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## **Supplemental Material**

# **Pesticide Exposure and Risk of Rheumatoid Arthritis among Licensed Male Pesticide Applicators in the Agricultural Health Study**

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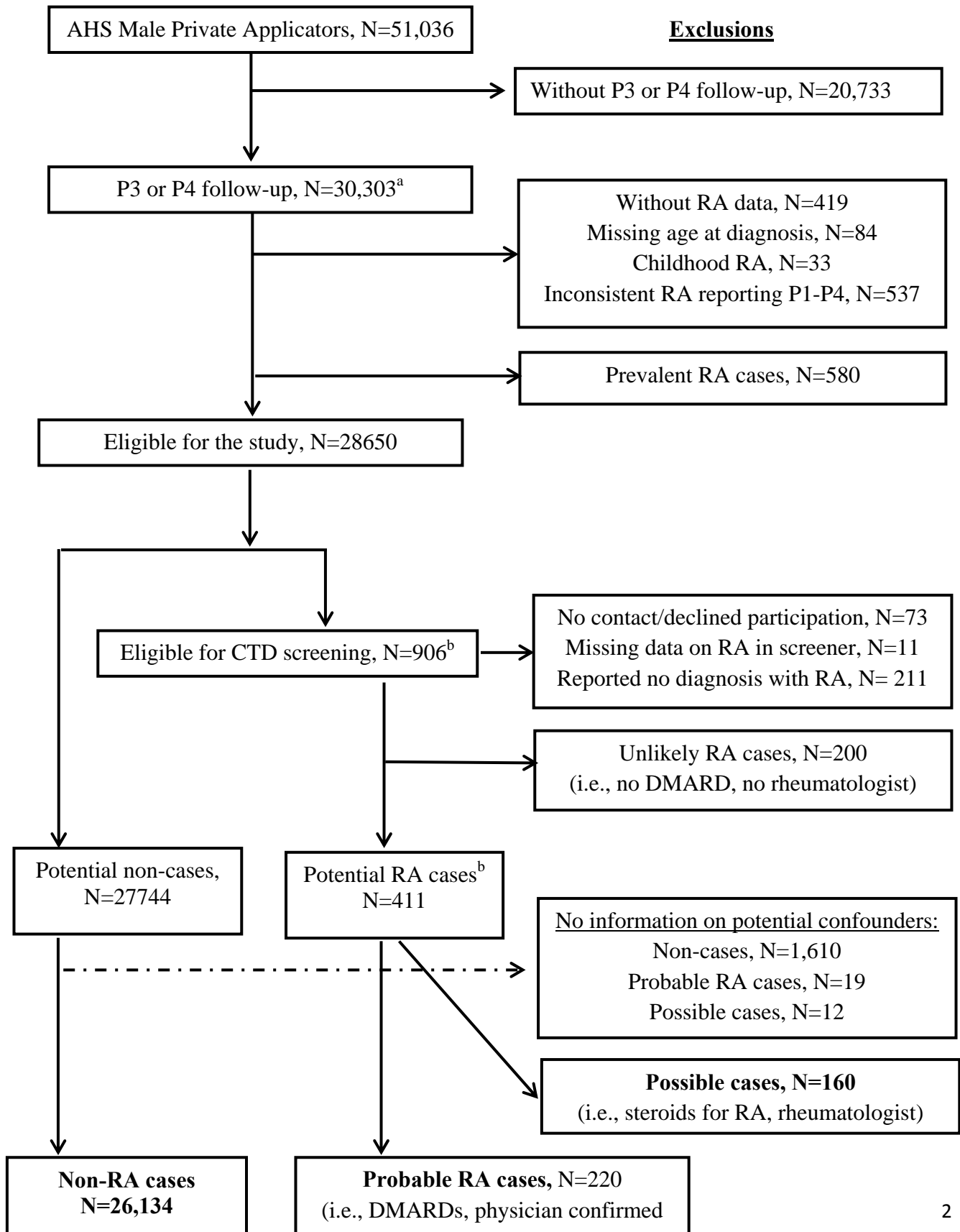
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**Figure S1 – Identification of cases and non-cases**



<sup>a</sup>Includes 25 self-reported cases of systemic autoimmune diseases (19 with RA) who were identified at enrollment, but lacked P3/P4 data. These were included because they were asked about RA in pilot validation study

<sup>b</sup>Includes 99 who were eligible to be screened for RA because they reported another systemic autoimmune disease (e.g., lupus); of these, 3 were identified as potential RA cases for the first time during screening

**Table S1 – Exposure-response relationship between intensity-weighted lifetime use of specific pesticides and incident RA among male licensed pesticide applicators in the Agricultural Health Study**

	Non-cases (N=26,134)	Incident RA cases (N=220)	
	N (%)	N (%)	OR <sup>a</sup> (95%CI)
<b>INSECTICIDES</b>			
<b>ORGANOCHLORINES</b>			
<b>Aldrin</b>			
Never	11257 (83)	99 (78)	Reference
T1	1095 (8)	14 (11)	1.38 (0.77,2.50)
T2	116 (1)	2 (2)	1.73 (0.41,7.23)
T3	1084 (8)	12 (10)	1.18 (0.63,2.22)
p-trend			0.45
<b>Chlordane</b>			
Never	11035 (81)	93 (72)	Reference
T1	639 (5)	8 (6)	1.14 (0.54,2.38)
T2	1051 (8)	15 (12)	1.41 (0.81,2.48)
T3	825 (6)	13 (10)	1.48 (0.81,2.69)
p-trend			0.11
<b>Toxaphene<sup>b</sup></b>			
Never	12143 (89)	104 (83)	Reference
T1	859 (6)	7 (6)	0.82 (0.38,1.78)
T2	392 (3)	9 (7)	2.00 (0.99,4.04)
T3	202 (1)	6 (5)	2.42 (1.03,5.68)
p-trend			0.02
<b>ORGANOPHOSPHATES</b>			
<b>Chlorpyrifos</b>			
Never	14678 (57)	111 (52)	Reference
T1	3756 (15)	34 (16)	1.27 (0.86,1.88)
T2	3727 (15)	37 (17)	1.42 (0.97,2.06)
T3	3602 (14)	32 (15)	1.25 (0.84,1.86)
p-trend			0.09
<b>Dichlorvos</b>			
Never	21220 (88)	170 (87)	Reference
T1	1052 (4)	9 (5)	1.33 (0.67,2.63)
T2	936 (4)	7 (4)	1.14 (0.53,2.44)
T3	796 (3)	9 (5)	1.76 (0.87,3.48)
p-trend			0.11
<b>Fonofos</b>			
Never	18544 (77)	141 (71)	Reference

<b>T1</b>	1825 (8)	24 (12)	2.27 (1.44,3.57)
<b>T2</b>	2074 (9)	12 (6)	0.98 (0.54,1.80)
<b>T3</b>	1725 (7)	22 (11)	2.10 (1.32,3.36)
<b>p-trend</b>			0.005
<b>Terbufos</b>			
<b>Never</b>	14178 (59)	116 (58)	Reference
<b>T1</b>	3548 (15)	27 (14)	1.14 (0.74,1.75)
<b>T2</b>	3464 (14)	29 (15)	1.25 (0.82,1.89)
<b>T3</b>	2954 (12)	28 (14)	1.33 (0.88,2.03)
<b>p-trend</b>			0.12
<b>CARBAMATES</b>			
<b>Carbaryl<sup>b</sup></b>			
<b>Never</b>	7966 (59)	55 (44)	Reference
<b>T1</b>	2339 (17)	23 (18)	1.33 (0.81,2.18)
<b>T2</b>	1775 (13)	25 (20)	1.62 (0.96,2.71)
<b>T3</b>	1434 (11)	23 (18)	1.61 (0.90,2.86)
<b>p-trend</b>			0.05
<b>HERBICIDES</b>			
<b>Alachlor</b>			
<b>Never</b>	10698 (45)	80 (40)	Reference
<b>T1</b>	5931 (25)	50 (25)	1.25 (0.87,1.79)
<b>T2</b>	3001 (13)	24 (12)	1.17 (0.74,1.86)
<b>T3</b>	4362 (18)	44 (22)	1.37 (0.94,1.98)
<b>p-trend</b>			0.11
<b>Atrazine</b>			
<b>Never</b>	6685 (26)	52 (24)	Reference
<b>T1</b>	8199 (32)	60 (28)	1.10 (0.75,1.62)
<b>T2</b>	6002 (23)	49 (23)	1.28 (0.85,1.91)
<b>T3</b>	4914 (19)	54 (25)	1.62 (1.09,2.40)
<b>p-trend</b>			0.01
<b>Chlorimuron ethyl</b>			
<b>Never</b>	9217 (68)	81 (63)	Reference
<b>T1</b>	1116 (8)	12 (9)	1.47 (0.79,2.72)
<b>T2</b>	1881 (14)	20 (16)	1.34 (0.82,2.21)
<b>T3</b>	1397 (10)	16 (12)	1.42 (0.83,2.45)
<b>p-trend</b>			0.10
<b>Imazethapyr</b>			
<b>Never</b>	12842 (54)	111 (57)	Reference
<b>T1</b>	4910 (21)	50 (26)	1.74 (1.18,2.55)
<b>T2</b>	599 (3)	5 (3)	1.42 (0.56,3.59)
<b>T3</b>	5582 (23)	30 (15)	0.97 (0.62,1.52)
<b>p-trend</b>			0.82
<b>Trifluralin</b>			

<b>Never</b>	10781 (45)	89 (45)	Reference
<b>T1</b>	4520 (19)	35 (18)	1.11 (0.75,1.66)
<b>T2</b>	3572 (15)	31 (16)	1.34 (0.87,2.06)
<b>T3</b>	5204 (22)	43 (22)	1.25 (0.85,1.84)
<b>p-trend</b>			0.18
<b>FUNGICIDES</b>			
<b>Chlorothalonil</b>			
<b>Never</b>	24052 (93)	194 (91)	Reference
<b>T1</b>	862 (3)	7 (3)	0.82 (0.38,1.77)
<b>T2</b>	572 (2)	6 (3)	1.02 (0.44,2.35)
<b>T3</b>	304 (1)	7 (3)	2.35 (1.07,5.14)
<b>p-trend</b>			0.18
<b>Metalaxyl</b>			
<b>Never</b>	11164 (83)	96 (74)	Reference
<b>T1</b>	714 (5)	7 (5)	1.05 (0.48,2.28)
<b>T2</b>	991 (7)	14 (11)	1.20 (0.64,2.23)
<b>T3</b>	616 (5)	13 (10)	1.76 (0.91,3.41)
<b>p-trend</b>			0.13
<b>OTHERS</b>			
<b>Methyl bromide</b>			
<b>Never</b>	22116 (86)	166 (76)	Reference
<b>T1</b>	1004 (4)	16 (7)	1.74 (1.00,3.02)
<b>T2</b>	1469 (6)	17 (8)	1.18 (0.68,2.05)
<b>T3</b>	1203 (5)	19 (9)	1.54 (0.90,2.63)
<b>p-trend</b>			0.11

<sup>a</sup>Adjusted for age, state of enrollment, pack-years smoking, and education

<sup>b</sup>From take-home questionnaire

<sup>c</sup>Tertiles (T1, T2, T3) based on intensity-weighted lifetime days of use.

**Table S2 - Ever use of specific pesticides and incident RA among smokers and non-smokers male licensed pesticide applicators in the Agricultural Health Study**

	Non,smokers			Smokers		
	Non-cases	Incident cases	OR <sup>a</sup> (95%CI)	Non-cases	Incident cases	OR <sup>a</sup> (95%CI)
	N=15046	N=100		N=11088	N=120	
	N (%)	N (%)		N (%)	N (%)	
<b>INSECTICIDES</b>						
<b>ORGANOCHLORINES</b>						
	7364 (50)	61 (62)	1.37 (0.88,2.15)	6533 (62)	72 (63)	0.99 (0.65,1.49)
Aldrin	1230 (15)	13 (25)	1.80 (0.96,3.37)	1140 (20)	16 (21)	1.25 (0.69,2.29)
Chlordane	1305 (16)	17 (31)	2.37 (1.33,4.23)	1285 (23)	19 (25)	1.05 (0.61,1.80)
DDT	1387 (17)	18 (33)	2.41 (1.36,4.25)	1533 (27)	22 (29)	0.92 (0.53,1.59)
Dieldrin	258 (3)	7 (13)	4.48 (2.01,10.01)	233 (4)	1 (1)	0.35 (0.05,2.56)
Heptachlor	943 (12)	8 (15)	1.34 (0.63,2.85)	780 (14)	7 (9)	0.74 (0.32,1.68)
Lindane	1176 (15)	10 (19)	1.32 (0.66,2.63)	830 (15)	8 (11)	0.75 (0.36,1.59)
Toxaphene	765 (10)	10 (19)	2.21 (1.11,4.42)	722 (13)	13 (18)	1.26 (0.68,2.35)
<b>ORGANOPHOSPHATES</b>						
	13571 (90)	96 (96)	NC	9927 (90)	107 (89)	1.20 (0.66,2.16)
Chlorpyrifos	6577 (44)	53 (54)	1.47 (0.99,2.18)	4684 (43)	51 (44)	1.13 (0.78,1.64)
Coumaphos	1368 (10)	6 (6)	0.63 (0.27,1.44)	891 (9)	7 (7)	0.81 (0.38,1.76)
Diazinon	1421 (18)	12 (23)	1.39 (0.73,2.66)	1370 (25)	22 (29)	1.13 (0.67,1.90)
Dichlorvos	1784 (13)	16 (17)	1.40 (0.82,2.40)	1071 (11)	10 (10)	1.22 (0.62,2.38)
Fonofos	3447 (24)	29 (30)	1.34 (0.86,2.07)	2292 (23)	29 (28)	1.94 (1.21,3.13)
Malathion	5181 (64)	33 (61)	0.88 (0.51,1.52)	3802 (68)	54 (71)	1.27 (0.77,2.12)
Parathion	510 (6)	4 (7)	1.18 (0.42,3.27)	512 (9)	7 (9)	0.79 (0.36,1.77)
Phorate	2626 (33)	20 (37)	1.22 (0.70,2.12)	1719 (31)	20 (27)	1.03 (0.59,1.79)
Terbufos	6230 (44)	43 (44)	1.01 (0.68,1.51)	3939 (39)	43 (41)	1.37 (0.91,2.06)
<b>CARBAMATES</b>						
	9485 (64)	72 (72)	1.25 (0.79,1.99)	7817 (73)	91 (78)	1.16 (0.73,1.84)
Aldicarb	454 (6)	3 (6)	0.98 (0.31,3.16)	496 (9)	10 (14)	1.19 (0.57,2.45)
Carbaryl	3011 (37)	25 (47)	1.50 (0.87,2.57)	2697 (48)	49 (64)	1.74 (1.03,2.92)
Carbofuran	4007 (28)	24 (25)	0.85 (0.54,1.36)	3059 (31)	39 (38)	1.40 (0.94,2.09)
<b>PYRETHROIDS</b>						
	3877 (28)	24 (26)	0.98 (0.61,1.57)	2360 (24)	28 (29)	1.46 (0.93,2.28)
Permethrin C <sup>b</sup>	1966 (14)	12 (13)	0.91 (0.50,1.67)	1386 (14)	19 (19)	1.40 (0.84,2.33)
Permethrin P <sup>b</sup>	2395 (17)	15 (15)	0.90 (0.52,1.57)	1242 (12)	10 (10)	1.07 (0.55,2.09)

<b>HERBICIDES</b>						
<b>2,4-D</b>	11894 (80)	85 (86)	1.55 (0.88,2.73)	8542 (78)	84 (73)	0.93 (0.60,1.44)
<b>2,4,5-T</b>	1418 (18)	17 (31)	2.10 (1.18,3.72)	1194 (21)	12 (16)	0.74 (0.39,1.41)
<b>2,4,5-TP</b>	360 (4)	5 (9)	2.13 (0.84,5.37)	315 (6)	2 (3)	0.47 (0.12,1.95)
<b>Alachlor</b>	7989 (56)	60 (64)	1.38 (0.91,2.11)	5706 (56)	62 (57)	1.14 (0.77,1.68)
<b>Atrazine</b>	11346 (76)	81 (83)	1.52 (0.90,2.57)	7974 (72)	85 (71)	1.09 (0.72,1.64)
<b>Butylate</b>	2372 (29)	16 (29)	0.99 (0.55,1.77)	1540 (28)	19 (25)	1.04 (0.61,1.78)
<b>Chlorimuron_ethyl</b>	2645 (33)	24 (44)	1.59 (0.93,2.72)	1825 (33)	27 (35)	1.23 (0.76,1.98)
<b>Cyanazine</b>	6677 (47)	39 (41)	0.80 (0.53,1.21)	4283 (42)	37 (35)	1.02 (0.65,1.60)
<b>Dicamba</b>	8297 (58)	51 (55)	0.87 (0.57,1.30)	5105 (51)	41 (38)	0.80 (0.50,1.27)
<b>EPTC</b>	3210 (23)	12 (13)	0.51 (0.28,0.93)	1971 (20)	12 (12)	0.68 (0.36,1.27)
<b>Glyphosate</b>	11353 (76)	75 (75)	0.96 (0.61,1.51)	8707 (79)	91 (77)	0.88 (0.57,1.37)
<b>Imazethapyr</b>	7034 (50)	47 (51)	1.04 (0.69,1.56)	4258 (43)	39 (38)	1.27 (0.77,2.09)
<b>Metolachlor</b>	7179 (50)	47 (50)	0.99 (0.66,1.48)	4730 (47)	38 (36)	0.74 (0.49,1.12)
<b>Metribuzin</b>	3378 (42)	24 (44)	1.08 (0.63,1.84)	2128 (38)	24 (32)	0.98 (0.58,1.67)
<b>Paraquat</b>	1116 (14)	9 (17)	1.25 (0.61,2.56)	1015 (18)	15 (19)	0.85 (0.47,1.56)
<b>Pendimethalin</b>	2995 (37)	20 (36)	0.97 (0.56,1.69)	2188 (39)	29 (38)	0.93 (0.58,1.49)
<b>Trifluralin</b>	8097 (57)	57 (59)	1.09 (0.73,1.63)	5493 (54)	55 (53)	1.25 (0.83,1.90)
<b>FUNGICIDES</b>						
<b>Benomyl</b>	484 (6)	1 (2)	0.29 (0.04,2.13)	495 (9)	8 (10)	0.89 (0.41,1.92)
<b>Captan</b>	1752 (12)	13 (14)	1.12 (0.62,2.01)	1181 (12)	8 (8)	0.68 (0.33,1.42)
<b>Chlorothalonil</b>	895 (6)	11 (11)	1.96 (1.05,3.69)	966 (9)	13 (11)	1.04 (0.56,1.90)
<b>Maneb</b>	491 (6)	3 (5)	0.89 (0.28,2.85)	540 (10)	11 (15)	1.19 (0.59,2.39)
<b>Metalaxyl</b>	1184 (15)	9 (16)	1.14 (0.55,2.33)	1327 (24)	26 (34)	1.33 (0.76,2.32)
<b>OTHERS</b>						
<b>80/20_mix</b>	300 (4)	3 (6)	1.52 (0.47,4.89)	295 (5)	2 (3)	0.51 (0.12,2.12)
<b>Methylbromide</b>	1642 (11)	15 (15)	1.45 (0.83,2.51)	2183 (20)	38 (32)	1.57 (0.98,2.53)
<b>Petroleum_oil</b>	1801 (22)	9 (16)	0.68 (0.33,1.39)	1227 (22)	10 (13)	0.61 (0.31,1.19)
<b>No. of pesticides applied</b>						
<b>0 – 5</b>	3042 (20)	16 (16)	Reference	2699 (24)	28 (23)	Reference
<b>6 – 9</b>	4425 (30)	27 (27)	1.30 (0.69,2.44)	3095 (28)	41 (34)	1.51 (0.92,2.49)
<b>10 – 13</b>	3740 (25)	26 (26)	1.53 (0.80,2.92)	2534 (23)	21 (18)	1.04 (0.58,1.87)
<b>14+</b>	3768 (25)	30 (30)	1.67 (0.89,3.16)	2726 (25)	30 (25)	1.44 (0.83,2.47)



NC = not calculated due to fewer than 5 exposed or unexposed cases

<sup>a</sup>Adjusted for age, state of enrollment, pack-years smoking, and education

<sup>b</sup> Permethrin C: used on crops; Permethrin P: used on poultry and livestock

**Table S3 – Ever use of specific pesticides and incident RA among male licensed pesticide applicators in the Agricultural Health Study, according to age**

	Age < 50			Age >= 50		
	Non-cases N=15814	Incident RA N=111	OR <sup>a</sup> (95%CI)	Non-cases N=10320	Incident RA N=109	OR <sup>a</sup> (95%CI)
	N (%)	N (%)		N (%)	N (%)	
<b>INSECTICIDES</b>						
<b>ORGANOCHLORINES</b>	6340 (41)	52 (48)	1.03 (0.69,1.54)	7557 (77)	81 (77)	1.13 (0.71,1.80)
<b>Aldrin</b>	490 (6.3)	4 (7.0)	0.81 (0.29,2.30)	1880 (32.4)	25 (35.2)	1.38 (0.82,2.35)
<b>Chlordane</b>	947 (12.1)	15 (26.3)	1.82 (0.97,3.39)	1643 (28.4)	21 (29.2)	1.01 (0.60,1.71)
<b>DDT</b>	455 (5.8)	8 (13.8)	1.15 (0.51,2.59)	2465 (42.3)	32 (44.4)	1.15 (0.71,1.87)
<b>Dieldrin</b>	61 (0.8)	0 (0.0)	NC	430 (7.5)	8 (11.3)	1.91 (0.89,4.11)
<b>Heptachlor</b>	443 (5.7)	1 (1.8)	0.28 (0.04,2.01)	1280 (22.0)	14 (19.7)	0.99 (0.53,1.85)
<b>Lindane</b>	1006 (12.9)	7 (11.9)	0.91 (0.41,2.03)	1000 (17.3)	11 (15.5)	0.95 (0.49,1.83)
<b>Toxaphene</b>	558 (7.1)	8 (14.0)	1.35 (0.62,2.95)	929 (16.0)	15 (21.4)	1.40 (0.78,2.51)
<b>ORGANOPHOSPHATES</b>	14323 (91)	104 (94)	1.67 (0.77,3.63)	9175 (89)	99 (91)	1.42 (0.73,2.77)
<b>Chlorpyrifos</b>	7148 (45.4)	54 (50.0)	1.21 (0.83,1.77)	4113 (40.3)	50 (46.7)	1.34 (0.91,1.97)
<b>Coumaphos</b>	1329 (9.0)	5 (5.0)	0.52 (0.21,1.28)	930 (10.3)	8 (8.6)	0.87 (0.42,1.81)
<b>Diazinon</b>	1421 (18.2)	18 (31.0)	1.42 (0.78,2.61)	1370 (23.7)	16 (22.9)	0.91 (0.51,1.62)
<b>Dichlorvos</b>	1731 (11.6)	10 (9.7)	0.94 (0.48,1.83)	1124 (12.3)	16 (17.2)	1.82 (1.04,3.20)
<b>Fonofos</b>	3469 (23.0)	29 (28.2)	1.56 (0.99,2.47)	2270 (24.7)	29 (30.2)	1.76 (1.09,2.87)
<b>Malathion</b>	5015 (64.0)	33 (55.9)	0.69 (0.41,1.16)	3968 (67.9)	54 (76.1)	1.55 (0.89,2.69)
<b>Parathion</b>	471 (6.1)	3 (5.1)	NC	551 (9.6)	8 (11.3)	1.06 (0.50,2.28)
<b>Phorate</b>	2251 (28.7)	15 (25.9)	0.99 (0.53,1.84)	2094 (36.1)	25 (35.2)	1.15 (0.67,1.96)
<b>Terbufos</b>	6477 (42.9)	42 (40.2)	1.03 (0.67,1.54)	3692 (40.0)	44 (44.9)	1.48 (0.97,2.26)
<b>CARBAMATES</b>	9919 (64)	81 (74)	1.22 (0.77,1.93)	7383 (74)	82 (77)	1.09 (0.68,1.74)
<b>Aldicarb</b>	593 (7.6)	7 (12.1)	0.94 (0.40,2.24)	357 (6.2)	6 (8.6)	1.12 (0.46,2.77)
<b>Carbaryl</b>	2892 (36.9)	32 (55.2)	1.44 (0.81,2.59)	2816 (48.2)	42 (59.2)	1.50 (0.89,2.52)
<b>Carbofuran</b>	3869 (25.8)	30 (29.4)	1.05 (0.68,1.62)	3197 (34.9)	33 (34.4)	1.03 (0.67,1.58)
<b>PYRETHROIDS</b>	12505 (79)	80 (72)	0.76 (0.49,1.17)	8518 (83)	95 (87)	1.74 (0.97,3.13)

<b>Permethrin C<sup>b</sup></b>	2332 (15.7)	18 (17.7)	1.09 (0.65,1.82)	1020 (11.3)	13 (14.1)	1.24 (0.69,2.25)
<b>Permethrin P<sup>b</sup></b>	2710 (18.0)	13 (12.5)	0.78 (0.43,1.42)	927 (10.1)	12 (12.8)	1.51 (0.81,2.83)
<b>HERBICIDES</b>	15497 (98)	107 (96)	0.52 (0.19,1.42)	10090 (98)	106 (97)	0.89 (0.28,2.87)
<b>2,4-D</b>	12219 (77.7)	78 (72.9)	0.86 (0.55,1.35)	8217 (80.6)	91 (85.1)	1.66 (0.95,2.89)
<b>2,4,5-T</b>	947 (12.1)	9 (15.3)	1.16 (0.56,2.39)	1665 (28.5)	20 (28.6)	1.08 (0.63,1.83)
<b>Alachlor</b>	8236 (54.7)	58 (55.2)	1.01 (0.68,1.49)	5459 (58.4)	64 (66.0)	1.51 (0.99,2.33)
<b>Atrazine</b>	11707 (74.3)	86 (77.5)	1.35 (0.85,2.15)	7613 (74.2)	80 (74.8)	1.12 (0.71,1.78)
<b>Butylate</b>	2254 (28.8)	11 (18.3)	0.57 (0.29,1.11)	1658 (28.4)	24 (33.3)	1.41 (0.85,2.35)
<b>Chlorimuron ethyl</b>	2846 (36.4)	20 (33.3)	0.93 (0.54,1.61)	1624 (27.7)	31 (43.1)	2.04 (1.27,3.27)
<b>Cyanazine</b>	6810 (45.2)	31 (30.1)	0.57 (0.37,0.93)	4150 (44.8)	45 (46.9)	1.49 (0.94,2.37)
<b>Dicamba</b>	8510 (56.7)	42 (41.2)	0.62 (0.39,0.98)	4885 (52.9)	50 (51.0)	1.26 (0.78,2.02)
<b>EPTC</b>	3445 (23.1)	10 (9.6)	0.41 (0.21,0.79)	1736 (19.1)	14 (15.4)	0.93 (0.51,1.67)
<b>Glyphosate</b>	12344 (78.3)	85 (77.3)	0.82 (0.52,1.31)	7716 (75.3)	81 (75.0)	0.96 (0.61,1.50)
<b>Imazethapyr</b>	7593 (50.6)	43 (42.2)	0.99 (0.62,1.58)	3699 (40.5)	43 (45.3)	1.85 (1.11,3.05)
<b>Metholachlor</b>	7685 (50.9)	42 (40.4)	0.73 (0.49,1.09)	4224 (45.6)	43 (45.3)	1.01 (0.72,1.66)
<b>Metribuzin</b>	3219 (41.1)	15 (25.4)	0.53 (0.29,1.00)	2287 (39.2)	33 (45.8)	1.65 (0.98,2.76)
<b>Paraquat</b>	1201 (15.3)	11 (18.3)	0.73 (0.36,1.50)	930 (15.9)	13 (18.3)	1.02 (0.53,1.95)
<b>Pendimethalin</b>	3172 (40.5)	21 (35.0)	0.78 (0.46,1.33)	2011 (34.3)	28 (38.9)	1.19 (0.74,1.93)
<b>Trifluralin</b>	8434 (55.8)	54 (52.4)	1.04 (0.69,1.58)	5156 (55.7)	58 (59.2)	1.43 (0.93,2.21)
<b>FUNGICIDES</b>	5541 (35)	48 (43)	1.04 (0.67,1.60)	3771 (37)	43 (39)	1.00 (0.65,1.54)
<b>Benomyl</b>	508 (6.5)	3 (5.1)	0.40 (0.12,1.31)	471 (8.1)	6 (8.3)	0.85 (0.35,2.06)
<b>Captan</b>	1891 (12.7)	12 (11.5)	0.90 (0.50,1.65)	1042 (11.4)	9 (9.8)	0.90 (0.45,1.80)
<b>Chlorothalonil</b>	1132 (7.2)	13 (11.8)	1.25 (0.67,2.33)	729 (7.1)	11 (10.2)	1.30 (0.67,2.53)
<b>Maneb</b>	495 (6.3)	6 (10.3)	0.96 (0.39,2.39)	536 (9.2)	8 (11.1)	0.97 (0.43,2.17)
<b>Metalaxyl</b>	1411 (18.0)	16 (27.1)	1.00 (0.51,1.96)	1100 (18.9)	19 (26.4)	1.35 (0.74,2.44)
<b>OTHERS</b>						
<b>80/20 Mix</b>	182 (2.3)	0 (0.0)	...	413 (7.1)	5 (6.9)	1.06 (0.42,2.67)
<b>Methylbromide</b>	2166 (13.8)	26 (23.4)	1.21 (0.70,2.09)	1659 (16.2)	27 (25.0)	1.61 (0.94,2.75)
<b>Petroleum oil</b>	1840 (23.6)	11 (18.3)	0.77 (0.40,1.50)	1188 (20.5)	8 (11.1)	0.51 (0.24,1.07)
<b>No. of pesticides applied</b>						
<b>0 – 5</b>	3386 (22)	25 (23)	Reference	2355 (23)	19 (18)	Reference
<b>6 – 9</b>	4745 (30)	42 (38)	1.37 (0.82,2.28)	2775 (27)	26 (24)	1.36 (0.74,2.50)

<b>10 – 13</b>	3994 (25)	21 (19)	0.86 (0.47,1.57)	2280 (22)	26 (24)	1.79 (0.96,3.33)
<b>14+</b>	3617 (23)	23 (21)	0.99 (0.55,1.78)	2877 (28)	37 (34)	2.16 (1.19,3.90)

NC = not calculated due to fewer than 5 exposed or unexposed cases

<sup>a</sup>Adjusted for age, state of enrollment, pack-years smoking, and education

<sup>b</sup>Permethrin C: used on crops; Permethrin P: used on poultry and livestock

**Table S4 – Main characteristic of probable and possible RA cases and non-cases among male licensed private applicators in the Agricultural Health Study**

	<b>Non-cases (N=26,134) N (%)</b>	<b>Probable + Possible Incident RA cases (N=380) N (%)</b>	<b>OR<sup>a</sup> (95%CI)</b>
<b>Age at enrollment (years)</b>			
<40	7858 (30)	54 (14)	Reference
40-49	7956 (30)	118 (31)	2.16 (1.56,2.98)
50-59	6035 (23)	122 (32)	2.94 (2.13,4.06)
60+	4285 (16)	86 (23)	2.92 (2.07,4.11)
<b>Body Mass Index (Kg/m<sup>2</sup>)</b>			
< 25	6157 (26)	62 (17)	Reference
25 to <30	12353 (51)	193 (54)	1.51 (1.13,2.01)
≥ 30	5551 (23)	101 (28)	1.79 (1.30,2.47)
<b>State of enrollment</b>			
Iowa	17972 (69)	209 (55)	Reference
North Carolina	8162 (31)	171 (45)	1.66 (1.35,2.04)
<b>Race</b>			
White	25454 (97)	359 (95)	Reference
Non-white	659 (3)	20 (5)	2.01 (1.27,3.17)
<b>Education</b>			
High school or less	14117 (54)	250 (66)	Reference
> high school	12017 (46)	130 (34)	0.70 (0.56,0.87)
<b>Smoking status at enrollment</b>			
Never Smoked	15046 (58)	171 (45)	Reference
Past Smoker	7637 (29)	138 (36)	1.33 (1.06,1.68)
Current Smoker	3392 (13)	70 (19)	1.83 (1.38,2.42)
<b>Pack-years of cigarette smoked</b>			
None	15046 (58)	172 (45)	Reference
<5	3460 (13)	44 (12)	1.07 (0.76,1.49)
5-18	3760 (15)	69 (18)	1.46 (1.10,1.93)
> 18	3868 (15)	95 (25)	1.80 (1.39,2.33)
<b>Alcohol consumption (times/week)</b>			
None	8143 (32)	143 (39)	Reference
≤ 1	11778 (47)	148 (40)	0.84 (0.67,1.07)
> 1	5331 (21)	77 (21)	0.97 (0.73,1.29)

<sup>a</sup>Adjusted for continuous age

**Table S5 – Ever use of specific pesticides associated with possible and probable incident RA among male licensed pesticide applicators in the Agricultural Health Study**

	Non-cases (N=26,134) N (%)	Probable + Possible Incident RA cases (N=380) N (%)	OR <sup>a</sup> (95%CI)
<b>INSECTICIDES</b>			
<b>ORGANOCHLORINES</b>	13897 (55)	241 (66)	1.25 (0.98,1.58)
Aldrin	2370 (17)	47 (22)	1.23 (0.86,1.75)
Chlordane	2590 (19)	54 (26)	1.18 (0.85,1.64)
DDT	2920 (21)	69 (32)	1.23 (0.88,1.70)
Dieldrin	491 (4)	10 (5)	1.17 (0.61,2.27)
Heptachlor	1723 (13)	27 (13)	0.97 (0.63,1.49)
Lindane	2006 (15)	29 (14)	0.95 (0.64,1.42)
Toxaphene	1487 (11)	29 (14)	1.03 (0.69,1.55)
<b>ORGANOPHOSPHATES</b>	23498 (90)	344 (91)	1.32 (0.93,1.88)
Chlorpyrifos	11261 (43)	157 (42)	1.03 (0.84,1.27)
Coumaphos	2259 (10)	23 (7)	0.71 (0.46,1.08)
Diazinon	2791 (21)	52 (25)	1.06 (0.76,1.48)
Dichlorvos	2855 (12)	38 (11)	1.14 (0.80,1.61)
Fonofos	5739 (24)	90 (26)	1.44 (1.11,1.87)
Malathion	8983 (66)	140 (65)	0.99 (0.74,1.32)
Parathion	1022 (8)	20 (9)	0.96 (0.59,1.44)
Phorate	4345 (32)	73 (35)	1.34 (0.99,1.83)
Terbufos	10169 (42)	140 (40)	1.13 (0.91,1.42)
<b>CARBAMATES</b>	17302 (68)	254 (73)	1.07 (0.83,1.38)
Aldicarb	950 (7)	24 (11)	1.26 (0.79,2.01)
Carbaryl	5708 (42)	107 (50)	1.09 (0.80,1.47)
Carbofuran	7066 (29)	117 (34)	1.17 (0.93,1.46)
<b>PYRETHROIDS</b>	6237 (26)	61 (25)	1.07 (0.89,1.44)
Permethrin C <sup>b</sup>	3352 (14)	50 (15)	1.11 (0.82,1.51)
Permethrin P <sup>b</sup>	3637 (15)	36 (11)	0.89 (0.62,1.27)
<b>HERBICIDES</b>			
2,4,D	20436 (79)	290 (78)	1.09 (0.84,1.41)
2,4,5,T	2612 (19)	43 (20)	0.96 (0.68,1.37)
2,4,5,TP	675 (5)	18 (9)	1.65 (1.01,2.70)
Alachlor	13695 (56)	205 (58)	1.14 (0.92,1.42)
Atrazine	19320 (74)	284 (75)	1.25 (0.98,1.60)
Butylate	3912 (29)	55 (26)	0.99 (0.72,1.36)
Chlorimuron ethyl	4470 (33)	81 (38)	1.39 (1.05,1.84)
Cyanazine	10960 (45)	135 (39)	1.00 (0.78,1.27)
Dicamba	13402 (55)	155 (45)	0.87 (0.67,1.11)
EPTC	5181 (22)	50 (15)	0.77 (0.57,1.06)
Glyphosate	20060 (77)	285 (76)	0.90 (0.70,1.14)
Imazethapyr	11292 (45)	141 (41)	1.20 (0.92,1.55)
Metolachlor	11909 (49)	158 (46)	1.04 (0.83,1.29)
Metribuzin	5506 (40)	81 (38)	1.15 (0.85,1.56)
Paraquat	2131 (16)	42 (20)	1.06 (0.73,1.53)

<b>Pendimethalin</b>	5183 (38)	86 (40)	1.10 (0.83,1.45)
<b>Trifluralin</b>	13590 (56)	182 (52)	1.06 (0.85,1.33)
<b>FUNGICIDES</b>			
<b>Benomyl</b>	979 (7)	16 (8)	0.75 (0.44,1.28)
<b>Captan</b>	2933 (12)	34 (10)	0.86 (0.60,1.22)
<b>Chlorothalonil</b>	1861 (7)	46 (12)	1.58 (1.05,2.06)
<b>Maneb</b>	1031 (8)	18 (9)	0.75 (0.45,1.27)
<b>Metalaxyl</b>	2511 (18)	56 (26)	1.17 (0.82,1.66)
<b>OTHERS</b>			
<b>80/20 Mix</b>	595 (4)	8 (4)	0.76 (0.37,1.55)
<b>Methylbromide</b>	3825 (15)	85 (23)	1.19 (0.89,1.59)
<b>Petroleum oil</b>	3028 (22)	34 (16)	0.73 (0.51,1.06)
<b>No. of pesticides applied</b>			
<b>0 – 5</b>	5741 (22)	86 (23)	Reference
<b>6 – 9</b>	7520 (29)	106 (28)	1.15 (0.86,1.54)
<b>10 – 13</b>	6274 (24)	85 (23)	1.19 (0.87,1.64)
<b>14+</b>	6494 (25)	100 (27)	1.32 (0.97,1.80)

<sup>a</sup> Adjusted for age, state of enrollment, pack-years smoking, and education

<sup>b</sup> Permethrin C: used on crops; Permethrin P: used on poultry and livestock

**Table S6 – Lifetime days of pesticide use in relation to probable + possible incident RA among male licensed pesticide applicators in the Agricultural Health Study**

	<b>Non-cases (N=26,134) N (%)</b>	<b>Probable + Possible Incident RA Cases (N=380) N (%)</b>	<b>OR<sup>a</sup> (95%CI)</b>
<b>INSECTICIDES</b>			
<b>ORGANOCHLORINES</b>			
<b>Aldrin</b>			
Never	11257 (83)	164 (78)	Reference
< 20	1095 (8)	21 (10)	1.21 (0.75,1.96)
>=20 to <24.5	116 (1)	13 (6)	1.18 (0.65,2.12)
>=24.5	1084 (8)	11 (5)	1.28 (0.68,2.41)
<b>p-trend</b>			0.32
<b>Chlordane</b>			
Never	11035 (81)	158 (75)	Reference
< 8.75	639 (5)	12 (6)	1.03 (0.56,1.88)
>=8.75 to <20	1051 (8)	19 (9)	1.08 (0.66,1.75)
>=20	825 (6)	22 (10)	1.48 (0.93,2.34)
<b>p-trend</b>			0.15
<b>Toxaphene<sup>b</sup></b>			
Never	12143 (89)	180 (87)	Reference
< 8.75	859 (6)	4 (2)	0.90 (0.33,2.46)
>=8.75 to <38.75	392 (3)	10 (5)	0.66 (0.34,1.25)
>=38.75	202 (1)	14 (7)	1.87 (1.06,3.30)
<b>p-trend</b>			0.47
<b>ORGANOPHOSPHATES</b>			
<b>Chlorpyrifos</b>			
Never	14678 (57)	216 (58)	Reference
< 14.5	3756 (15)	47 (13)	0.93 (0.68,1.28)
>=14.5 to <56	3727 (15)	57 (15)	1.14 (0.85,1.53)
>=56	3602 (14)	51 (14)	1.05 (0.77,1.43)
<b>p-trend</b>			0.54
<b>Dichlorvos</b>			
Never	21220 (88)	304 (90)	Reference
< 20	1052 (4)	9 (3)	0.92 (0.47,1.80)
>=20 to <116	936 (4)	11 (3)	0.96 (0.52,1.77)
>=116	796 (3)	15 (4)	1.33 (0.79,2.26)
<b>p-trend</b>			0.46
<b>Fonofos</b>			
Never	18544 (77)	254 (74)	Reference
< 20	1825 (8)	31 (9)	1.62 (1.10,2.39)
>=20 to <56	2074 (9)	25 (7)	1.12 (0.73,1.71)
>=56	1725 (7)	34 (10)	1.76 (1.21,2.56)
<b>p-trend</b>			0.005
<b>Terbufos</b>			
Never	14178 (59)	208 (60)	Reference
< 24.5	3548 (15)	48 (14)	1.14 (0.83,1.58)
>=24.5 to <63.75	3464 (14)	44 (13)	1.07 (0.76,1.49)
>=63.75	2954 (12)	46 (13)	1.24 (0.89,1.72)
<b>p-trend</b>			0.22
<b>CARBAMATES</b>			
<b>Carbaryl<sup>b</sup></b>			
Never	7966 (59)	106 (50)	Reference
< 14.5	2339 (17)	31 (15)	0.93 (0.62,1.39)
>=14.5 to <56	1775 (13)	30 (14)	1.25 (0.81,1.94)
>=56	1434 (11)	43 (20)	1.18 (0.77,1.79)
<b>p-trend</b>			0.36



<b>HERBICIDES</b>			
<b>2,4,5-T<sup>b</sup></b>			
Never	11038 (81)	169 (81)	Reference
< 8.75	474 (3)	4 (2)	0.48 (0.18,1.32)
>=8.75 to <24.5	1555 (11)	18 (9)	0.93 (0.56,1.52)
>=24.5	499 (4)	18 (9)	1.18 (0.72,1.95)
<b>p-trend</b>			0.84
<b>Alachlor</b>			
Never	10698 (45)	147 (43)	Reference
< 24.5	5931 (25)	50 (14)	0.92 (0.67,1.28)
>=24.5 to <108.5	3001 (13)	67 (19)	1.15 (0.85,1.54)
>=108.5	4362 (18)	81 (23)	1.36 (1.03,1.79)
<b>p-trend</b>			0.03
<b>Atrazine</b>			
Never	6685 (26)	93 (25)	Reference
< 29.5	8199 (32)	85 (23)	1.16 (0.85,1.57)
>=29.5 to <116	6002 (23)	92 (25)	1.13 (0.84,1.52)
>=116	4914 (19)	102 (27)	1.51 (1.13,2.03)
<b>p-trend</b>			0.01
<b>Imazethapyr</b>			
Never	12842 (54)	203 (59)	Reference
< 14.5	4910 (21)	68 (20)	1.29 (0.95,1.76)
>=14.5 to <29.5	599 (3)	42 (12)	1.17 (0.81,1.69)
>=29.5	5582 (23)	29 (8)	1.09 (0.72,1.65)
<b>p-trend</b>			0.49
<b>Trifluralin</b>			
Never	10781 (45)	167 (49)	Reference
< 29.5	4520 (19)	62 (18)	1.05 (0.78,1.42)
>=29.5 to <108.5	3572 (15)	44 (13)	1.01 (0.71,1.43)
>=108.5	5204 (22)	71 (21)	1.11 (0.82,1.48)
<b>p-trend</b>			0.56
<b>FUNGICIDES</b>			
<b>Chlorothalonil</b>			
Never	24052 (93)	327 (89)	Reference
< 12.5	862 (3)	11 (3)	1.09 (0.59,2.01)
>=12.5 to <87.5	572 (2)	14 (4)	1.50 (0.86,2.63)
>=87.5	304 (1)	16 (4)	1.71 (1.01,2.89)
<b>p-trend</b>			0.02
<b>Metalaxyl</b>			
Never	11164 (83)	160 (74)	Reference
< 12.25	714 (5)	12 (6)	0.95 (0.52,1.72)
>=12.25 to <28	991 (7)	12 (6)	1.07 (0.58,2.00)
>=28	616 (5)	31 (14)	1.70 (1.07,2.69)
<b>p-trend</b>			0.05
<b>OTHERS</b>			
<b>Methyl bromide</b>			
Never	22116 (86)	292 (78)	Reference
< 15.5	1004 (4)	29 (8)	1.27 (0.84,1.92)
>=15.5 to <54.25	1469 (6)	20 (5)	0.91 (0.56,1.48)
>=54.25	1203 (5)	35 (9)	1.49 (1.00,2.21)
<b>p-trend</b>			0.12

<sup>a</sup>Adjusted for age, state of enrollment, pack-years smoking, and education

<sup>b</sup>From take-home questionnaire