Tables and Figures

Table 1: Summary of study population characteristics ($N = 38\,549$; 1.51 million person-years). The cohort was restricted to individuals who were hired in or after 1938 and for whom at least half of their work history data was available. Individuals were considered lost to follow-up once they reach the maximum observed age at death.

	n	%	
Study population size (N)	38 549	100%	
Race			
White	22816	59%	
African American	7131	18%	
Unknown	8602	22%	
Sex			
Male	33792	88%	
Female	4757	12%	
$\mathrm{Plant}^{\natural}$			
Plant 1	9 090	24%	
Plant 2	17087	44%	
Plant 3	12372	32%	
Ever exposed to MWFs			
Straight	20352	53%	
Soluble	31795	82%	
Synthetic	12523	32%	
Deceased by end of follow-up	20565	53%	
	Median	25 th %tile	75^{th} %tile
Years of follow-up	39	34	47
Years at work*	16.62	7.52	27.33
Year of hire	1965	1952	1973
Age at hire (years)	28	23	36
Year of birth	1937	1922	1948
Year of death among deceased	1996	1984	2006
Age at death (years) among deceased	70	60	79
Cumulative exposure [#] to MWFs (mg/m ³ ·y)			
Straight	0.66	0.21	2.34
Soluble	4.41	1.74	10.71
Synthetic	0.44	0.15	1.56

 $^{^{\}natural}$ For individuals who worked at several plants, plant was taken to be the site where they accrued the most work record time.

^{*} Among those with known date of worker exit.

 $[\]sharp$ Summary statistics calculated for ever-exposed individuals at end of follow-up only. Exposures were lagged 21 years.

Table 2: Standardized mortality ratios calculated for the GM-UAW cohort followed from 1941 to 2015. NIOSH LTAS-extracted reference rates were used for the years 1940 through 2009. CDC Mortality data were used as reference rates for 2010 onwards.

Cause of death		SMR	(95% CI) [♭]
All causes	20 565	0.91	(0.89, 0.92)
All natural causes	18857	0.89	(0.88, 0.91)
All cancers	5472	0.96	(0.94, 0.99)
Esophageal cancer	184	1.06	(0.92, 1.23)
Stomach cancer	192	1.10	(0.95, 1.27)
Intestinal cancer	418	0.90	(0.82, 0.99)
Rectal cancer	89	0.86	(0.70, 1.06)
Bladder and urinary organ cancers	146	0.95	(0.81, 1.12)
Bile duct, liver, and gallbladder cancers	162	0.88	(0.76, 1.03)
Pancreatic cancer	315	1.05	(0.94, 1.17)
Laryngeal cancer	74	1.17	(0.93, 1.47)
Lung cancer	1891	1.07	(1.02, 1.12)
Skin cancer	73	0.66	(0.52, 0.83)
Prostate cancer	417	0.82	(0.75, 0.91)
Brain and nervous system cancers	128	0.99	(0.84, 1.18)
Leukemia	200	0.98	(0.85, 1.12)
Breast cancer	76	0.79	(0.63, 0.99)
All nonmalignant respiratory diseases	1682	0.84	(0.81, 0.89)
Chronic obstructive pulmonary disease		0.93	(0.87, 0.99)
Cirrhosis and other chronic liver disease		0.90	(0.81, 1.00)
All heart diseases	6743	0.75	(0.73, 0.77)
Ischemic heart disease		0.89	(0.87, 0.92)
Cerebrovascular disease	1080	0.83	(0.78, 0.88)
All external causes	1671	1.03	(0.98, 1.08)

 $^{^{\}flat}$ Variance estimates assume Poisson-distributed rates in the observed population. Reference rates were assumed to be parameters.

Figure 1: Adjusted hazard ratio estimates for cancers and cumulative exposure to straight metalworking fluids.

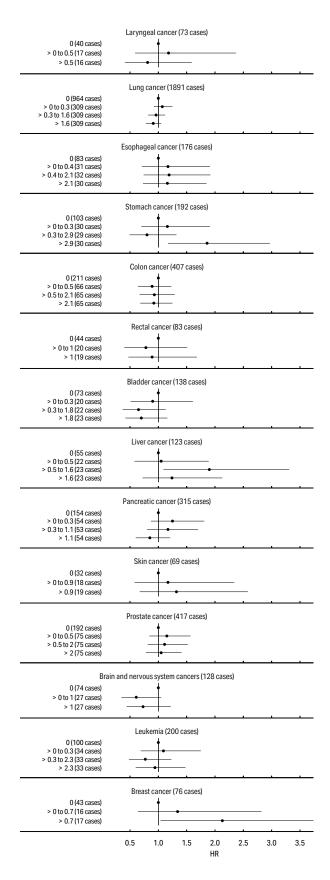


Figure 2: Adjusted hazard ratio estimates for cancers and cumulative exposure to soluble metalworking fluids.

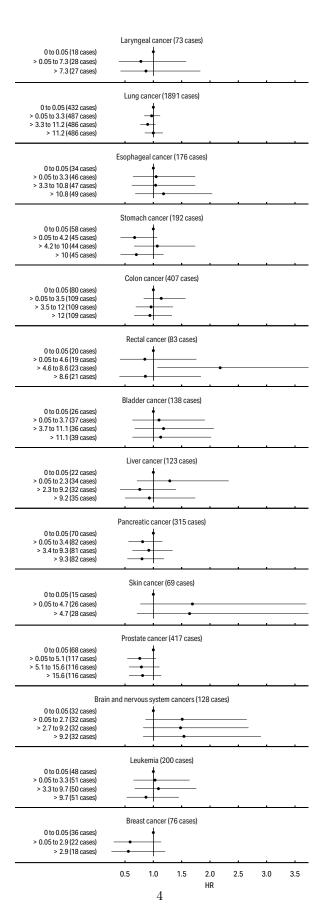
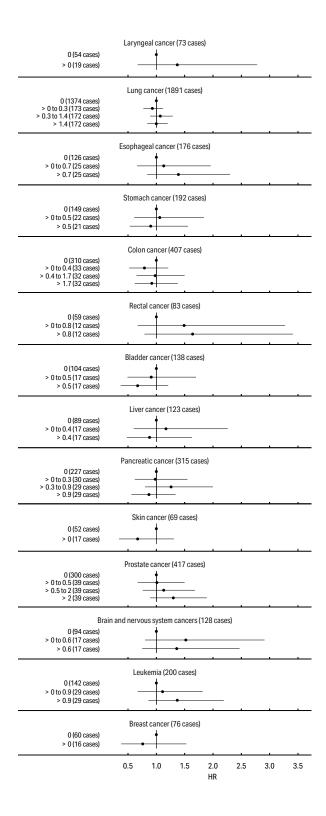


Figure 3: Adjusted hazard ratio estimates for cancers and cumulative exposure to synthetic metalworking fluids.



Appendix 1

Table 3: International Classification of Diseases (ICD) codes used to define cases. Deaths prior to 1999 were defined with respect to the $9^{\rm th}$ revision. Deaths in or after 1999 were defined with respect to the $10^{\rm th}$.

Cause of death	ICD 9	ICD 10
All causes	_	_
All natural causes	001-799	A00-R99, U00-Z99
All cancers	140-239	C00-C99, D00-D49
Esophageal cancer	150	C15
Stomach cancer	151	C16
Intestinal cancer	152-153	C17-C18
Rectal cancer	154	C19–C21
Kidney cancer	189.0, 189.2	C64-C66
Bladder and urinary organ cancers	188,189.3–189.9	C67–C68
Bile duct, liver, and gallbladder cancers	155-156	C22-C24
Pancreatic cancer	157	C25
Laryngeal cancer	161	C32
Lung cancer	162	C33–C34
Skin cancer	172–173	C43-C44, C46.0, C46.9
Prostate cancer	185	C61
Brain and nervous system cancers	191 – 192	C47, C70–C72
Leukemia	204-208	C91.0-C91.3, C91.5-C91.9, C92-C95
Breast cancer	174 – 175	C50
All nonmalignant respiratory diseases	460–466, 470–478, 480–487, 490–495, 496–519	A48.1, J00–J01, J02.8–J02.9, J03.8–J03.9, J04–J06, J10–J18, J20–J22, J40–J46, J30–J33, J34.1–J34.8, J35–J39, J47, J60:J95, J98, R09.1
Chronic obstructive pulmonary disease	490–492, 496	J40-J44
Pneumonia	480-486	A48.1, J12–J18
Cirrhosis and other chronic liver disease	571	K70, K73–K74, K76.0
All heart diseases	390–398, 402, 404, 410–414, 420–429	I00–I09, I11–I13, I20–I22, I24–I25, I30–I38, I40, I42, I44–I52, I97.0–I97.1, I97.8–I97.9, R00.1, R00.8
Ischemic heart disease	410–414	I20–I22, I24–I25, I51.3, I51.6
Rheumatic heart disease	390–398	I00-I09
Cerebrovascular disease	430–438	G45.0–G45.2, G45.4–G45.9, I60–I69
All external causes	E800–E848, E850–E888, E929–E978, E980–E999	V00-V99, W00-W99, X00-X84, X85-X99, Y00-Y36, Y40-Y89

Appendix 2

Table A2.1: Cox model estimates of the hazard ratio for selected cancer outcomes associated with exposure to **straight** metalworking fluids, controlling for other fluid types, calendar year, calendar year of hire, age, race, sex, and plant, when possible.

		Number of cases	HR	95% CI	
Laryngeal cance	er (73 cases)				
0	mg/m ³ ·years	40			
> 0 to 0.5	mg/m ³ ·years	17	1.18	(0.59, 2.37)	
> 0.5	mg/m ³ ·years	16	0.81	(0.41, 1.59)	
Lung cancer (18	-, -	-		(- ,)	
0	mg/m ³ ·years	964			
> 0 to 0.3	mg/m ³ ·years	309	1.07	(0.92, 1.25)	
> 0.3 to 1.6	mg/m ³ ·years	309	0.96	(0.82, 1.12)	
> 1.6	mg/m ³ ·years	309	0.91	(0.78, 1.05)	
Esophageal can	-,			, ,	
0	mg/m ³ ·years	83			
> 0 to 0.4	mg/m ³ ·years	31	1.17	(0.71, 1.91)	
> 0.4 to 2.1	mg/m ³ ·years	32	1.19	(0.74, 1.92)	
> 2.1	mg/m ³ ·years	30	1.16	(0.73, 1.85)	
Stomach cancer	· ·			, ,	
0	mg/m ³ ·years	103			
> 0 to 0.3	mg/m ³ ·years	30	1.16	(0.70, 1.91)	
> 0.3 to 2.9	mg/m ³ ·years	29	0.80	(0.49, 1.32)	
> 2.9	mg/m ³ ·years	30	1.86	(1.17, 2.97)	*
Colon cancer (4	-, -			(', ', ', ',	
0	mg/m ³ ·years	211			
> 0 to 0.5	mg/m ³ ·years	66	0.89	(0.64, 1.23)	
> 0.5 to 2.1	mg/m ³ ·years	65	0.93	(0.67, 1.29)	
> 2.1	mg/m ³ ·years	65	0.92	(0.68, 1.25)	
Rectal cancer (8	-, -			, , ,	
0	mg/m ³ ·years	44			
> 0 to 1	mg/m ³ ·years	20	0.78	(0.40, 1.51)	
> 1	mg/m ³ ·years	19	0.89	(0.47, 1.68)	
Bladder cancer	-, -			, ,	
0	mg/m ³ ·years	73			
> 0 to 0.3	mg/m ³ ·years	20	0.90	(0.51, 1.61)	
> 0.3 to 1.8	mg/m ³ ·years	22	0.65	(0.37, 1.13)	
> 1.8	mg/m ³ ·years	23	0.70	(0.42, 1.16)	
Liver cancer (12	-, -			, , ,	
0	mg/m ³ ·years	55			
> 0 to 0.5	mg/m ³ ·years	22	1.05	(0.58, 1.89)	
> 0.5 to 1.6	mg/m ³ ·years	23	1.90	(1.09, 3.31)	*
> 1.6	mg/m ³ ·years	23	1.24	(0.72, 2.13)	
Pancreatic canc	-, -			, , ,	
0	mg/m ³ ·years	154			
> 0 to 0.3	mg/m ³ ·years	54	1.25	(0.87, 1.81)	
> 0.3 to 1.1	mg/m ³ ·years	53	1.17	(0.80, 1.70)	
> 1.1	mg/m ³ ·years	54	0.85	(0.60, 1.21)	

Table A2.1: Cox model estimates of the hazard ratio for selected cancer outcomes associated with exposure to **straight** metalworking fluids, controlling for other fluid types, calendar year, calendar year of hire, age, race, sex, and plant, when possible.

		Number of cases	HR	95% CI	
Skin cancer (69	cases)				
0	mg/m^3 ·years	32			
> 0 to 0.9	mg/m^3 ·years	18	1.17	(0.58, 2.34)	
> 0.9	mg/m^3 ·years	19	1.32	(0.67, 2.58)	
Prostate cancer	(417 cases)				
0	mg/m^3 ·years	192			
> 0 to 0.5	mg/m^3 ·years	75	1.15	(0.84, 1.57)	
> 0.5 to 2	mg/m^3 ·years	75	1.11	(0.81, 1.52)	
> 2	mg/m^3 ·years	75	1.05	(0.78, 1.41)	
Brain and nervo	ous system canc	ers (128 cases)			
0	mg/m^3 ·years	74			
> 0 to 1	mg/m^3 ·years	27	0.61	(0.35, 1.05)	
> 1	mg/m^3 ·years	27	0.73	(0.44, 1.22)	
Leukemia (200 d	cases)				
0	mg/m^3 ·years	100			
> 0 to 0.3	mg/m^3 ·years	34	1.09	(0.69, 1.75)	
> 0.3 to 2.3	mg/m^3 ·years	33	0.77	(0.48, 1.23)	
> 2.3	mg/m^3 ·years	33	0.94	(0.60, 1.48)	
Breast cancer (76 cases)					
0	mg/m^3 ·years	43			
> 0 to 0.7	mg/m^3 ·years	16	1.34	(0.64, 2.82)	
> 0.7	mg/m^3 ·years	17	2.13	(1.04, 4.39)	*

Table A2.2: Cox model estimates of the hazard ratio for selected cancer outcomes associated with exposure to soluble metalworking fluids, controlling for other fluid types, calendar year, calendar year of hire, age, race, sex, and plant, when possible.

		Number of cases	HR	95% CI
Laryngeal cancer	(73 cases)			
0 to 0.05	mg/m^3 ·years	18		
> 0.05 to 7.3	mg/m^3 ·years	28	0.78	(0.39, 1.58)
> 7.3	mg/m^3 ·years	27	0.87	(0.42, 1.83)
Lung cancer (189	91 cases)			
0 to 0.05	mg/m^3 ·years	432		
> 0.05 to 3.3	mg/m^3 ·years	487	0.97	(0.84, 1.12)
> 3.3 to 11.2	mg/m^3 ·years	486	0.90	(0.77, 1.04)
> 11.2	mg/m^3 ·years	486	1.00	(0.85, 1.17)
Esophageal cance	er (176 cases)			
0 to 0.05	mg/m^3 ·years	34		
> 0.05 to 3.3	mg/m^3 ·years	46	1.05	(0.64, 1.74)
> 3.3 to 10.8	mg/m^3 ·years	47	1.04	(0.62, 1.74)
> 10.8	mg/m^3 ·years	49	1.18	(0.68, 2.04)
Stomach cancer ((192 cases)			

Table A2.2: Cox model estimates of the hazard ratio for selected cancer outcomes associated with exposure to **soluble** metalworking fluids, controlling for other fluid types, calendar year, calendar year of hire, age, race, sex, and plant, when possible.

		Number of cases	HR	95% CI	
0 to 0.05	mg/m ³ ·years	58			
> 0.05 to 4.2	mg/m ³ ·years	45	0.67	(0.42, 1.07)	
> 4.2 to 10	mg/m ³ ·years	44	1.07	(0.66, 1.74)	
> 10	mg/m ³ ·years	45	0.70	(0.42, 1.18)	
Colon cancer (40	7 cases)				
0 to 0.05	mg/m^3 ·years	80			
> 0.05 to 3.5	mg/m ³ ·years	109	1.14	(0.83, 1.57)	
> 3.5 to 12	mg/m ³ ·years	109	0.96	(0.69, 1.35)	
> 12	mg/m ³ ·years	109	0.94	(0.66, 1.33)	
Rectal cancer (83	3 cases)				
0 to 0.05	mg/m ³ ·years	20			
> 0.05 to 4.6	mg/m ³ ·years	19	0.85	(0.41, 1.76)	
> 4.6 to 8.6	mg/m ³ ·years	23	2.18	(1.07, 4.48)	*
> 8.6	mg/m ³ ·years	21	0.86	(0.40, 1.84)	
Bladder cancer (138 cases)			,	
0 to 0.05	mg/m ³ ·years	26			
> 0.05 to 3.7	mg/m ³ ·years	37	1.10	(0.63, 1.91)	
> 3.7 to 11.1	mg/m ³ ·years	36	1.18	(0.67, 2.07)	
> 11.1	mg/m ³ ·years	39	1.13	(0.63, 2.02)	
Liver cancer (123	3 cases)			,	
0 to 0.05	mg/m ³ ·years	22			
> 0.05 to 2.3	mg/m ³ ·years	34	1.29	(0.71, 2.33)	
> 2.3 to 9.2	mg/m ³ ·years	32	0.76	(0.41, 1.40)	
> 9.2	mg/m ³ ·years	35	0.93	(0.50, 1.74)	
Pancreatic cancer	-, -			, ,	
0 to 0.05	mg/m ³ ·years	70			
> 0.05 to 3.4	mg/m ³ ·years	82	0.81	(0.56, 1.16)	
> 3.4 to 9.3	mg/m ³ ·years	81	0.92	(0.63, 1.34)	
> 9.3	mg/m ³ ·years	82	0.80	(0.54, 1.19)	
Skin cancer (69 c	0, 0			, , ,	
0 to 0.05	mg/m ³ ·years	15			
> 0.05 to 4.7	mg/m ³ ·years	26	1.69	(0.77, 3.70)	
> 4.7	mg/m ³ ·years	28	1.64	(0.71, 3.80)	
Prostate cancer (, , ,	
0 to 0.05	mg/m ³ ·years	68			
> 0.05 to 5.1	mg/m ³ ·years	117	0.76	(0.54, 1.05)	
> 5.1 to 15.6	mg/m ³ ·years	116	0.79	(0.57, 1.11)	
> 15.6	mg/m ³ ·years	116	0.81	(0.57, 1.14)	
Brain and nervou	-, -			, ,	
0 to 0.05	mg/m^3 ·years	32			
> 0.05 to 2.7	$mg/m^3 \cdot years$	32	1.51	(0.86, 2.65)	
> 2.7 to 9.2	$mg/m^3 \cdot years$	32	1.48	(0.82, 2.68)	
> 9.2	$mg/m^3 \cdot years$	32	1.54	(0.82, 2.90)	
Leukemia (200 ca	-, -	- -	~ -	(, =:00)	
0 to 0.05	mg/m ³ ·years	48			
	G/ J ====				

Table A2.2: Cox model estimates of the hazard ratio for selected cancer outcomes associated with exposure to soluble metalworking fluids, controlling for other fluid types, calendar year, calendar year of hire, age, race, sex, and plant, when possible.

		Number of cases	HR	95% CI
> 0.05 to 3.3	mg/m ³ ·years	51	1.03	(0.65, 1.64)
> 3.3 to 9.7	mg/m^3 ·years	50	1.09	(0.67, 1.76)
> 9.7	mg/m^3 ·years	51	0.87	(0.53, 1.45)
Breast cancer (76	i cases)			
0 to 0.05	mg/m^3 ·years	36		
> 0.05 to 2.9	mg/m^3 ·years	22	0.59	(0.30, 1.14)
> 2.9	mg/m^3 ·years	18	0.56	$(0.26,\ 1.21)$

Table A2.3: Cox model estimates of the hazard ratio for selected cancer outcomes associated with exposure to **synthetic** metalworking fluids, controlling for other fluid types, calendar year, calendar year of hire, age, race, sex, and plant, when possible.

		Number of cases	HR	95% CI
Laryngeal cance	er (73 cases)			
0	mg/m^3 ·years	54		
> 0	mg/m^3 ·years	19	1.37	(0.67, 2.78)
Lung cancer (1	891 cases)			
0	mg/m^3 ·years	1374		
> 0 to 0.3	mg/m^3 ·years	173	0.93	(0.77, 1.12)
> 0.3 to 1.4	mg/m^3 ·years	172	1.07	(0.89, 1.29)
> 1.4	mg/m^3 ·years	172	1.00	(0.84, 1.20)
Esophageal can	cer (176 cases)			
0	mg/m^3 ·years	126		
> 0 to 0.7	mg/m^3 ·years	25	1.13	(0.66, 1.96)
> 0.7	mg/m^3 ·years	25	1.39	(0.84, 2.30)
Stomach cancer	(192 cases)			
0	mg/m^3 ·years	149		
> 0 to 0.5	mg/m^3 ·years	22	1.06	(0.61, 1.84)
> 0.5	mg/m^3 ·years	21	0.90	(0.53, 1.56)
Colon cancer (4	07 cases)			
0	mg/m^3 ·years	310		
> 0 to 0.4	mg/m^3 ·years	33	0.79	(0.52, 1.21)
> 0.4 to 1.7	mg/m^3 ·years	32	0.98	(0.65, 1.50)
> 1.7	mg/m^3 ·years	32	0.92	(0.62, 1.38)
Rectal cancer (8	33 cases)			
0	mg/m^3 ·years	59		
> 0 to 0.8	mg/m^3 ·years	12	1.49	(0.67, 3.27)
> 0.8	mg/m^3 ·years	12	1.64	(0.79, 3.41)
Bladder cancer	(138 cases)			
0	mg/m^3 ·years	104		
> 0 to 0.5	mg/m^3 ·years	17	0.91	(0.49, 1.70)
> 0.5	mg/m^3 ·years	17	0.67	(0.37, 1.21)
Liver cancer (12	23 cases)			

Table A2.3: Cox model estimates of the hazard ratio for selected cancer outcomes associated with exposure to **synthetic** metalworking fluids, controlling for other fluid types, calendar year, calendar year of hire, age, race, sex, and plant, when possible.

		Number of cases	HR	95% CI	
0	mg/m ³ ·years	89			
> 0 to 0.4	mg/m^3 ·years	17	1.17	(0.60, 2.26)	
> 0.4	mg/m^3 ·years	17	0.88	(0.48, 1.63)	
Pancreatic canc	er (315 cases)				
0	mg/m^3 ·years	227			
> 0 to 0.3	mg/m^3 ·years	30	0.98	(0.62, 1.55)	
> 0.3 to 0.9	mg/m^3 ·years	29	1.26	(0.80, 2.00)	
> 0.9	mg/m^3 ·years	29	0.87	(0.56, 1.34)	
Skin cancer (69	cases)				
0	mg/m^3 ·years	52			
> 0	mg/m^3 ·years	17	0.67	(0.34, 1.31)	
Prostate cancer	(417 cases)				
0	mg/m^3 ·years	300			
> 0 to 0.5	mg/m^3 ·years	39	1.01	(0.67, 1.50)	
> 0.5 to 2	mg/m^3 ·years	39	1.13	(0.76, 1.68)	
> 2	mg/m^3 ·years	39	1.30	(0.89, 1.89)	
Brain and nerve	ous system cance	ers (128 cases)			
0	mg/m^3 ·years	94			
> 0 to 0.6	mg/m^3 ·years	17	1.52	(0.80, 2.91)	
> 0.6	mg/m^3 ·years	17	1.36	(0.75, 2.47)	
Leukemia (200 cases)					
0	mg/m^3 ·years	142			
> 0 to 0.9	mg/m^3 ·years	29	1.11	(0.67, 1.82)	
> 0.9	mg/m^3 ·years	29	1.37	(0.86, 2.19)	
Breast cancer (76 cases)					
0	mg/m^3 ·years	60			
> 0	mg/m ³ ·years	16	0.76	(0.38, 1.53)	