Table 2.1 Cohort studies of inorganic acid mists and cancer

Reference, location, name of study	Cohort description	Exposure assessment	Organ site (ICD code)	Exposure categories	No. of cases/deaths	Relative risk (95% CI)*	Adjustment for potential confounders	Comments
	Metal pickling							
Ahlborg <i>et al.</i> (1981) Sweden	181 men employed 1951- 1979 in a stainless steel pickling house	Sulfuric and nitric acid in 1950s; oxalic acid, ammonium bifluoride, soap 1960s and 1970s	Larynx (161)	Cases in entire cohort	3	[50 (16.13- 155.03)]		
Beaumont et al. (1987), Steenland (1997), Steenland & Beaumont (1989), Steenland et al. (1988) USA	steelworkers employed ≥6 months 1940-1965 in pickling-related jobs at 3 steel-manufacturing facilities, vital status through 1986, cancer status for subset of 1031 through 1994	Sulfuric acid in 1975, 1977, 1979 averaged 0.19 mg/m³ for personal and 0.29 mg/m³ for area samples	Lung (162) Larynx (161)	Ever-exposed Those with 20 years latency  Ever-exposed Only sulfuric acid Only other acid Sulfuric acid daily	41 14 7 3 10	SMR 1.36 (0.97-1.84) 1.50 (1.05-2.27) RR 2.19 (1.2-3.7) 2.5 (1.7-4.7)	Smoking based on estimates from smoking survey of 841 workers or next of kin.	Incidence study did not include linking to cancer registries Other acid = hydrochloric, nitric, hydrofluoric, and hydrocyanic, "hydrochloric being the most common"
Mazumdar <i>et al.</i> (1975) USA	8465 men employed in 1953 in sheet and tin mill areas of 7 steel plants, vital status through 1966	Workers in pickling, coating, or specialty finishing areas (n=2822, 2763 white, 59 nonwhite) exposed to sulfuric, hydrochloric, or phosphoric acid fumes	All cancer Respiratory cancer	White males White males	248 85	1.08 1.15		Combined risk for departments exposed to acid mists: batch pickling and sheet dryers, coating, continuous pickling and electric cleaning, sheet finishing and shipping, stainless annealing, pickling and processing, and tin finishing and shipping not shown.

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Moulin et al. (2000) France	4288 male workers employed ≥1 year 1968- 1991 at a stainless steel and metal alloy production factory, vital status through 1992; nested case-control study of 54 lung cancer deaths with 3 controls per case, matched on sex and date of birth; Nested case- control study for lung cancer with 3 controls per case, matched on sex + birth date  Isopropyl or ethanol	Work histories assessed for exposure to metals, acid mists, other agents. JEM developed.	Larynx (161) Lung (162)	Employed in plant Employed in plant Exposed to acid mists	17 54 5	SMR 1.47 (0.9-2.4) SMR 1.20 (0.9-1.6) OR 0.57 (0.16-1.20)	France regional comparison rates  Smoking, PAH and silica exposure	Comparison region not specified No women died of lung or laryngeal cancer. Smoking history from medical records.
	alcohol manufacture							
Hueper (1966) USA	779 workers in isopropyl alcohol plant 1927-1950	Department	Nasal sinus (160) Larynx (161)	Entire cohort Entire cohort	2	<b>SIR</b> [21.3, 7.9-56.1]		Risk calculation for both sites combined
Lynch, <i>et al</i> . (1979) USA	Process workers (333 male, 2 female) employed ≥1 month 1950-76 + cohort of chemical mechanics, supervisors, & refinery workers (408 male) who worked on isopropyl or ethyl alcohol process units 1950-77; follow-up through 1977	Department & job title + personal recall by retirees & company supervisors & personnel employees	Larynx (161)	Entire cohort Process workers	7 4	SIR 3.2 [1.5-6.7] 5.0 [1.4-21.6]	US white male comparison rates	A nasopharyngeal cancer also was reported

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Soskolne et al. (1984) USA	50 men among those (n unspecified) who worked ≥12 months at plant, 1944-80. Ascertained: 14 while case employed, 12 as deceased annuitants, 3 ethanol unit follow-up, 8 living annuitant follow-up, 2 living annuitants who notified medical department, 11 noncompany tumour registry, all histologically confirmed; controls, 225 men from same cohort, matched (3+:1) by duration of employment ± 4 years, year of first employment ± 3 years, age ± 3 years, race)	Plant industrial hygienist assigned ordinal grade (0-5) of likely sulfuric acid, ethyl alcohol, isopropyl alcohol, asbestos, nickel, and wood dust exposures to department- job-era categories, analyzed by mean grade over work history. Analysis presented for sulfuric acid only. Moderate 1.10-2.09 mean grade, high 2.10-4.88 mean grade	nose, nasal	"moderate"	15 29	OR 5.2 (1.23-22.1) 2.9 (0.74-11.3)	Alcoholism, smoking (from plant medical records for those who did not leave employment <1955)	
Enterline (1982) USA	433 workers employed ≥3 months 1941-1965 in an isopropyl alcohol unit; 125 of 433 workers subsequently worked in jobs involving probable exposure to epichlorohydrin (ECH); follow-up through 1978	Department	Buccal cavity and pharynx (140-149)	Entire cohort Workers not subsequently employed in ECH unit	2 2	SMR 4.0 [1.0-16.0] 5.4 [1.4-21.6]	Texas comparison rates	

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Weil, <i>et al</i> . (1952) USA	182 workers employed 1928-1950 in an isopropyl alcohol unit	Department	Larynx (161) Nasal sinus (160)	Entire cohort Entire cohort	1 4			p value for 4 cases among 71 workers
05/1	isopropyr arconor ann		(100)	71 workers employed >5 years	4	P<1.0x10 <sup>-7</sup>		
Ott <i>et al</i> . (1989) USA	129 lymphohemato- poietic malignancies (excluding Hodgkin disease) on 5785 death certificates from cohort	Job histories evaluated to start of final case 5- year period of employment. 21 "suspect" substances	Non-Hodgkin lymphoma (200)	Alkyl sulfates Ethanol unit	8 5	OR 5.1 p<0.05 5.4 p<0.05		
	mortality study; Controls randomly selected from cohort of 29,139 men, matched to cases (5:1) by decade of 1 <sup>st</sup> hire, survival at least to start of same 5-year period of employment as cases.	and 1020 substances in 52 groups rated as ever/never in each	Multiple myeloma (202)	Alkyl sulfates	1	1.7		
Teta <i>et al</i> . (1992) USA	Isopropyl alcohol and ethyl alcohol male manufacturing workers 538 workers employed 1928-49 in isopropyl	Department and era	Buccal cavity and pharynx (140-149)	Entire cohort	2	SMR 1.5 expected [1.33] (0.2-4.8)	U.S. national comparison rates	SMRs not all presented but confidence intervals were
	strong acid unit and/or 1930-68 in ethyl strong		Larynx (161)	Entire cohort	1	[1.43] (0-8.0)		
	1930-68 in ethyl strong acid unit		Lympho- sarcoma, reticulo- sarcoma (200)	Entire cohort	5	5.6 (1.8-13.0)		

Table 2.1 Cohort studies of inorganic acid mists and cancer

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Teta <i>et al</i> . (1992) USA	Isopropyl alcohol and ethyl alcohol manufacturing male workers 493 workers (92	Department and era	Buccal cavity and pharynx (140-149)	Entire cohort	1	<b>SMR</b> [1.43] (0-8.4)	U.S. national comparison rates	SMRs not presented but confidence intervals were
	isopropyl, 316 ethyl) employed 1941-50 in		Larynx (161)	Entire cohort	1	[3.33] ( 0.1-18.6)		
	isopropyl strong acid unit, 1949-92 in isopropyl weak acid unit, 1941-68 in ethyl strong acid unit, and/or 1969-85 in ethyl weak acid unit		Leukemia, aleukemia (204-ICD7)	Entire cohort	2	[2.86] (0.3-10.3)		
Alderson & Rattan (1980) UK	262 men employed 1949- 80 in isopropyl alcohol unit for ≥1 continuous year	Department	Buccal cavity and pharynx (140-149)	Entire cohort	0	SMR	England and Wales national comparison rates	Average follow-up 15.5 years – possible insufficient latency for some workers
			Nasal sinus (160)	Entire cohort	1	50.0 p= 0.017		

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	Acid manufacture							
Englander <i>et al.</i> (1988) Sweden	400 men employed ≥6 months 1961-1981 in a sulfuric acid factory, vital status through 1985	Sulfuric acid area samples 0.1-3.1 mg/m³ , personal samples <0.1-2.9 mg/m³	Mortality all cancer  Morbidity Respiratory cancer	Entire cohort ≥5 years latency  Entire cohort ≥5 years latency	8 7 5 4	SMR 0.88, p>0.5 0.89, p>0.5 SIR 2.00, p=0.11 1.83, p=0.18	Mortality: Malmöhus county (excluding Malmo) rates. Incidence: Southern Swedish Regional Tumor Register	
			Gastroin-	Entire cohort	6	1.69, p=0.18		
			testinal cancer	≥5 years latency	5	1.64, p=0.24		
			Bladder	Entire cohort	5	3.77, p=0.01		
			cancer	≥5 years latency	5	4.36, p=0.006		
Hilt <i>et al</i> . (1985) Norway	287 men who worked at a nitric acid production plant 1928-1980, in Cancer Registry 1953- 1980	Exposed to nitric acid vapors, nitrous gases, ammonia, asbestos	Lung and pleural (162/3)	Entire cohort	17	<b>O/E</b> 4.5 (2.6-7.2)	Norwegian Cancer Registry	There was also a case of laryngeal cancer
Pesatori et al.	1409 (1372 male,	Sulfuric acid area				SMR	Tuscany comparison	505 of 1096 traceable male
(2006) Italy	including 1124 production workers and	sample medians: 1977, 1.4 mg/m <sup>3</sup> ; 1979, 0.9	Larynx (161)	Male production workers	4	1.30 (0.35-3.33)	rates	production workers previously worked in pyrite
	248 white-collar workers, 37 female) employed ≥1	mg/m <sup>3</sup> ; 1981, 1.2 mg/m <sup>3</sup> . Threshold limit		No prior pyrite exposure	2	1.61 (0.20-5.82)		mining; 594 did not.
	year 1962-1997 in a sulfuric acid factory, vital	value (1 mg/m <sup>3</sup> )	Lung (162)	Male production workers	27	0.82 (0.54-1.20)		
	status through 2000.			No prior pyrite exposure	5	0.39 (0.13-0.90)		
			Myeloid leukemia	Male production workers	4	3.10 (0.84-7.94)		
			(205)	No prior pyrite exposure	3	5.23 (1.08-15.3)		

Table 2.1 Cohort studies of inorganic acid mists and cancer

Reference, location, name of study	Cohort description	Exposure assessment	Organ site (ICD code)	Exposure categories	No. of cases/deaths	Relative risk (95% CI)*	Adjustment for potential confounders	Comments
Rapiti, et al. (1997) Italy	505 men employed for any time from 17 February 1954 through 31 August 1970, vital	125 ever worked in acid mixtures work process (sulfuric acid, nitric acid)	Cancer	Entire acid mix group		<b>SMR</b> 1.04 (0.67-1.53)	Italian regional comparison rates; 90% confidence intervals	Factory opened 1913, no pre- 1954 data, no analysis by duration of exposure or separately for those
status through 1991		Lung (162)	Entire acid mix group 30+ years latency	8	1.62 (0.81-2.92) 1.99 (0.87-3.92)		employed in earlier years Comparison region not specified	
	Soap manufacture							
Forastiere <i>et al.</i> (1987) Italy	361 men employed ≥1 year 1964-1972, vital status 1969-1983, cancer incidence 1972-1983	Sulfuric acid vapors in hydrolysis and saponification areas, 0.64-1.12 mg/m <sup>3</sup>	Larynx (161)	Entire cohort	1 death 5 cases	SMR 2.30 (0.1-11.4) 0.72 or 1.44 expected, depending on reference rates	All laryngeal cancer cases were smokers	
			Lung (162)	Entire cohort	5 deaths	1.69 (0.55-3.9)		
	Phosphate processing & fertilizer manufacture							
Block et al.	3451 males (2610 white,	Industrial hygiene				SMR	US national	
(1988)	841 black) employed	analysis found only	Lung (162)	Nonwhite males	9	1.04	comparison rates	
USA	≥6 months between 1950	Maintenance and		White males	37	1.62 p<0.05	No lung cancer in non-	
	and 1979 in phosphate	Chemical/ Fertilizer		Employed ≥1 yr	27	1.42	smokers in group for	
	fertilizer manufacture,	jobs had potential		1-9 years	9	0.86	which smoking data	
	vital status through 1981	exposure to dust, fumes [mists?], or		With 20+ years latency	4	1.22	were available	
		radiation		10-19 years	8	1.64		
				With 20+ years	3	1.61		
				latency				
			Larynx (161)	≥20 years	10	2.48 p<0.05		
				Nonwhite males	0	1.01		
				White males	2	1.91		

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Checkoway, et al. (1996), Checkoway et al. (1985a), Checkoway et al. (1985b) USA	22992 males (18446 white, 4546 nonwhite) employed in phosphate processing at 16 companies ≥3 months 1949-1978 and ≥12 months total, vital status through 1992	Job and department classified by exposure level to 6 agents	Lung (162)	White males Nonwhite males ≥5 years sulfuric acid production: Whites Nonwhites ≥5 years phosphoric acid production: Whites	354 105 5 1	SMR 1.19 (1.07-1.32) 1.13 (0.92-1.37)  1.15 2.07	US national comparison rates	Vital status was updated but work histories were not	
				Nonwhites	1	0.49			
Stayner <i>et al.</i> (1985) USA	3199 (2981 male, 212 female, 6 sex unknown) employed 1953-1976 in phosphate fertilizer manufacture, vital status through 1977	Industrial hygiene survey in 1976, 8 sulfuric acid samples, 0.013-0.22 mg/m <sup>3</sup>	Lung (162)	Entire cohort White males Nonwhite males	10 5 5	SMR (90% CI) 1.13 (0.6-1.9) 0.85 1.82	US national comparison rates		
	Nitrate and/or phosphate fertilizer manufacture								
Al-Dabbagh <i>et al</i> . (1986) UK	1448 (1327 male, 121 female) employed ≥1 year between 1946 and March 1981 in nitric	Exposure to nitrates but not to nitric acid mists estimated; compared with salivary	Oesophagus (150)	Men heavily exposed (to nitrates) All other men	3	[2.38 (0.77-7.38)] [2.27 (0.57-9.09)]	England and Wales national comparison	Includes only a 35% random sample of unvested employees who left work 1962-1972. No records for	
	acid or nitrate fertilizer	nitrate and nitrite in	Stomach (151)	Men heavily exposed	7	[0.97 (0.46-2.03)]	rates both used	pre World War II	
	manufacture, vital status through 1981	current workers		(to nitrates) All other men	5	[1.03 (0.43-2.48])		employment (plant opened 1923). No separate analysis for nitric acid plant.	
					Men heavily exposed	25	[1.19 (0.80-1.76)]		r
				(to nitrates) All other men	10	[0.68 (0.37-1.26)]			

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Bulbulyan <i>et al</i> . (1996) Russia	4996 (2039 men, 2957 women) employed ≥2 years between 1945 and 1985 in sulfuric acid	Cumulative exposure calculated for nitrogen oxides, SO <sub>2</sub> , and arsenic, but not acid	Stomach (151)	Sulfuric acid tower department Men	13 11	<b>SMR</b> 1.69 (0.90-2.89) 2.04 (1.02-3.66)	Moscow region comparison rates	Insufficient latency for any workers starting after 1980. No separate results for sulfuric acid contact,
	tower, sulfuric acid contact, ammonia, phosphoric acid, nitric	mists	Lung (162)	Sulfuric acid tower department	11	1.86 (0.93-3.34)		phosphoric acid, or nitric acid departments
	acid, or fertilizer manufacture departments of a large plant, vital status through 1990			Men	10	1.87 (0.90-3.44)		
Fandrem et al.	2023 men employed	Measured nitrate dust		Nitrate index		SIR	Norway national	Factory complex includes
(1993) Norway	>1 year in calcium	levels used to construct	Stomach (151)		6	0.85 (0.31-1.85)	comparison rates	nitric acid manufacture plant;
(-,,-,-,-,-,,	nitrate, fertilizer, or	job exposure matrix;		20-39 level-years	3	0.83 (0.17-2.44)	1	this was not one of the
	related departments and	exposure to nitric acid		40-59 level-years	3	1.07 (0.21-3.14)		"related departments" in the
	first employed between 1945 and 1979, cancer	mists not estimated		60+ level years	3	0.88 (0.17-2.51)		cohort
	incidence 1953-1988		Lung (162)	1-19 level-years	14	1.19 (0.65-1.99)		
				20-39 level-years	5	0.91 (0.29-2.13)		
				40-59 level-years	5	1.16 (0.37-2.72)		
				60+ level years	6	1.02 (0.37-2.22)		
Hagmar et al.	2,131 males employed >3	Nitrate exposure				SIR	>10 years latency	
(1991) Sweden	months between 1963	calculated &	Salivary gland	Nitrate workers	2	6.45 (0.78-23.3)	period	
	and 1985 in nitrate	cumulative dose	(142)	Phosphate workers	2	4.41 (0.53-15.9)	Southern Sweden	
	fertilizer manufacture;	estimated for 1460		•			comparison rates	
	1,236 males employed $\geq 3$	nitrate workers;	Lung (162)	Nitrate workers	10	0.90 (0.4-1.7)	•	
	months between 1906 and 1962 in phosphate	exposure to nitric acid mists not estimated		Phosphate workers	24	1.50 (0.98-2.3)		
	fertilizer manufacture, cancer morbidity 1958- 1986	mists not estimated						

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Rafnsson & Gunnarsdóttir (1990) Iceland	603 men hired from 1954 through 1985 at a nitrate fertilizer plant	Plant produces nitric acid and uses it to make fertilizer, but exposure to nitric acid mists not estimated	Lung (162) Stomach (151)	Entire cohort	4	<b>SMR</b> 1.39 (0.38-3.56) 0.93 (0.25-2.37)	Icelandic national comparison rates	Vital status through 1985? (not stated explicitly). 1974- 85 hires (n=252) have insufficient latency
Zandjani <i>et al.</i> (1994) Norway	1756 men employed ≥1 year between 1947 and 1980 in a nitrate fertilizer plant, vital status and cancer incidence through 1992	Departments include nitric acid, but exposure to nitric acid mists not estimated	Stomach (151) Testis (178)	Hired before 1960 Hired 1960 or later Hired before 1960 Hired 1960 or later	27 1 2 5	SIR 1.50 p<0.05 0.53 1.25 3.33 p<0.05	Rural Norway comparison rates	Results presented for some departments but not the nitric acid department
	Battery manufacture							
Coggon <i>et al.</i> (1996) UK	4401 men who worked 1950-1990 at two battery manufacture (1243 workers) and two steel	Work histories reviewed and classified by likelihood of acid mist exposure	Lung (162)	Definitely exposed to acid mists	83	<b>SMR</b> 0.98(0.78-1.2)	UK national comparison rates	Results not presented separately for battery and steel plants
	works factories (3158 workers) (different dates for each factory), vital status to 1993; 2678 (772 battery, 1906 steel works) definitely exposed to acid mists		Larynx (161)	Definitely exposed to acid mists	1	0.48 (0.01-2.7)		

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Coggon et al. (1996) (contd.)	Nested case-control study 15 men with upper aerodigestive cancer from a cohort of 4401 men	reviewed and classified by likelihood of acid mist exposure. High	Upper aerodigestive (140-141, 143-149, 160-	Maximum acid exposure: Low High	2 9	OR 1.9 (0.2-15.8) 1.3 (0.3-5.7)	>	
	who worked 1950-1990 at 2 battery manufacture and 2 steel works factories, 75 controls	exposure=>1 mg/m <sup>3</sup>	161)	Uncertain Cumulative acid exposure: Intermediate or	5	1.0 (0.1-12) 1.0 (0.2-4.6)		
	from the cohort, matched by date of birth and factory			uncertain ≥5 years high exposure	7	2.0 (0.4-10)		
Cooper <i>et al</i> . (1985) USA	4519 men employed ≥1 year 1947-1970 in 10 battery plants, vital		Lung (162)	Entire cohort employment:	109	<b>SMR</b> 1.24 (1.02-1.5)	US white male comparison rates	No mention of acid in paper
	status through 1980			1-9 years 10-19 years ≥20 years	20 21 53	1.67 p<0.05 1.56 1.05		
			Larynx (161)	Entire cohort employment: 1-9 years ≥20 years	6 1 4	1.28 (0.47-2.8) 1.85 1.41		
Malcolm & Barnett (1982) UK	1898 pensioners (1644 men, 254 women) employed 1892-1976 at 4 acid battery factories	Lead exposure classified as none, light, high. Acid exposure not categorized	All cancer	Males Females	293 25		England and Wales national comparison rates	No records on unvested former workers

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Sorahan & Esmen (2004) UK	926 men employed ≥1 year 1947-1975 in a nickel-cadmium battery plant, vital status through	Cumulative cadmium exposure from work histories and jobexposure matrix.	Pharynx (146- 149)	Entire cohort	4	<b>SMR</b> 5.59 (1.52-14.3)	England and Wales national comparison rates	
	2000	Sulfuric acid used 1947-1972 to dissolve	Larynx (161)	Entire cohort	2	1.95 (0.24-7.06)		
		nickel powder	Lung (162)	Entire cohort Hired 1947-1956 Hired 1957-1966 Hired 1967-1975  1-9 years from year of hire 10-19 years from year of hire 20-29 years from year of hire ≥30 years from year of hire	45 29 12 4 7 6 15	1.11 (0.81-1.48) 1.31 0.94 0.69 p-trend=0.15 1.49 0.63 1.21 1.21 p-trend=0.72		
Bond et al. (1986), Bond et al. (1991) USA	OTHER INDUSTRIES  308 male decedents from cancers of the lung, trachea, and bronchus from a cohort of 19,608 employees  308 male decedents (other causes) and 308 living from same cohort, matched to cases on race, birth year ± 5 years, year of hire, death year  ≤5 years (for decedents)	Industrial hygienist reviewed each job for hydrochloric acid exposure, based on monitoring data 1974- 1983	Lung (162)	Hydrochloric acid ppm-year 0.1-3.9 4.0-12.4 ≥12.5	62 45 22	RR  0.9 (0.6-1.3) 1.2 (0.8-1.9) 1.0 (0.6-1.8) p-trend = 0.35	Smoking	Referent group is the unexposed Matching on year of birth, year of first employment, and year of death may be overmatching "Living" controls could also be deceased if they died after the cases. Fifty decedent controls and 14 of the 97 dead "living" controls died of cancer.

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Grimsrud et al. (2005) Norway	213 lung cancer cases who worked ≥1 year in a nickel refinery 525 controls (worked ≥1 year in the refinery, matched by sex and birth year, free of lung cancer at time case diagnosed	Refinery exposure assessment, interview re other jobs	Lung (162)	Sulfuric acid mist $(mg/m^3)$ x years Low (0.02-0.7, mean $0.3 \pm 0.2$ ) Medium (0.7-4.0, mean $2.3 \pm 1.0$ ) High (4.1-37, mean $9.1 + 7.0$ )	9 11 14	Adjusted OR 1.0 (0.4-2.3) 1.0 (0.4-2.4) 0.8 (0.3-2.0)	Smoking; exposures to nickel, arsenic, asbestos, and cobalt in the refinery; exposure to carcinogens in other employment	Unexposed are the referent group