

Pesticide use and adult-onset asthma among male farmers in the Agricultural Health Study

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Abbreviations:

AHS Agricultural Health Study
BMI body mass index
CI confidence interval
HPEE High Pesticide Exposure Event
OR odds ratio
PPE Personal Protective Equipment

Abstract:

Although specific pesticides have been associated with wheeze in farmers, little is known about pesticides and asthma. We used data from 19,704 male farmers in the Agricultural Health Study to evaluate lifetime use of 48 pesticides and prevalent adult-onset asthma, defined as doctor-diagnosed asthma after age 20. We categorized asthma cases as allergic (N=127) and non-allergic (N=314) based on their history of eczema or hayfever. We used polytomous logistic regression controlling for age, state, smoking, and body mass to assess pesticide associations.

High pesticide exposure events were associated with a doubling of both allergic and non-allergic asthma. For ever use, 12 individual pesticides were associated with allergic asthma and four with non-allergic asthma. For allergic asthma, coumaphos (odds ratio (OR) =2.34, 95% Confidence Interval (CI) =1.49,3.70), heptachlor (OR=2.01, 95%CI=1.30,3.11), parathion (OR=2.05, 95%CI=1.21,3.46), 80/20 mix (carbon tetrachloride/carbon disulfide) (OR=2.15, 95%CI=1.23,3.76) and ethylene dibromide (OR=2.07, 95%CI=1.02,4.20), all had odds ratios greater than 2.0 and significant exposure-response trends. For non-allergic asthma, DDT had the strongest association (OR=1.41, 95%CI=1.09,1.84) but with little evidence of increasing asthma with increasing use. Current animal handling and farm activities did not confound these results. We saw little evidence that allergy alone was driving these associations.

Pesticides may be an overlooked contributor to asthma risk among farmers.

Keywords: allergy, farming, occupational exposure, pesticides, respiratory disease

Introduction

Pesticides may contribute to asthma among farmers (1), farm women (2), and insecticide applicators (3), but data for individual pesticides are limited to a few studies. Pesticide use, particularly use of organophosphate insecticides, has been associated with wheeze among US farmers (4), US commercial pesticide applicators (5), Kenyan farmworkers (6), and with atopic asthma in farm women (2). In guinea pigs, organophosphate insecticides induce airway hyperreactivity at doses below those causing acetylcholinesterase inhibition (7, 8). These effects are stronger in allergen-sensitized animals (9).

Farmers are more likely to be diagnosed with non-atopic asthma than atopic asthma when compared to other occupational groups (10). This observation along with evidence of differential response to pesticides in allergen-sensitized animals (9) indicates the importance of evaluating risk factors asthma separately based on atopic status. In a cross-sectional analysis of enrollment data from the Agricultural Health Study (AHS), we evaluated the association of pesticide use with allergic and non-allergic asthma in farmers. We used self-reported allergy as a surrogate for atopy because we lacked clinical measurement of atopy.

Materials and Methods

The AHS is a large, prospective study of Iowa and North Carolina pesticide applicators and their spouses (11). The study enrolled over 52,000 licensed private pesticide applicators, mostly farmers, from 1993 to 1997. After completing the enrollment questionnaire, 22,916 (44%) applicators returned a second mailed questionnaire; applicators who did or did not return this second questionnaire were similar regarding demographics, farming practices, and medical history (12). Our analysis was limited to private applicators who returned both questionnaires

because information on respiratory disease was obtained in the second questionnaire.

Questionnaires are available at www.aghealth.org/questionnaires.html.

Participants provided information on medical history and potential confounders, including smoking, body mass, respiratory and allergic symptoms, and other respiratory diseases, as well as detailed information on their farming practices. Applicators provided detailed information on lifetime pesticide use for 50 pesticides (ever use, frequency of use, number of years used, and decade of first use). Individuals also provided information on their pesticide use practices, including application method, mixing, equipment repair, and use of personal protective equipment (PPE). We used this information to create two metrics of lifetime pesticide use: lifetime days of use and intensity-adjusted lifetime days of use. The intensity-adjusted metric accounted for potential differential exposure to pesticides based on application methods, mixing habits, repairs, and use of PPE (13). For general farming exposures, we collected information on current crops and animals raised as well as current farm tasks and maintenance activities.

In addition to pesticide application information, we also had two measures indicative of ever having elevated pesticide exposure: pesticide poisoning and high pesticide exposure events (HPEE). Pesticide poisoning was based on a self-report of doctor diagnosis of pesticide poisoning. HPEE was based on the response to the question “Have you ever had an incident or experience while using any type of pesticide which caused you unusually high personal exposure?”

All male private applicators age 20 and older with complete information on smoking, asthma history, age, body mass index (BMI), and HPEE were included in this analysis; female applicators were excluded because there were too few cases to evaluate (N=487 female applicators, 19 asthma cases). We included as cases those subjects who reported a doctor-

diagnosis of asthma after age 19 and then further subdivided the cases by allergic status based on a history of doctor-diagnosed eczema or hayfever. Given our interest in adult asthma and occupational exposures, we excluded individuals diagnosed with asthma before age 20 (N=545).

We used polytomous logistic regression to evaluate associations between the farming exposures and prevalent adult-onset asthma. This allowed us to assess separate associations for allergic and non-allergic asthma simultaneously and formally test differences between odds ratios. Using goodness-of-fit tests, we developed a base model that consisted of age (4 categories: 20-39, 40-49, 50-59, 60+), state (IA, NC), smoking status (current, past, never), and BMI (<25, 25-29, 30+). We assessed each exposure individually adjusting for base-model covariates. We restricted our analysis to those exposures with at least five exposed cases. We also evaluated exposure response by creating a three-level variable for each pesticide. For each pesticide, we split the distribution of intensity-adjusted days among those who had ever used the pesticide at the median to create two exposed groups (\leq median, $>$ median); never users were the referent group. We used these ordinal categories for chi-square tests for trend.

To evaluate potential confounding by related exposures, we first assessed pairwise Spearman correlations between those exposure variables that were statistically significant individually. Second, for pairs of moderately correlated variables ($r > 0.3$), we added both exposure variables simultaneously to the base model to assess changes in the odds ratio estimates compared to modeling each exposure separately.

We conducted two sensitivity analyses to evaluate the impact of our choice of disease definition. First, because our allergic-asthma group contained two different conditions (asthma and allergy), we evaluated whether the results for pesticides might reflect an association with allergy alone. We removed the controls with allergy from the comparison group and constructed

a polytomous model with four groups: controls without allergy, allergy alone, asthma with allergy, and asthma alone. Second, because asthma can co-occur with other respiratory diseases, we repeated our analyses after excluding individuals with other respiratory diseases (i.e., chronic bronchitis, farmers lung)

Analyses were done using SAS v9.1 (SAS Institute Inc., Cary, NC) and AHS dataset release P1REL0310.

Results

Of the 19,704 male farmers included in this analysis, 441 individuals (2.2%) reported adult-onset asthma, with 127 classified as allergic and 314 classified as non-allergic. Men with asthma tended to be older and heavier and slightly more likely to have smoked compared to those without asthma (Table 1). Wheeze was reported by 17% of controls and 74% of those with asthma. Allergic symptoms were most common in those with allergic asthma. While use of any PPE for pesticide application was similar for those with and without asthma, individuals with asthma were much more likely to report the use of respiratory protection than those without asthma.

While overall pesticide use history (e.g., total years applied pesticides) was not related to either type of asthma, a history of HPEE was associated with an almost doubling of asthma risk for both types of asthma ($OR_{\text{allergic}}=1.98$, 95% CI=1.30, 2.99; $OR_{\text{non-allergic}}=1.96$, 95% CI=1.49, 2.56) (Table 2). After adjusting for HPEE and other base model covariates, a history of doctor-diagnosed pesticide poisoning was non-significantly associated with an almost two-fold increased prevalence of allergic asthma ($OR=1.95$, 95%CI=0.86, 4.39. Due to the strong association of HPEE with asthma, all subsequent analyses controlled for HPEE.

We evaluated 48 pesticides for asthma risk among farmers (ziram and trichlorfon had too few exposed cases to be assessed). While a minority of the cases were allergic asthma, more individual pesticides were associated with allergic asthma than non-allergic asthma based on ever use (12 vs. 4, Table 3). For allergic asthma, three herbicides (2,4,5-TP, EPTC, and paraquat), six insecticides (organochlorines: chlordane, heptachlor, and lindane; and organophosphates: diazinon, parathion, and coumaphos), one fungicide (captan), and two fumigants (ethylene dibromide and 80/20 mix – carbon tetrachloride and carbon disulfide) were positively associated. For non-allergic asthma, one herbicide (petroleum oil) and three insecticides (organochlorine: DDT and organophosphates; phorate and malathion) were associated. No chemical was significantly associated with both asthma subgroups, though the odds ratios were almost identical in both groups for DDT and phorate. Four pesticides had odds ratios that were significantly different (coumaphos, paraquat, captan, and lindane) between allergic and non-allergic asthma.

We evaluated exposure-response for pesticides and adult asthma using three cumulative pesticide exposure metrics: total years of use, lifetime days of use, and intensity-adjusted lifetime days of use. The results for all metrics were similar and because the intensity-adjusted metric accounts for potential differences in exposure as a result of application practices, we present those results in Figure 1 for those pesticides with significant trend tests for either allergic or non-allergic asthma (the online supplement Table E1 includes the exposure-response models for all 48 pesticides evaluated). Ten of the 12 pesticides associated with allergic-asthma also had significant exposure-response trends, and, with the exception of chlordane and parathion, all had higher odds ratios at the higher exposure levels. The herbicides 2,4-D and 2,4,5-T had significant exposure-response trends for allergic asthma, though the models for ever use were not

statistically significant. For non-allergic asthma, DDT and malathion had significant dose-response trends.

We were also able to assess the association for two fungicides (captan and metalaxyl) related to use of treated seed. Captan-treated seed was associated with allergic asthma only (OR=2.49, 95% CI=1.42, 4.36). Similarly, users of metalaxyl-treated seed were over 5 times more likely to report allergic asthma (OR=5.18, 95% CI =2.48, 10.8).

We saw some evidence of correlations among specific pesticides and attenuation of some associations, but no strong evidence of confounding for any specific pesticide. The highest observed correlation among the controls was 0.37 for 2,4,5-T and 2,4,5-TP. Below, we present the results for the odds ratios that changed in the multiple chemical models. For allergic asthma, parathion attenuated the association of paraquat (OR from 1.67 to 1.36, 95% CI=0.82, 2.25). For the organochlorine pesticides, there was evidence of confounding for the associations with allergic asthma. Chlordane attenuated the associations with DDT and allergic asthma (OR from 1.42 to 1.17, 95% CI=0.75,1.83) and diazinon (OR from 1.57 to 1.39, 95% CI=0.91,2.12) while heptachlor attenuated the association of chlordane (OR from 1.77 to 1.53, 95%CI=1.02,2.31). In a model containing DDT, heptachlor, and chlordane, the association for DDT was reduced to 1.05 (95%CI=0.67, 1.67), while the ORs for heptachlor and chlordane remained elevated (1.93 and 1.52, respectively). When heptachlor and chlordane were included in a model with diazinon, the OR_{allergic asthma} for diazinon was reduced to 1.25 (95% CI=0.82,1.93) with little change in the estimates for heptachlor and chlordane (1.93 and 1.46, respectively). We saw similar attenuation of the exposure-response models for DDT when chlordane and heptachlor were included in the model. None of the significant associations for non-allergic asthma could be explained by correlations with other pesticides.

We saw little evidence of association between prevalent asthma and current farming activities (data not shown). Allergic asthma was associated with using gasoline as a solvent to clean (OR= 1.48, 95%CI=1.04, 2.13) and with performing veterinary services (OR=1.51, 95%CI=1.03, 2.21). Veterinary services (OR=0.77, 95%CI =0.60,0.97) and driving combines (OR=0.72, 95%CI=0.53,0.97) were inversely associated with non-allergic asthma. There were no positive associations between non-allergic asthma and current farming activities. Current farm activities did not appear to confound the pesticide results.

To determine whether the differential results for allergic and non-allergic asthma were due to allergy alone, we removed allergic individuals from our control group and reran the analysis (Online supplement Table E2). While some pesticides were associated with allergy alone, the odds ratios were stronger for allergic asthma than for allergy alone for all pesticides associated with allergic asthma in earlier models.

To evaluate whether our results were related to asthma, or to some co-morbid respiratory disease, we excluded all individuals with chronic bronchitis and farmers lung and reran our models (online supplement Table E3). A larger proportion of asthma cases, both allergic and non-allergic, reported some other respiratory illness compared to controls (35% vs. 5%). With these individuals excluded, the odds ratios for allergic asthma were statistically significant for five pesticides (2,4,5-T, parathion, coumaphos, captan, and 80/20 mix) and similar to the values reported in Table 3. For four other pesticides, the resulting odds ratios were attenuated: heptachlor (OR from 2.01 to 1.30), chlordane (OR from 1.77 to 1.27), paraquat (OR=1.67 to 1.36), and 2,4,5-TP (OR from 1.91 to 1.68). Of the four pesticides significantly associated with non-allergic asthma, only phorate remained statistically significant when the individuals with other lung diseases were removed. After removing the other respiratory diseases, the odds ratio

for fonofos and non-allergic asthma increased from 1.22 to 1.39 (95% CI=1.00, 1.94) and the association between petroleum oil and non-allergic asthma went away (OR from 1.35 to 1.15).

Discussion

Grains, hays, and animals have been identified as important etiologic agents for respiratory disease among farmers for centuries (14); however, few studies have evaluated the association between specific pesticides and respiratory disease. Here we offer additional evidence that pesticides may also be a risk factor for asthma among farmers. Building on previous work in farmers (1), farmworkers (6, 15, 16), pesticide applicators (3), and farm women (2), we observed that specific pesticides were associated with allergic asthma and that self-reported high pesticide exposure events were associated with both allergic and non-allergic asthma. We also saw associations with lifetime days of use of individual pesticides suggesting an exposure-response relationship.

High pesticide exposure events are infrequent events on farms, however 14% of farmers report having at least one HPEE in their lifetime. These events often do not result in seeking care for pesticide poisoning (17). HPEE was an important risk factor for both allergic and non-allergic asthma. While we saw no evidence of an interaction between HPEE and individual pesticides, a history of HPEE was an important confounder of the pesticide findings. Before adjusting for HPEE, 14 pesticides were associated with allergic asthma and 10 with non-allergic asthma. HPEE has been associated with a twofold increased prevalence of the respiratory diseases chronic bronchitis (18) and farmers lung (19) as well as with a number of neurological outcomes including depression (20) and neurological symptoms (21).

Our findings are consistent with results from other respiratory analyses from the AHS and other studies. The herbicides, EPTC and paraquat, were associated with wheeze among

farmers (4). Paraquat was also associated with allergic asthma among farm women (2) as well as respiratory symptoms and oxygen desaturation in studies of farmworkers in Costa Rica and South Africa (22, 23). Paraquat has also been associated with allergic symptoms in grape farmers in Crete (24). The organochlorine insecticides DDT and lindane were associated with allergic asthma in farm women; DDT was also associated with non-allergic asthma among farm women. The organophosphate insecticides coumaphos and parathion were associated with allergic asthma in farm women; coumaphos was associated with wheeze in the commercial pesticide applicators (5), while parathion was associated with wheeze among farmers (4). Other investigators have also found organophosphate insecticides to be associated with asthma or wheeze. Saskatchewan farmers who used organophosphate insecticides had increased asthma, though the association was stronger with users of carbamate insecticides (1); our findings suggest stronger evidence for organophosphates. Farmworkers with depressed acetyl cholinesterase (a measure of organophosphate exposure) have increased respiratory symptoms (6, 16).

For the pesticides associated with non-allergic asthma, all were previously associated with wheeze or asthma in the AHS. Petroleum oil was associated with non-allergic asthma in farm women (2) and with wheeze among farmers and commercial pesticide applicators (4, 5). Both phorate and malathion were associated with allergic, but not non-allergic, asthma in farm women (2) and were associated with wheeze. One pesticide that was not associated with asthma in this analysis was carbofuran; this pesticide was strongly associated with asthma among Saskatchewan farmers (1) and among AHS farm women (2).

Allergic asthma is far less common in adults than children (25) and far less common in farmers than in other occupational groups (10). Yet in this study the associations with pesticides

were much stronger for allergic than non-allergic asthma; our analysis of farm women (2) showed a similar relationship. Pesticides may modulate inflammatory responses to farm bioaerosols such as endotoxin and allergens. Allergen-sensitized guinea pigs had a lower threshold for parathion-induced airway responsiveness (9). Carbaryl enhanced the allergenicity of dust mites in animals (26). Captan and metalaxyl, both commonly used seed treatments, were strongly associated with allergic asthma when pre-applied to seed, suggesting a possible interaction of allergen and pesticide. Given the variety of pesticides associated with allergic asthma, we cannot rule out the possibility that some common ingredient in the pesticidal products may explain these results.

We relied on self-reported exposure and outcome data. Farmers provide reliable information regarding their personal pesticide use (27, 28). By using an intensity-adjusted exposure metric, we were able to account for potential differences in exposure resulting from different application practices. The intensity metric is able to distinguish between high and low exposure intensity in field studies (29), thus we are better able to classify individuals with respect to total exposure. Even though individuals with allergic asthma appeared to have reduced their exposure through fewer pesticide application days annually and increased use of respiratory protection, we were still able to observe significant exposure-response trends using this intensity metric. We have no reason to believe that pesticide use reporting would differ by asthma or allergy status. Self-reported doctor-diagnosed asthma has been shown to be a reliable and valid endpoint (30, 31). Among farmers, self-reported asthmatic subjects have a higher symptom prevalence and lower lung function than non-asthmatics (1). In our sample, individuals with asthma were more likely to report wheeze than those without asthma and individuals classified as allergic had more allergic symptoms than those classified as non-allergic. Allergy was defined

as a history of doctor-diagnosed eczema or hayfever. While we have no clinical measures to assess allergic status such as skin prick testing or IgE measurement, our measure is similar to ones used previously. Our two case groups are based on responses to three questions (asthma, eczema, and hayfever), similar the classification that Upton and colleagues used to define atopic and non-atopic asthma among adults in Scotland (25). We classified 71% of adult asthma among farmers as non-allergic, an estimate very similar to the 70% reported among Norwegian adults where atopy was defined by IgE (10). By limiting our definition of allergy to those with doctor-diagnosis of hayfever or eczema it is likely that these individuals received allergy testing at the time of diagnosis, however we have no data to evaluate that.

Farms represent a complex occupational setting with opportunities for a number of concurrent exposures, including multiple pesticides. When we took into account correlated pesticides, the associations for DDT were attenuated, but the associations for other organochlorines remained elevated. While we saw no strong associations with current farm activities, we lack data on lifetime farming exposures to animals, hays, and grains, which may contribute to asthma. Hence, the findings for some specific pesticides may be confounded by previous exposures. For example, coumaphos is an insecticide used on animals, and thus, it may be a surrogate for historic animal exposures. However, a majority of the pesticides associated with asthma have not been used in animal production. Given the lack of commonality to the pesticides, both in their use patterns and toxicity, it seems unlikely that all the findings are due to uncontrolled confounding.

In this analysis of 19,704 male farmers with good information on lifetime pesticide use, specific pesticides were associated with allergic asthma. One of the pesticides, parathion, has also been associated with allergic asthma in farm women, wheeze in farmers, and with airway

hyperresponsiveness in allergen-sensitized animals. Additionally individuals with a history of a high pesticide exposure event were twice as likely to report having adult-onset asthma. Because the analysis was cross-sectional we cannot evaluate the temporal order of exposure and disease, however, given the consistency of the results for epidemiologic and animal studies, more detailed studies are warranted to identify patterns of pesticide use which may contribute to asthma among farmers.

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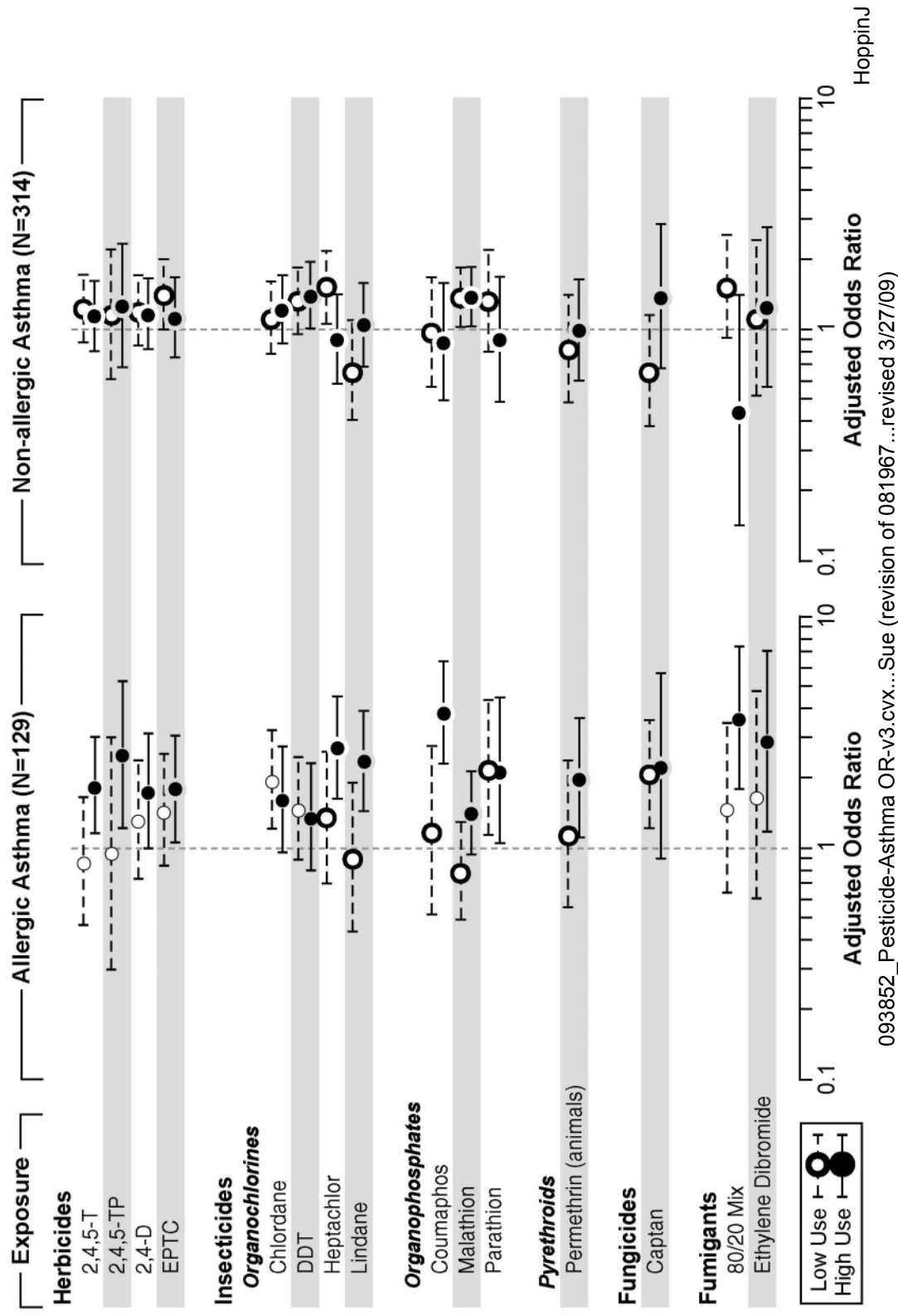
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Tables and Figures

Table 1	Demographic, Medical and Farming Characteristics by adult-onset asthma status for 19,704 male farmers from the Agricultural Health Study
Table 2	Associations of Overall Pesticide Exposure variables with Allergic and Non-allergic Asthma among 19,704 male farmers from the Agricultural Health Study
Table 3	Associations of ever-use of individual pesticides with allergic and non-allergic asthma among 19,704 male farmers in the Agricultural Health Study

Figure 1 Selected Pesticide Exposure-Response Models for Adult Asthma



Legend Pesticides selected for presentation had a significant p-value for trend for either allergic or non-allergic asthma. For allergic asthma, 2,4,5-T; 2,4,5-TP; 2,4-D; EPTC; chlordane; lindane; heptachlor; parathion; permethrin on animals; captan; 80/20 mix; and ethylene dibromide had significant trend tests ($p < 0.05$). For non-allergic asthma, DDT and malathion had significant trend tests ($P < 0.05$).

Cutpoints for pesticide use were based on the users of that specific pesticide and were then split at the median number of intensity-adjusted lifetime days for that pesticide.

Online Supplement

Table E1	Complete exposure-response models for all 48 pesticides evaluated.
Table E2	Sensitivity analysis for allergy alone
Table E3	Odds ratios for individual pesticide and allergic and non-allergic asthma after exclusion of other respiratory disease cases

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Table 1: Demographic, Medical and Farming Characteristics by adult-onset asthma status
for 19,704 male farmers from the Agricultural Health Study

Characteristic	Control (N=19263)		Allergic Asthma (N=127)		Non-allergic Asthma (N=314)	
	n	%	n	%	n	%
Age (years)						
20-39	5143	27	19	15	31	10
40-49	5003	26	28	22	68	22
50-59	4520	24	33	26	83	26
60+	4597	24	47	37	132	42
State						
Iowa	13046	68	90	71	226	72
North Carolina	6217	32	37	29	88	28
Body Mass Index (kg/m ²)						
<25	5023	26	25	20	68	22
25-29	9874	51	68	54	157	50
30+	4366	23	34	27	89	28
Smoking History						
Never	10408	54	66	52	145	46
Past	6359	33	54	43	143	46
Current	2496	13	7	6	26	8
Race						
White	18907	98	125	98	310	99
Other Race	301	2	2	2	2	1
Education						
High School Graduate or less	10811	57	64	53	198	64
Greater than High School	8055	43	58	48	111	36
Grew up on a farm	17728	92	118	93	287	92
Respiratory and Allergic Conditions						
Farmer's Lung	386	2	22	17	32	10
Chronic Bronchitis	603	3	33	26	86	28
Allergy*	1856	10	127	100		0
Eczema	406	2	36	28		0
Hayfever	1519	8	105	83		0
Wheeze	3152	17	105	84	216	70
Runny Nose in the Past Year	12413	66	110	88	224	74
Watery Itchy Eyes in the Past	6343	34	85	69	119	40
PPE Use, 10 years before enrollment**						
Never used PPE	4071	21	24	19	62	20
Cartridge respirator	920	5	11	9	20	6
Dust mask	2117	11	20	16	56	18
Chemical gloves	6971	36	49	39	119	38
PPE Use at enrollment						
Never use PPE	1768	9	15	12	31	10
Cartridge respirator	1827	9	14	11	25	8
Dust mask	3937	20	36	28	95	30
Chemical gloves	13361	69	92	72	208	66

*History of doctor-diagnosed eczema or hayfever

**PPE= Personal Protective Equipment; PPE use is not mutually exclusive. Individuals may report more than one method.

Table 2: Associations of Overall Pesticide Exposure variables with Allergic and Non-allergic Asthma among 19,704 male farmers from the Agricultural Health Study

Exposure	Controls (N=19,263)	Allergic asthma (N=127)				Non-allergic asthma (N=314)				
		n	%	Odds	95% Confidence Interval	n	%	Odds	95% Confidence Interval	
Days per year apply pesticides	<5	3464	19	27	21	1.00	52	18	1.00	
	5-9	4649	25	29	23	0.78	69	23	0.95	0.66
	10-19	5683	31	41	32	0.94	97	33	1.18	0.84
	20-39	3303	18	22	17	0.92	55	19	1.29	0.87
	40+	1366	7	8	6	0.84	23	8	1.40	0.84
Total Years applied pesticides	<5	2206	12	11	9	1.00		7	1.00	
	6-10	2640	14	7	6	0.51		11	1.29	0.75
	11-20	6102	33	36	28	1.07		29	1.23	0.75
	21-30	4710	25	48	38	1.53		28	1.11	0.68
	31+	2908	16	25	20	0.99		25	1.18	0.71
HPEE**		2744	14	30	24	1.98	73	23	1.96	2.56
HPEE w/o respiratory exp		1745	9	16	13	1.70	39	12	1.72	2.44
HPEE w/ respiratory exposure		955	5	14	11	2.53	34	11	2.41	3.49
Decade of first HPEE	1940s-1960s	363	2	6	5	2.11		5	2.09	1.24
	1970s	700	4	9	7	2.26		5	1.58	0.94
	1980s-1990s	1419	7	14	11	2.10		10	1.95	1.32
Pesticide Poisoning		381	2	7	6	1.95	11	4	1.10	0.58
										2.07

*Odds ratios adjusted for age, state, smoking, HPEE, and BMI

**HPEE=High Pesticide Exposure Event

Table 3: Associations of ever-use of individual pesticides with allergic and non-allergic asthma among 19,704 male farmers in the Agricultural Health Study

Exposure	Controls (N=19,263)		Allergic asthma (N=127)				Non-allergic asthma (N=314)				p-for difference **	
	n	%	n	%	Odds Ratio*	95% Confidence Interval	n	%	Odds Ratio*	95% Confidence Interval		
Herbicides	3564	19	38	31	1.44	0.96	2.14	88	29	1.20	0.93	1.56
	925	5	13	11	1.91	1.06	3.44	23	8	1.24	0.80	1.91
	15054	79	110	87	1.56	0.91	2.69	264	85	1.19	0.86	1.64
	10219	56	69	57	0.93	0.64	1.34	180	62	1.15	0.90	1.47
	14034	73	94	75	0.95	0.63	1.45	241	78	1.12	0.85	1.49
	5058	27	42	33	1.23	0.84	1.81	96	31	1.10	0.85	1.42
	6124	32	45	36	1.21	0.83	1.75	99	32	1.05	0.82	1.35
	7842	43	58	49	1.14	0.77	1.70	149	51	1.24	0.96	1.60
	9607	53	71	59	1.19	0.78	1.81	173	61	1.28	0.97	1.69
	3611	20	35	29	1.61	1.06	2.43	71	25	1.25	0.94	1.66
	14788	77	104	82	1.37	0.86	2.17	247	79	1.15	0.87	1.51
	8042	45	52	44	0.97	0.64	1.48	123	43	0.88	0.67	1.14
	8624	47	58	48	0.99	0.69	1.44	148	51	1.12	0.88	1.43
	7179	38	55	44	1.16	0.79	1.70	127	42	1.01	0.79	1.29
	3068	16	28	22	1.67	1.05	2.65	40	13	0.82	0.58	1.18
Insecticides	7104	38	44	35	0.94	0.64	1.36	120	39	1.14	0.90	1.44
	3933	21	33	27	1.28	0.85	1.92	84	28	1.35	1.04	1.74
	9964	55	62	53	0.79	0.54	1.16	164	55	0.89	0.70	1.14
Carbamates												
Aldicarb	1508	8	7	6	0.79	0.35	1.79	14	5	0.66	0.37	1.17
Carbaryl	8089	43	63	50	1.26	0.85	1.85	140	45	0.98	0.77	1.26
Carbofuran	5266	29	42	35	1.10	0.75	1.61	106	37	1.17	0.92	1.49

Organochlorines												
Aldrin	3247	17	33	27	1.19	0.77	1.86	79	26	0.95	0.72	1.26
Chlordane	3592	19	42	34	1.77	1.19	2.63	86	28	1.22	0.94	1.59
DDT	4344	23	47	37	1.42	0.93	2.17	124	40	1.41	1.09	1.84
Dieldrin	704	4	10	8	1.47	0.75	2.90	13	4	0.61	0.35	1.09
Heptachlor	2289	12	34	27	2.01	1.30	3.11	64	21	1.20	0.89	1.61
Lindane	2528	14	28	23	1.57	1.01	2.41	44	14	0.85	0.61	1.17
Toxaphene	2156	11	18	15	1.06	0.63	1.78	43	14	0.96	0.69	1.35
Organophosphates												
Chlorpyrifos	8037	42	60	48	1.26	0.89	1.80	135	43	1.08	0.86	1.36
Coumaphos	1646	9	24	21	2.34	1.49	3.70	26	9	0.88	0.58	1.32
Diazinon	3891	21	38	30	1.57	1.05	2.35	70	23	1.03	0.78	1.36
Dichlorvos	2092	12	21	18	1.47	0.90	2.39	39	14	1.05	0.74	1.49
Fonofos	4079	23	37	31	1.43	0.95	2.16	84	29	1.22	0.93	1.60
Malathion	12150	64	87	69	1.08	0.74	1.59	229	74	1.35	1.04	1.75
Parathion	1501	8	19	16	2.05	1.21	3.46	30	10	1.11	0.75	1.66
Phorate	5776	31	48	39	1.23	0.84	1.81	126	41	1.29	1.01	1.65
Terbufos	7224	40	51	43	1.05	0.72	1.53	132	45	1.16	0.91	1.48
Pyrethroids												
Permethrin (animals)	2365	13	21	18	1.51	0.92	2.45	33	11	0.91	0.63	1.32
Permethrin (crops)	2328	13	20	18	1.52	0.93	2.48	42	15	1.28	0.91	1.79
Fungicides												
Benomyl	1519	8	10	8	0.97	0.49	1.94	22	7	0.87	0.54	1.38
Captan	1951	11	22	19	1.83	1.15	2.94	29	10	0.88	0.60	1.30
Chlorothalonil	1427	7	6	5	0.64	0.27	1.51	22	7	1.10	0.69	1.76
Maneb/Mancozeb	1530	8	13	11	1.40	0.74	2.67	20	7	0.80	0.49	1.31
Metalaxyl	3695	20	26	21	1.26	0.77	2.06	50	16	0.89	0.63	1.25
Fumigants												
80/20 mix***	818	4	15	12	2.15	1.23	3.76	24	8	1.25	0.81	1.93
Aluminum phosphide	641	3	6	5	1.34	0.58	3.06	12	4	1.14	0.63	2.04
Ethylene dibromide	818	4	10	8	2.07	1.02	4.20	16	5	1.25	0.73	2.15
Methyl bromide	2838	15	16	13	0.86	0.46	1.60	41	13	0.98	0.66	1.45

*Odds ratios adjusted for age, state, smoking, HPEE, and BMI

**P for difference shown if p-value <0.1

***80/20 mix is carbon tetrachloride/carbon disulfide.

*Odds ratios adjusted for age, state, smoking, HPEE, and BMI

**P for difference shown if p-value <0.1

***80/20 mix is carbon tetrachloride/carbon disulfide.

