center rows of each plot. The center two rows of each plot were mechanically harvested and weighed to determine seedcotton yield. Seedcotton grab samples from each plot were collected and ginned using a tabletop gin to calculate lint percentage and thus determine lint yield. To evaluate effects of herbicide treatments on fiber length, fiber length uniformity, fiber strength, and micronaire, 10-g lint subsamples were subjected to high-volume instrumentation analysis (Sasser 1981). All data were subjected to ANOVA using the GLIMMIX procedure of SAS software (version 9.4; SAS Institute Inc., Cary, NC) and means were separated using Fisher's protected LSD at $P = 0.05$ where appropriate. Main effects of herbicide treatment and year-by-herbicide treatment interactions were observed only for visual estimates of cotton injury. Therefore, all data, except cotton injury, are presented pooled over years.

Results and Discussion

Cotton Stand

Cotton stand in the nontreated control was 10 plants m row⁻¹ 21 DAP (Table 3). Cotton stand in plots treated with a residual herbicide PRE ranged from 9 to 10 plants m row⁻¹. Residual herbicides applied PRE, including the grower standard of acetochlor + diuron + fomesafen, IFT alone, and IFT + acetochlor, diuron, diuron +

pendimethalin, fluometuron, fluridone, fomesafen, and pyriithiobac, had no effect on cotton stand.

Cotton Response

Year-by-herbicide treatment interactions for cotton injury 9 to 14 DAP, 7 and 15 to 18 DA EPOST were significant; therefore, data are presented by year (Table 4). The grower standard PRE treatment, acetochlor + diuron + fomesafen (4%), caused the greatest injury 9 to 14 DA PRE in 2019. At this same time, only IFT + diuron and IFT + diuron + pendimethalin caused similar injury to cotton as that of acetochlor + diuron + fomesafen. All other treatments, including IFT alone (0%), were less injurious than the grower standard applied PRE. Cotton injury observed PRE in 2020 was less than in 2019 and ranged just 0% to 2%. It is important to note, like 2019, IFT alone caused 0% cotton injury 9 to 14 DA PRE. Cotton injury observed in this study is similar to that observed by Cahoon et al. (2015a) and Foster (2021). Previous research carried out in North Carolina found that cotton was injured $\leq 3\%$ 18 to 27 DA PRE by acetochlor, diuron, fomesafen, and fluometuron and two- and three-way combinations of the residual herbicides (Cahoon et al. 2015a). Foster (2021) evaluated HPPD-resistant cotton in Texas and reported 1% injury 14 DA PRE from IFT + fluometuron and IFT + pendimethalin.

Despite little cotton response to IFT and other PRE herbicides in 2019 and 2020, PRE rates required for effective weed control can sometimes injure cotton, especially when growing conditions are less than ideal (Schrage et al. 2012). Culpepper et al. (2012) reported that seedling vigor, rainfall/irrigation intensity, and/or planting depth influenced cotton response to diuron, fomesafen, pyriithiobac, and pendimethalin in Georgia. For example, under

Table 3. Cotton stand as affected by isoxaflutole alone and isoxaflutole combinations applied preemergence and early postemergence.^a

Herbicides and application times ^{b,c}		Cotton stand 21 DAP plants m row ⁻¹
PRE	EPOST ^d	
None	None	10 NS
Acet + diuron + fome	dimet + gluf	9
IFT + diuron	dimet + gluf	10
IFT + pendi	dimet + gluf	10
IFT + diuron + pendi	dimet + gluf	10
IFT	dimet + gluf	10
IFT + acet	dimet + gluf	10
IFT + fome	dimet + gluf	10
IFT + fluometuron	dimet + gluf	10
IFT + fluridone	dimet + gluf	10
IFT + pyriithiobac	dimet + gluf	10
None ^e	IFT LR ^f	10
None ^e	IFT HR ^f	10
None ^e	IFT HR + glyph	10
None	IFT HR + gluf	10
IFT	IFT HR ^f	10

^aData are averaged over 2 yr. Means within a column followed by the same letter are not different according to Fisher's protected LSD test at $P = 0.05$. For columns beginning with NS, means are not statistically different.

^bAbbreviations: Acet, acetochlor; Fome, fomesafen; IFT, isoxaflutole; Pendi, pendimethalin; Dimet, dimethenamid-*P*; Gluf, glufosinate; Glyph, glyphosate; NS, no significant difference; PRE, preemergence; EPOST, early postemergence; DAP, days after planting; HR, high rate; LR, low rate.

^cAcetochlor, diuron, fomesafen, fluometuron, fluridone, isoxaflutole, pendimethalin, and pyriithiobac applied at 1,261, 560, 210, 1,121, 168, 105, 925, and 59 g ha⁻¹ PRE. Dimethenamid-*P*, glufosinate, isoxaflutole low rate, and isoxaflutole high rate applied at 841, 881, 53, and 105 g ha⁻¹ EPOST. Glyphosate applied at 1,734 g ae ha⁻¹ EPOST.

^dAll treatments, except the nontreated control, received glufosinate at 881 g ha⁻¹ plus glyphosate at 1,734 g ae ha⁻¹ mid-POST.

^eTreatments received no PRE herbicide in 2019, and acetochlor at 1,261 g ha⁻¹ PRE in 2020.

^fIsoxaflutole applied alone EPOST included crop oil concentrate at 1% vol/vol; no adjuvant was included when IFT was applied in combination with glufosinate or glyphosate.

Glufosinate and/or IFT applied EPOST included ammonium sulfate at 2.5 g ha⁻¹.

ideal conditions (normal planting depth and normal irrigation), cotton was injured 4% by fomesafen, whereas the herbicide caused 33% to 41% injury under adverse conditions (i.e., shallow planting depth and intense irrigation). Similarly, Main et al. (2012) observed up to 24% reduction in cotton stand 5 to 7 DA after a PRE application of fomesafen when rainfall coincided with cotton emergence. Because various factors including seedling vigor, environmental conditions, and planting depth can influence injury from residual herbicides, further evaluation of HPPD-resistant cotton across varying environments and production practices is warranted.

In 2019, cotton injury 7 DA EPOST was minimal across all treatments (0% to 5%), but cotton injury was greater from dimethenamid-*P* + glufosinate (3% to 5%), regardless of PRE treatment, compared to low rate of IFT (LR; 53 g ha⁻¹) or high rate of IFT (HR; 105 g ha⁻¹), which injured cotton 1% and 2%, respectively. Note that dimethenamid-*P* + glufosinate was applied only to treatments receiving a PRE, whereas IFT HR applied EPOST followed no PRE or IFT PRE. However, we can directly compare IFT PRE followed by (fb) dimethenamid-*P* + glufosinate or IFT HR EPOST. Following IFT PRE, dimethenamid-*P* + glufosinate EPOST injured cotton 4% 7 DA EPOST compared to only 2% injury from IFT PRE fb IFT HR EPOST. In the absence of a PRE, IFT HR + glyphosate (0%) and IFT HR + glufosinate (2%) were no more injurious than IFT alone 7 DA EPOST. IFT alone applied EPOST included crop oil concentrate and AMS; when IFT was applied in combination with glyphosate or glufosinate, only AMS was added. Regardless of adjuvant or herbicide partner, cotton response to IFT applied EPOST was

minimal. In 2020, similar trends in cotton injury were observed. Regardless of IFT rate or herbicide combination, cotton was injured 0% to 1%, with the exception of IFT HR + glufosinate, which resulted in 4% injury to cotton. No injury was observed from IFT PRE fb IFT HR EPOST in 2020, again confirming the safety of IFT when applied both PRE and EPOST. Notably, injury from dimethenamid-*P* + glufosinate was greater in 2020 than 2019 and ranged from 6% to 9%. When making a direct comparison between IFT PRE fb dimethenamid-*P* + glufosinate or IFT HR EPOST, cotton injury from dimethenamid-*P* + glufosinate (7%) again exceeded injury from IFT HR (0%) EPOST. Conditions were favorable for cotton growth in both 2019 and 2020; however, increased cotton response to dimethenamid-*P* + glufosinate in 2020 may be attributed to higher air temperature and relative humidity, and greater soil moisture at the time of EPOST applications (data not shown). Cahoon and York (2020) noted increased cotton response to *S*-metolachlor, another VLCFA inhibitor (Weed Science Society of America Group 15), when applications coincide with high temperatures and increased soil moisture.

In 2019, injury 15 to 18 DA EPOST ranged between 2% and 3% across all treatments and was not significant. Contrarily, herbicide treatments did affect cotton injury at this timing in 2020, though injury was less than that observed 7 DA EPOST. Dimethenamid-*P* + glufosinate resulted in 2% to 4% injury to cotton, whereas IFT applied EPOST resulted in just 0% to 2% injury to cotton. The lower and higher rates of IFT applied EPOST resulted in 0% and 1% cotton injury 15 to 18 DA EPOST, respectively. IFT HR + glyphosate (0%) and IFT + glufosinate (2%) applied at the same timing also caused little injury. Additionally, IFT PRE fb IFT HR EPOST again caused minimal injury (1%) and resulted in less injury than cotton treated with IFT PRE fb dimethenamid-*P* + glufosinate EPOST (3%). When cotton was 6- to 10-leaf (MPOST), glyphosate + glufosinate was applied to all treatments, except the nontreated control (data not shown), injuring cotton $\leq 1\%$.

Cotton response to EPOST treatments containing dimethenamid-*P* is unsurprising. EPOST applications of VLCFA inhibitors can cause cotton injury, but even in cases in which moderate or severe early season injury has been documented, cotton lint yield remains unaffected (Cahoon et al. 2014; Eure et al. 2013; Everman et al. 2007, 2009; Inman et al. 2014; Samples 2020). In Texas, Foster (2021) observed similar response of HPPD-resistant cotton to EPOST applications of dimethenamid-*P* + glufosinate, which resulted in 6% to 10% cotton injury 14 DA EPOST across 4-site years. However, Foster (2021) reported greater cotton injury following IFT + glufosinate (6% to 9%) applied EPOST compared to 2% to 4% injury from the combination in this study.

Cotton Height

Cotton in the nontreated control plots averaged 19 cm in height 15 DA EPOST; all herbicide treatments resulted in reduced cotton height compared to the nontreated control at that time. Cotton in plots treated with IFT PRE fb dimethenamid-*P* + glufosinate, acetochlor fb IFT LR, and IFT fb IFT HR averaged 18 cm in height. Cotton was shortest in plots treated with pendimethalin PRE (14 cm). In general, residual combinations applied PRE fb dimethenamid-*P* resulted in shorter cotton compared to treatments that did not include dimethenamid-*P*. Despite early season differences in height, cotton recovered, and by the date that harvest aids were applied, no difference in cotton height was observed. Final cotton height across all treatments ranged 75 to 81 cm.

Table 4. Cotton injury from isoxaflutole alone and isoxaflutole combinations applied preemergence and early postemergence.^a

Herbicides and application times ^{b,c}		Cotton injury					
		2019			2020		
		9 to 14 DA	7 DA	15 to 18 DA	9 to 14 DA	7 DA EPOST	15 to 18 DA
		PRE	EPOST	EPOST	PRE	EPOST	EPOST
PRE	EPOST ^d				—%		
Acet + diuron + fome	dimet + gluf	4 a	5 a	2	2 a	8 ab	4 a
IFT + diuron	dimet + gluf	3 ab	4 b	2	1 b	6 bc	3 ab
IFT + pend	dimet + gluf	2 b	5 a	2	0 c	9 a	4 a
IFT + diuron + pend	dimet + gluf	3 ab	3 c	2	0 c	7 ab	2 bc
IFT	dimet + gluf	1 cd	4 b	3	0 c	7 ab	3 ab
IFT + acet	dimet + gluf	1 cd	3 c	2	1 b	7 ab	3 ab
IFT + fome	dimet + gluf	1 cd	4 b	2	2 a	7 ab	3 ab
IFT + fluometuron	dimet + gluf	1 cd	4 b	2	0 c	7 ab	4 a
IFT + fluridone	dimet + gluf	2 bc	4 b	2	1 b	6 bc	4 a
IFT + pyriithiobac	dimet + gluf	1 cd	4 b	2	0 c	7 ab	4 a
None ^e	IFT LR ^f	—	2 d	3	—	0 d	0 d
None ^e	IFT HR ^f	—	1 e	3	—	1 d	1 c
None ^e	IFT HR + glyph	—	0 f	2	—	1 d	0 d
None	IFT HR + gluf	—	2 d	2	—	4 c	2 bc
IFT	IFT HR ^f	0 d	2 d	2	0 c	0 d	1 c

^aData are averaged over 2 yr. Means within a column followed by the same letter are not different according to Fisher's protected LSD test at $P = 0.05$. For columns beginning with NS, means are not statistically different.

^bAbbreviations: Acet, acetochlor; Fome, fomesafen; IFT, isoxaflutole; Pend, pendimethalin; Dimet, dimethenamid-P; Gluf, glufosinate; Glyph, glyphosate; PRE, preemergence; EPOST, early postemergence; HR, high rate; LR, low rate.

^cAcetochlor, diuron, fomesafen, fluometuron, fluridone, isoxaflutole, pendimethalin, and pyriithiobac applied at 1,261, 560, 210, 1,121, 168, 105, 925, and 59 g ha⁻¹ PRE. Dimethenamid-P, glufosinate, isoxaflutole low rate, and isoxaflutole high rate applied at 841, 881, 53, and 105 g ha⁻¹ EPOST. Glyphosate applied at 1,734 g ae ha⁻¹ EPOST.

^dAll treatments, except the nontreated control, received glufosinate at 881 g ha⁻¹ plus glyphosate at 1,734 g ae ha⁻¹ mid-POST.

^eTreatments received no PRE herbicide in 2019, and acetochlor at 1,261 g ha⁻¹ PRE in 2020.

^fIsoxaflutole applied alone EPOST included crop oil concentrate at 1% vol/vol; no adjuvant was included when IFT was applied in combination with glufosinate or glyphosate. Glufosinate and/or IFT applied EPOST included ammonium sulfate at 2.5 kg ha⁻¹.

Table 5. Cotton height and yield as affected by isoxaflutole alone and isoxaflutole combinations applied preemergence and early postemergence.^a

Herbicides and application times ^{b,c}		Cotton height		Yield
		15 DA EPOST height	Final plant height	
PRE	EPOST ^d	—cm		kg ha ⁻¹
None	None	19 a	80	1,470
Acet + diuron + fome	dimet + gluf	15 e	78	1,340
IFT + diuron	dimet + gluf	16 d	76	1,370
IFT + pend	dimet + gluf	14 f	75	1,370
IFT + diuron + pend	dimet + gluf	14 f	76	1,300
IFT	dimet + gluf	18 b	81	1,440
IFT + acet	dimet + gluf	16 d	76	1,340
IFT + fome	dimet + gluf	16 d	76	1,380
IFT + fluometuron	dimet + gluf	17 c	75	1,430
IFT + fluridone	dimet + gluf	17 c	77	1,400
IFT + pyriithiobac	dimet + gluf	16 d	76	1,340
None ^e	IFT LR ^f	18 b	77	1,440
None ^e	IFT HR ^f	17 c	79	1,470
None ^e	IFT HR + glyph	17 c	76	1,470
None	IFT HR + gluf	17 c	76	1,380
IFT	IFT HR ^f	18 b	81	1,430

^aData are averaged over 2 yr. Means within a column followed by the same letter are not different according to Fisher's protected LSD test at $P = 0.05$. For columns beginning with NS, means are not statistically different.

^bAbbreviations: Acet, acetochlor; Fome, fomesafen; IFT, isoxaflutole; Pend, pendimethalin; Dimet, dimethenamid-P; Gluf, glufosinate; Glyph, glyphosate; PRE, preemergence; EPOST, early postemergence; HR, high rate; LR, low rate.

^cAcetochlor, diuron, fomesafen, fluometuron, fluridone, isoxaflutole, pendimethalin, and pyriithiobac applied at 1,261, 560, 210, 1,121, 168, 105, 925, and 59 g ha⁻¹ PRE. Dimethenamid-P, glufosinate, isoxaflutole low rate, and isoxaflutole high rate applied at 841, 881, 53, and 105 g ha⁻¹ EPOST. Glyphosate applied at 1,734 g ae ha⁻¹ EPOST.

^dAll treatments, except nontreated control, received glufosinate at 881 g ha⁻¹ plus glyphosate at 1,734 g ae ha⁻¹ mid-POST.

^eTreatments received no PRE herbicide in 2019 and acetochlor at 1,261 g ha⁻¹ PRE in 2020.

^fIsoxaflutole applied alone EPOST included crop oil concentrate at 1% vol/vol; no adjuvant was included when IFT was applied in combination with glufosinate or glyphosate. Glufosinate and/or IFT applied EPOST included ammonium sulfate at 2.5 kg ha⁻¹.

Cotton Yield

Cotton lint yield ranged from 1,300 to 1,470 kg ha⁻¹ (Table 5), and no treatment differences were observed. Like cotton yield, herbicide treatments did not affect micronaire, fiber length, fiber length uniformity, or fiber strength (data not shown).

The HPPD-resistant cotton cultivar used in this experiment demonstrated excellent tolerance to IFT PRE and EPOST. IFT, applied alone or in combination with other herbicides, caused minimal cotton injury, thus demonstrating potential for safe use at either timing. These findings are further reinforced by research conducted in Arkansas and Texas, which also reported no decrease in cotton stand, minimal injury, and no reductions in lint yield from IFT applied PRE or EPOST when used alone or combined with other herbicides (Fleming et al. 2021; Foster 2021). Additionally, the experimental cotton cultivar demonstrated tolerance to commonly used cotton herbicides PRE and POST.

Despite considerable evidence that HPPD-resistant cotton responds minimally to IFT applied PRE and EPOST at rates up to 105 g ha⁻¹, further research is warranted to explore the response of HPPD-resistant cultivars to higher rates of IFT. Presently, 105 g ha⁻¹ is expected to be the 1× rate of IFT for use in cotton (J Sanderson, BASF Corporation, personal communication); however, evaluation of cotton response exceeding this rate is needed to ensure a considerable margin of crop safety is present.

Although further research is required, tolerance of the experimental HPPD-resistant cultivar observed in this experiment indicates that IFT can likely be integrated safely into cotton weed management systems and used alongside common PRE and POST cotton herbicides with minimal risk to producers, while providing another effective MOA for controlling troublesome weeds in cotton.

Acknowledgments. Funding for this work was provided by BASF. No conflicts of interest have been declared.

References

- Anonymous (2019a) Brake herbicide. Carmel, IN: SePRO Corporation. 4 p
- Anonymous. (2019b) Balance flexx herbicide. St. Louis, MO: Bayer CropScience. 21 p
- Anonymous. (2019c) Corvus herbicide. St. Louis, MO: Bayer CropScience. 21 p
- Anonymous. (2019d) Alite 27 herbicide. Research Triangle Park, NC: BASF Corporation. 18 p
- Armel GR (2002) Weed management in conventional, no-till, and transgenic corn with mesotrione combinations and other herbicides. Ph.D. dissertation. Blacksburg, VA: Virginia Tech University. 17 p
- Bartels PG, Watson CW (1978) Inhibition of carotenoid synthesis by fluridone and norflurazon. *Weed Sci* 26:198–203
- Beckie HJ, Harker KN (2017) Our top 10 herbicide-resistant weed management practices. *Pest Manage Sci* 73:1045–1052
- Bhowmik PC, Kushwaha S, Mitra S (1999) Response of various weed species and corn (*Zea mays*) to RPA 201772. *Weed Technol* 13:504–509
- Brabham C, Norsworthy JK, Houston MM, Varanasi VK, Barber T (2019) Confirmation of S-metolachlor resistance in palmer amaranth (*Amaranthus palmeri*). *Weed Technol* 33:720–726
- Bond JA, Oliver LR, Stephenson DO (2006) Response of Palmer Amaranth (*Amaranthus palmeri*) Accessions to Glyphosate, Fomesafen, and Pyriithobac. *Weed Technol* 20:885–892
- Boudec P, Rodgers M, Dumas F, Sailland A, Bourdon H; Aventis Crop Science, assignee (2001) November 7. Mutated hydroxyphenylpyruvate dioxxygenase, DNA sequence and isolation of plants which contain such a gene, and which are tolerant to herbicides. U.S. patent 6,245,968 B1
- Busi R, Powles SB, Beckie HJ, Renton M (2020) Rotations and mixtures of soil-applied herbicides delay resistance. *Pest Manage Sci* 76:487–496
- Cahoon CW, York AC (2020) Weed management in cotton. Pages 85–135 in 2020 cotton information. Raleigh: North Carolina Cooperative Extension
- Cahoon CW, York AC, Jordan DL, Braswell LR (2014) Chloroacetamide tank mixes with pyriithobac in glyphosate- and glufosinate-based herbicide systems. Pages 1058–1060 in Proceedings of the Beltwide Cotton Conference, National Cotton Council of America. New Orleans, Louisiana, January 6–8, 2014
- Cahoon CW, York AC, Jordan DL, Everman WJ, Seagroves RW, Braswell LR, Jennings KM (2015a) Weed control in cotton by combinations of microencapsulated acetochlor and various residual herbicides applied preemergence. *Weed Technol* 29:740–750
- Cahoon CW, York AC, Jordan DL, Seagroves RW, Everman WJ, Jennings KM (2015b) Fluridone carryover to rotational crops following application to cotton. *J Cotton Sci* 1:631–640
- Chamovitz D, Sandmann G, Hirschberg J (1993) Molecular and biochemical characterization of herbicide-resistant mutants of cyanobacteria reveals the phytoene desaturation is a rate-limiting step in carotenoid biosynthesis. *J Biol Chem* 268:17348–17353
- Copeland JD, Giacomini DA, Tranel PJ, Montgomery GB, Steckel LE (2018) Distribution of PPX2 mutations conferring PPO-inhibitor resistance in palmer amaranth populations of Tennessee. *Weed Technol* 32:592–596
- Copeland DJ, Montgomery GB, Steckel LE (2019) Evaluation of the time-of-day effect of herbicides applied POST on protoporphyrinogen IX oxidase-resistant and -susceptible palmer amaranth (*Amaranthus palmeri*). *Weed Technol* 33:651–657
- Culpepper AS (2019) Weed management in cotton. Pages 14–17 in 2019 Georgia cotton production guide. J. Whitaker, ed. Athens: University of Georgia Extension
- Culpepper AS, Moore T, Ethredge R, Briggs W (2012) Cotton injury as influenced by herbicides, irrigation, seedling vigor, seeding depth, and environmental stresses. Pages 1515–1516 in Proceedings of the 2012 Beltwide Cotton Conferences, National Cotton Council of America. Orlando, Florida, January 3–6, 2012
- Culpepper AS, Webster TM, Sosnoskie LM, York AC, Nandula VK (2010) Glyphosate-resistant Palmer amaranth in the United States. Pages 195–212 in Glyphosate Resistance in Crops and Weeds: History, Development, and Management. Hoboken, NJ: John Wiley & Sons
- Dayan FE (2019) Current status and future prospects in herbicide discovery. *Plants* 8:341
- Duke SO (2012) Why have no new herbicide modes of action appeared in recent years? *Pest Manage Sci* 68:505–512
- Duke SO, Heap I (2017) Evolution of weed resistance to herbicides. What have we learned after 70 years? Pages 63–86 in Biology, physiology and molecular biology of weeds. M. Jugulam, ed. Boca Raton, FL: CRC Press.
- Eure PM, Culpepper AS, Merchant RM (2013) An assessment of cotton tolerance to pyroxasulfone, acetochlor, and S-metolachlor. Pages 660–661 in Proceedings of the 2013 Beltwide Cotton Conferences, National Cotton Council of America. San Antonio, Texas, January 7–10, 2013
- Everman WJ, Burke IC, Allen JR, Collins J, Wilcut JW (2007) Weed control and yield with glufosinate-resistant cotton weed management systems. *Weed Technol* 2:695–701
- Everman WJ, Clewis SB, York AC, Wilcut JW (2009) Weed control and yield with flumioxazin, fomesafen, and S-metolachlor systems for glufosinate-resistant cotton residual weed management. *Weed Technol* 23:391–397
- Foster DC (2021) Crop response, weed management systems, and tank mix partners with isoxaflutole in HPPD tolerant cotton. M.S. Thesis. Lubbock, TX: Texas Tech University. 109 p
- Fleming JA, Norsworthy JK, Barber LT, Farr RB (2021) Evaluating tolerance of HPPD-tolerant cotton to PRE and POST applications of isoxaflutole. Proceedings of the Beltwide Cotton Conference, National Cotton Council of America. Virtual, January 5–7, 2021
- Frans RE, Talbert R, Marx D, Crowley H (1986) Experimental design and techniques for measuring and analyzing plant responses to weed control practices. Pages 29–46 in N.D. Camper, ed. Research Methods in Weed Science. Champaign, IL: South Weed Science Society
- Garcia AM, Hoos AB, Terziotti S (2011) A regional modeling framework of phosphorus sources and transport in streams of the southeastern United States. *J Am Water Resour Assoc* 47:991–1010

- Gianessi LP (2008) Economic impacts of glyphosate-resistant crops. *Pest Manage Sci* 64:346–352
- Heap I (2021) Cotton herbicide resistant weeds globally. <http://www.weedscience.org/Pages/Crop.aspx>. Accessed: September 25, 2020
- Inman M, Jordan DL, Edmisten K, York AC, Bachelor J, Wells R (2014) Cotton and weed response to combinations of glyphosate and glufosinate applied with acephate and 116 residual herbicides Page 1044 in Proceedings of the Beltwide Cotton Conference, National Cotton Council of America. New Orleans, Louisiana, January 6–8, 2014
- Knezevic SZ, Sikkema PH, Tardif F, Hamill AS, Chandler K, Swanton CJ (1998) Biologically effective dose and selectivity of RPA 201772 for preemergence weed control in corn (*Zea mays*). *Weed Technol* 12:670–676
- Kniss AR (2018) Genetically engineered herbicide-resistant crops and herbicide-resistant weed evolution in the United States. *Weed Sci* 66:260–273
- Kowalczyk-Schroder S, Sandmann G (1992) Interference of fluridone with the desaturation of phytoene by membranes of the cyanobacterium *Aphanocapsa*. *Pestic Biochem Phys* 42:7–12
- Mahoney DJ, Jordan DL, Roma-Burgos N, Jennings KM, Leon RG, Vann MC, Everman WJ, Cahoon CW (2020) Susceptibility of Palmer amaranth (*Amaranthus palmeri*) to herbicides in accessions collected from the North Carolina Coastal Plain. *Weed Sci* 68:582–593
- Main CL, Faircloth JC, Steckel LE, Culpepper AS, York AC (2012) Cotton tolerance to fomesafen applied preemergence. *J Cotton Sci* 16:80–87
- Matringe M, Sailland A, Pelissier B, Rolland A, Zink O (2005) p-Hydroxyphenylpyruvate dioxygenase inhibitor-resistant plants. *Pest Manage Sci* 61:269–276
- Mehlich A (1984) Photometric determination of humic matter in soils, a proposed method. *Commun Soil Sci Plant Anal* 15:1417–1422
- NASA [National Aeronautics and Space Administration] (2016) Soil composition across the U.S. <https://earthobservatory.nasa.gov/images/87220/soil-composition-across-the-us>. Accessed: September 29, 2020
- Neve P, Norsworthy JK, Smith KL, Zelaya IA (2011) Modeling glyphosate resistance management strategies for palmer amaranth (*Amaranthus palmeri*) in cotton. *Weed Technol* 25:335–343
- Norsworthy JK, Griffith G, Griffin T, Bagavathiannan M, Gbur EE (2014) In-field movement of glyphosate-resistant palmer amaranth (*Amaranthus palmeri*) and its impact on cotton lint yield: evidence supporting a zero-threshold strategy. *Weed Sci* 62:237–249
- Samples CA (2020) Evaluation of pesticide application technology in cotton production. Ph.D. dissertation. Starkville, MS: Mississippi State University. 150 p
- Sasser PE (1981) Basics of high volume instruments for fiber testing. In Proceedings of the Beltwide Cotton Production Research Conferences, National Cotton Council of America. New Orleans, Louisiana, January 6–8, 1981
- Schrage BW, Norsworthy JK, Smith KL, Johnson DB, Bagavathiannan MV, Riar DS (2012) Factors contributing to cotton injury from soil-applied residual herbicides. Pages 102–106 in Summaries of Cotton Research 2012. Fayetteville: University of Arkansas Research and Extension
- Smith A (2019) Annual weed control in isoxaflutole-resistant soybean. M.S. Thesis. Guelph, Ontario, Canada: The University of Guelph. 232 p
- Starkey CE, Norsworthy JK, Schwartz LM (2016) Use of HPPD-inhibiting herbicides for control of troublesome weeds in the mid-southern United States. *Adv Crop Sci Technol* 4:1–8
- Stephenson DO, Bond JA (2012). Evaluation of thien carbazono-methyl- and isoxaflutole-based herbicide programs in corn. *Weed Technol* 26:37–42
- Takano HK, Oliveira RJ, Constantin J, Silva V, Mendes RR (2018) Chemical control of glyphosate-resistant goosegrass. *Planta Daninha*, 36. doi: <https://www.scielo.br/j/pda/9CBNtHPPYrSFpQgS76wTr/?lang=en#:~:text=The%20use%20of%20residual%20herbicides,effective%20in%20controlling%20this%20species>. Accessed: September 27, 2020
- [USDA-APHIS] U.S. Department of Agriculture–Animal and Plant Health Inspection Service (2017) Bayer Crop Science. Petition for a determination of nonregulated status for herbicide tolerant cotton transformation event GHB811. https://www.aphis.usda.gov/brs/aphisdocs/17_13801p.pdf. Accessed: September 27, 2020
- [USDA-NASS] U.S. Department of Agriculture–National Agricultural Statistics Service (2020) Adoption of genetically engineered crops in the U.S. <https://www.ers.usda.gov/data-products/adoption-of-genetically-engineered-crops-in-the-us/>. Accessed: September 27, 2020
- [USDA-NASS] U.S. Department of Agriculture–National Agricultural Statistics Service (2019) Upland cotton 2019 production by county for selected states. https://www.nass.usda.gov/Charts_and_Maps/Crops_County/ctu-pr.php. Accessed: September 27, 2020
- Van Wychen L (2019) 2019 survey of the most common and troublesome weeds in broadleaf crops, fruits & vegetables in the United States and Canada. Weed Science Society of America national weed survey dataset. http://wssa.net/wp-content/uploads/2019-Weed-Survey_broadleaf-crops.xlsx. Accessed: September 27, 2020
- Varanasi VK, Brabham C, Norsworthy JK, Nie H, Young BG, Houston M, Scott RC (2018) A statewide survey of PPO-inhibitor resistance and the prevalent target-site mechanisms in palmer amaranth (*Amaranthus palmeri*) accessions from Arkansas. *Weed Sci* 66:149–158
- Vrabel TE, Streigel WL, Lavoy JD (1996) Efficacy of isoxaflutole as a burndown treatment in no-till corn. Page 67 in Proceedings of the North Central Weed Science Society. St. Louis, Missouri, December, 1996
- Ward SM, Webster TM, Steckel LE (2013) Palmer amaranth (*Amaranthus palmeri*): A review. *Weed Technol* 27:12–27
- Whitaker JR, York AC, Jordan DL, Culpepper AS, Sosnoskie LM (2011) Residual herbicides for Palmer amaranth control. *J Cotton Sci* 15:89–99
- Whitaker JR, York AC, Jordan DL, Culpepper AS (2010) Palmer amaranth (*Amaranthus palmeri*) control in soybean with glyphosate and conventional herbicide systems. *Weed Technol* 24:403–410
- Wicks GA, Knezevic SZ, Bernards M, Wilson RG, Klein RN, Martin AR (2007) Effect of planting depth and isoxaflutole rate on corn injury in Nebraska. *Weed Technol* 21:642–646
- Wiggins MS, Hayes RM, Steckel LE (2016) Evaluating cover crops and herbicides for glyphosate-resistant Palmer amaranth (*Amaranthus palmeri*) control in cotton. *Weed Technol* 30:415–422
- Wilcut JW, York AC, Jordan DL (1995) Weed management systems for oilseed crops Pages 355–358 in AE Smith, ed. Handbook of weed management systems. New York: Marcel-Dekker
- Wilson RG, Wicks GA, Klein RN, Roeth FW, Knezevic S, Martin AR (1999) Factors affecting isoxaflutole injury to corn in Nebraska: environment. Page 82 in Proceedings of the North Central Weed Science Society. Columbus, OH: North Central Weed Science Society
- York AC (2014) Brake F2 section 18 emergency exemption for cotton <https://cotton.ces.ncsu.edu/2014/03/brake-f2-section-18-emergency-exemption-for-cotton/>. Accessed: November 27, 2020
- York AC (2016) New or almost new herbicides for 2016 <https://cotton.ces.ncsu.edu/2016/04/new-or-almost-new-herbicides-for-2016-york/>. Accessed: November 27, 2020
- Young BG, Hart SE (1998) Optimizing foliar activity of isoxaflutole on giant foxtail (*Setaria faberi*) with various adjuvants. *Weed Sci* 46:397–402
- Zhao N, Zuo L, Li W, Guo W, Liu W, Wang J (2017) Greenhouse and field evaluation of isoxaflutole for weed control in maize in china. *Sci Rep* 7:12690