

Further follow up of mortality in a United Kingdom oil refinery cohort

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Abstract

The results of an extension of follow up (1976 to 1989) of a cohort of workers employed for at least one year between 1 January 1950 and 31 December 1975 at eight oil refineries in Britain are reported. Over 99% of the workers were successfully traced to determine their vital status at 31 December 1989. The mortality observed was compared with that expected from the death rates of all the male population of England and Wales and Scotland. The mortality from all causes of death for the total study population was lower than that of the comparison population, and reduced mortality was also found for many of the major non-malignant causes of death. Raised mortality patterns were found for diseases of the arteries, in particular aortic aneurysm, and accidental fire and explosion, for the total study population, and across several refineries and other subgroups. Mortality from all neoplasms was lower than expected overall, largely due to a deficit of deaths from malignant neoplasm of the lung. Raised mortality from all neoplasms was found for labourers and in particular for malignant neoplasms of the oesophagus, stomach, and lung, although the mortality was also high for all men in this social class in the national population. Regional variations may have accounted for some of the high mortality. There were other raised mortality patterns in malignant neoplasms of the intestine, rectum, larynx, and prostate but these tended to be isolated and not consistent across refineries and other subgroups. As in the earlier follow up there was raised mortality from melanoma in several job groups.

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Previous publications^{1,2} document the initial follow up from 1951 to 1975 of employees in eight United Kingdom oil refineries. This paper describes the results of extending this follow up to 31 December 1989. The detailed results for kidney cancer and leukaemia are reported elsewhere.³

Many studies have documented the carcinogenic properties of oil both in experimental^{4,5} and epidemiological studies.^{6,7} Others have investigated the carcinogenic and toxic properties of petroleum⁸ and its effect on the kidney.⁹

Several papers and unpublished reports have described past or ongoing studies of the mortality and less often the morbidity, of refinery populations.¹⁰⁻²¹ Four papers²²⁻²⁵ provide narrative epidemiological reviews. There is considerable variation in the dates, duration of follow up, and numbers in these studies. Those lost to follow up varies from 1% to 16%. Most of the studies show the healthy worker effect²⁶ to some extent, an effect that is reflected in many of the numerically large non-malignant diseases such as respiratory and cardiovascular diseases.

Most of the papers have focused on malignant diseases. The mortality from all malignant diseases tended to be lower than expected. In several studies the significant deficit from all malignant diseases was mainly due to a significant deficit from cancer of the lung.^{13,15,19,27} Smoking history is not usually available in these studies although there is some suggestion that oil refinery workers smoke less than comparable groups in other industries.²⁸ By contrast Bertazzi *et al*¹¹ found a significant excess from lung cancer when Italy was used as a comparison population.

Bearing in mind the problems of small numbers of deaths and multiplicity of statistical tests several studies have found excesses in specific cancer sites. Increased mortality (albeit mainly statistically non-significant) has been found for the total study population or in subgroups for cancers of the liver,^{10,14,29} pancreas,^{13,14,30-32} digestive system,¹⁶ stomach,^{14,33} intestines and rectum,^{34,35} prostate,^{21,30,31} kidney,^{11,13,30,36} and brain,^{10,11,13,14,17,19,20,29,30,33} and melanoma,^{16,27,29} and leukaemia.^{11-13,19-21,31,33}

A few studies of resident populations in areas where the petroleum industry is concentrated have

suggested links between certain diseases and the industry.³⁷⁻³⁹ As Harrington²³ points out, however, these types of studies are at best hypothesis generating exercises and should therefore be interpreted with caution.

Results of the Institute of Petroleum follow up study, 1951-75

The overall mortality in the previously reported results for the Institute of Petroleum refinery follow up study to 1975¹ was considerably lower than expected as was the mortality from heart disease, stroke, bronchitis, and pneumonia. The lowered mortality from all neoplasms was due to a large deficit of observed deaths from lung cancer. Raised mortality patterns were found in several refineries for cancers of the oesophagus, stomach, intestines, and rectum with significantly more deaths than expected from cancer of the nasal cavities and sinus and from melanoma.

Methods

Full details of the feasibility study carried out before the initial data collection, the reasoning behind the choice of data, items to be collected, and the methods used have been presented earlier.^{1,2} The study included all men with a duration of service of at least one continuous year between 1 January 1950 and 31 December 1975 at eight refineries in the United Kingdom. In this paper the refineries are denoted A, B, C, D, F, G, H, and J.

Data collected from personnel records comprised full name, date of birth, address, date of joining, last job (or present job for those in employment on 1 January 1976), whether a day or shift worker, National Insurance (NI) number and NHS number where available. For those who left before 1 January 1976, date of leaving and the reason for leaving were also collected.

In the previous follow up a one off tracing exercise was carried out with the National Health Service Central Registers (NHSCRs) at Southport and Edinburgh for men for whom the NHS number was available or the Social Security records at Newcastle otherwise. Full details on death were obtained for all those found to be dead either directly from the NHSCRs or from the Office of Population Census and Surveys (OPCS).

In the study reported here it was decided to carry out a flagging procedure at the NHSCRs. Details of all the study population not dead at 31 December 1975 were sent to the NHSCRs who identified those who had since died and provided the death certificates. Also it was decided to flag the records for the others. This flagging procedure enables the NHSCRs to identify the study population who die

in the future and to provide the details of death, a process that continues until the whole cohort dies. The death certificates provided multiple causes of death coded to the International Classification of Disease (ICD) revision in use at the time of death.

For those for whom the NHSCRs were unable to trace a death or general practitioner registration, details were sent to the Social Security records for one off tracing, from the NI number. For those identified as dead in this way a death certificate was then obtained.

When most of the flagging had been completed an examination of the records for those leaving before 1 January 1976 showed that of those flagged as alive at 31 December 1989, 13% were aged over 70 compared with 9% over 70 in the population of England and Wales. As incorrect flagging as alive could have resulted from failure to link the death record to the NHS record, it was decided to trace all these men (about 4000) through the National Insurance records for further deaths. Evidence of the incompleteness of the NHSCR records, particularly in the elderly, has also been shown by Darby *et al.*⁴⁰ As a result of this exercise a further 398 deaths that occurred before 1990 were identified.

Those who emigrated have been considered to be alive up to the date of emigration, this date therefore being their study end date. As all records were sent to the NHSCRs for flagging, more emigrations were identified than in the previous follow up in which much of the tracing was carried out through the National Insurance records, which do not identify emigrations.

As in the analysis of the first follow up,^{1,2} the expected deaths have been calculated by applying the five year age and calendar period specific death rates for England and Wales for the six English or Welsh refineries, and corresponding Scottish rates for the two Scottish refineries to the five year age and calendar period person-years at risk in the study cohort. Account was taken of the disparities between the four revisions of the ICD covered by the study period with standard bridging codes.

The standardised mortality ratio (SMR) was calculated from the ratio observed deaths: expected deaths multiplied by 100. The 95% confidence interval (95% CI) is given for each SMR.⁴¹ No formal adjustment has been made to take account of known regional variations in death rates. These, together with variations in death rates by social class, have been used to aid the interpretation of the results.

Many of the problems and limitations inherent in the interpretation of the results from this type of study have been discussed previously.^{1,2} This is mainly an a posteriori study and the data should be used principally to generate hypotheses to be further investigated in other studies on new data.

The study can, however, also be used as an a priori study to examine hypotheses suggested by other studies.

The use of hypothesis testing (leading to *p* values) is only strictly appropriate for the a priori approach. It is, however, widely and indiscriminately used in a posteriori epidemiological studies and can lead to a problem of multiple comparisons with the chance of producing a spurious statistically significant result in the absence of a real excess of mortality.⁴²

As pointed out by Gardner and Altman⁴¹ undue emphasis on hypothesis testing in medical studies has led to studies aimed at reaching statistical significance rather than determining the magnitude of any effects of interest. A preferable approach to significance testing is to use confidence intervals to examine the magnitude and patterns of mortality in the data across different variables and subgroups^{41, 42} rather than giving undue weight to isolated results.

In many industrial cohort studies of this type the overall SMR from all causes of death is found to be low—for example, around 90—and hence the cohort exhibits the healthy worker effect.²⁶ Consideration needs to be given in these cases to the interpretation of the SMRs for specific causes (which may be greater than 100).

The impact of the reanalysis of data in a cohort study after a further 14 years of follow up also needs to be considered because repeated analysis of accruing data also leads to a multiplicity problem. Allowance for this may be made in various ways, most simply by the use of techniques such as repeated confidence intervals.⁴³

Results

Table 1 shows the status of the study population at 31 December 1989. There were 34 569 men in the study (compared with 34 781 at 31 December 1975), 22 600 alive, 10 193 dead, 1 691 emigrated, and 85 untraced. As before untraced men have not been included in the analyses. Included in the alive group were 723 men whom it was not possible to flag at the NHSCRs but who have been traced as

Table 1 Vital status of refinery study population at 31 December 1989

Refinery	Alive	Dead	Emigrated	No trace	Total
A	1153	439	117	9	1718
B	2825	1480	277	40	4622
C	1606	712	50	1	2369
D	2909	1187	265	20	4381
F	2451	1803	130	1	4385
G	182	122	10	1	315
H	3540	1191	299	1	5031
J	7934	3259	543	12	11 748
Total	22 600	10 193	1691	85	34 569

alive at 31 December 1989 through the National Insurance records.

The smaller refinery population at the second follow up is because during the flagging and the merging of the new data base created for this follow up with old data files some duplicate records were identified. During the flagging process a few extra deaths were also identified for the previous follow up period. These, combined with the slightly smaller study population, have slightly altered the results for 1951-75 presented in this paper compared with those previously published.

Those dead included 70 for whom it was not possible to obtain a death certificate, 26 who were known to have died abroad, and 44 who were identified as dead by the National Insurance records but for whom no trace was found either in the English or Scottish death certificate records or the company pension records. In the analyses these 70 men were taken as dead, cause unknown.

Table 2 shows the person-years of observation for each follow up period for the total study population and for each refinery; there was an increase of roughly 360 000 person-years overall.

As described the information collected about job was the last job title for those who had left the refineries or the current title for those in post. Table 3 gives the distribution of the study population by job title. About a quarter of the total work-

Table 2 Person-years of observation of the refinery study population

Refinery	Person-years of observation	
	At 31 December 1989	At 31 December 1975
A	45 461	27 439
B	127 772	82 772
C	59 261	33 857
D	114 654	67 684
F	119 229	79 361
G	9093	6351
H	140 406	85 405
J	315 764	193 114
Total	931 640	575 983

Table 3 Refinery population by job classification

Job title	No (%)
Operator	9018 (26.1)
Maintenance	8211 (23.8)
General labourer	5098 (14.7)
Storeman	480 (1.4)
Driver	987 (2.9)
Fire and safety	1046 (3.0)
Foreman	2484 (7.2)
Scientific and technical	2110 (6.1)
Administrative and clerical	3438 (9.9)
Engineer	1697 (4.9)
Total	34 569 (100.0)

force were operators and a further quarter were craftsmen (analysed separately as fitters, pipefitters, riggers, and others). About 15% were labourers and 10% administrative, clerical and managerial groups. The other subgroups analysed separately, such as drivers, fire and safety workers, and foremen, consisted of small numbers, leading to small numbers of deaths and the corresponding problems of interpretation. Analyses by seven years of entry groups, (pre-1940, 1940-9, 1950-4, 1955-9, 1960-4, 1965-9, 1970-4) also yielded small numbers for

many disease groups making the patterns of mortality unclear. It should be noted that the study cohort is only complete for men joining after 1950 and before 1975 and staying for at least one year. Those that left before this date were excluded; this omission of shorter service workers starting before 1950 may thus have influenced the patterns of mortality.

Analyses by years since first employment (0-9, 10-19, 20-29, 30-39, 40-49, ≥ 50) are presented for selected causes. Time since first employment can be regarded as a proxy for duration of

Table 4 Observed (O) and expected (E) deaths, SMRs, and 95% CIs for non-malignant causes of death

Causes of death	Follow up 1951-75				Follow up 1976-89				Follow up 1951-89			
	O	E	SMR	(95% CI)	O	E	SMR	(95% CI)	O	E	SMR	(95% CI)
All causes	4498	4981.6	90	(88-93)	5695	6271.8	91	(88-93)	10 193	11 253.3	91	(89-92)
Respiratory TB	25	55.5	45	(29-67)	4	10.2	39	(11-100)	29	65.7	44	(30-63)
Diabetes	17	29.0	59	(34-94)	49	56.2	87	(65-115)	66	85.2	77	(60-99)
Cerebrovascular Disease	417	439.7	95	(86-104)	437	546.6	80	(73-88)	854	986.3	87	(81-93)
Chronic rheumatic heart disease	54	69.4	78	(58-102)	16	26.1	57	(32-95)	70	95.5	73	(57-93)
Ischaemic heart disease	1443	1495.8	96	(92-102)	1924	2125.4	91	(87-95)	3367	3621.2	93	(90-96)
Other heart disease	90	107.2	84	(68-103)	140	163.3	86	(72-101)	230	270.5	85	(74-97)
Hypertensive heart disease	76	96.5	79	(62-99)	34	51.3	66	(46-93)	110	147.8	74	(61-90)
Diseases of the arteries	93	87.5	106	(86-130)	205	160.6	128	(111-146)	298	248.1	120	(107-135)
Other circulatory disease	39	37.3	105	(74-143)	29	34.1	85	(56-122)	68	71.4	95	(74-121)
Pneumonia	163	181.2	90	(77-105)	228	269.2	85	(74-96)	391	450.4	87	(78-96)
Bronchitis and emphysema	257	345.4	74	(66-84)	168	234.8	72	(61-83)	425	580.2	73	(66-81)
Peptic ulcer	52	53.6	97	(72-127)	38	45.7	83	(59-114)	90	99.3	91	(73-111)
Cirrhosis of liver	14	21.4	65	(36-110)	24	37.2	65	(41-96)	38	58.6	65	(46-89)
Nephritis and nephrosis	32	38.3	84	(57-118)	7	9.6	73	(29-150)	39	47.9	82	(58-111)
Hyperplasia of prostate	15	16.1	93	(52-154)	13	11.0	118	(63-202)	28	27.1	103	(69-149)
Motor vehicle accidents	110	104.7	105	(86-127)	32	49.8	64	(44-91)	142	154.5	92	(77-108)
Accidental falls	23	30.1	76	(48-115)	27	29.4	92	(61-134)	50	59.5	84	(62-111)
Accidental fire and explosion	12	5.9	203	(105-355)	10	6.1	164	(79-301)	22	12.0	183	(115-278)
Suicide	64	85.9	75	(57-95)	43	58.8	73	(53-99)	107	144.7	74	(61-89)

Table 5 Observed (O) and expected (E) deaths, SMRs and 95% CIs for selected causes by time since first employment

Disease	Time Since First Employment (y)											
	0-9				10-19				20-29			
	O	E	SMR	(95% CI)	O	E	SMR	(95% CI)	O	E	SMR	(95% CI)
All causes	559	721.0	78	(71-84)	1585	1881.3	84	(80-89)	2934	3194.6	92	(86-95)
All neoplasms	154	165.7	92	(79-109)	446	486.7	92	(83-101)	854	866.8	99	(92-105)
Malignant neoplasm of stomach	24	19.9	121	(77-180)	61	55.5	110	(84-141)	94	86.7	108	(88-133)
Intestine	10	9.7	103	(50-190)	29	29.9	97	(65-139)	45	54.8	82	(60-110)
Rectum	6	6.7	90	(33-195)	18	21.2	85	(50-134)	39	39.2	99	(71-136)
Lung	46	58.4	79	(58-105)	157	196.8	80	(68-93)	349	356.7	98	(88-109)
Prostate	3	2.2	136	(28-396)	10	12.6	79	(38-146)	44	38.8	113	(82-152)
Melanoma	8	1.7	467	(202-920)	9	3.9	230	(105-437)	8	5.3	151	(65-298)
Leukaemia	10	7.6	132	(63-242)	13	13.5	96	(51-165)	19	19.1	99	(60-155)
Cerebrovascular disease	28	42.7	66	(44-95)	119	141.7	84	(70-101)	242	265.2	91	(80-104)
Ischaemic heart disease	140	158.7	88	(74-104)	487	584.8	83	(76-91)	1031	1100.0	94	(88-100)
Diseases of the arteries	4	6.0	67	(18-172)	20	26.1	77	(47-118)	78	65.6	119	(94-148)
Pneumonia	8	17.0	47	(20-93)	34	53.4	64	(44-89)	121	119.5	101	(84-121)
Bronchitis	8	31.7	25	(11-50)	88	109.4	80	(64-99)	132	171.1	77	(65-92)

exposure, although it was not possible to collect data on actual exposures to the multiplicity of chemicals and other potential hazards in an oil refinery for the total duration of the follow up.

NON-MALIGNANT CAUSES OF DEATH

Table 4 gives the observed and expected deaths, SMRs, and 95% CIs for the SMRs by non-malignant causes of death for the total refinery study population. These are presented separately for the first follow up, the second follow up, and the total follow up period.

The SMRs for all causes of death were similar for the two follow up periods, indicative of a continuation of the healthy worker effect. For many of the numerically large non-malignant causes of death, such as cerebrovascular disease, ischaemic heart disease, and pneumonia, the SMR for the second follow up period was lower than that of the first period. For other disease groups, such as bronchitis and cirrhosis of the liver, the SMRs for the two follow up periods were similar. A clear exception to this was diseases of the arteries where the SMR for the later follow up period was much larger than that of the first period.

The total follow up period results showed large deficits of observed deaths compared with expected deaths for the total study population for all causes, and many of the other disease groups. Only diseases of the arteries and accidental fire and explosion were clearly raised (as judged by the confidence interval).

The SMRs for all causes of death were very low for all the refineries with the exception of refinery F, where the observed deaths roughly equalled those expected. Mortality from all causes was also low for both shift and day workers and across all job groups except fire and safety workers, drivers, and storemen in whom the observed deaths roughly equalled those expected, and labourers in whom mortality was raised by about 10%. Table 5 gives

the observed and expected deaths, SMRs and 95% CIs by time since first employment for selected causes of death. There was a tendency for the all cause mortality to increase with increasing time since first employment and earlier year of entry.

Many of the numerically large non-malignant disease groups such as ischaemic heart disease, other heart disease, cerebrovascular disease, and pneumonia, showed the same patterns of mortality as the all cause mortality. Most of the job groups also had low mortality from these disease groups, with slightly higher mortality in fire and safety workers, drivers, and storemen.

Labourers, however, showed raised mortality both overall and from several disease groups. Table 6 gives the observed and expected deaths, SMRs and 95% CIs for labourers for several disease and refinery subgroups. The mortality from both all causes of death and all neoplasms was clearly raised both for the total labourer population and at several refineries. Similar patterns were found for several of the non-malignant disease groups—for example, cerebrovascular disease, ischaemic heart disease (though not pronounced), diseases of the arteries, other diseases of the circulatory system, pneumonia, and bronchitis and emphysema.

There were a few increases in mortality over expected numbers in some refinery and non-malignant cause of death combinations. Most of these were due to a corresponding increase in mortality in labourers—for example, this was true for ischaemic heart disease at refinery F (O=595, E=554.4, SMR=107, 95% CI 99-116) and pneumonia at refinery J (O=166, E=135.4, SMR=123, 95% CI 105-143).

An overall appreciable excess of observed deaths compared with those expected was found for accidental fire and explosion, (see table 4), especially at refinery F (O=10, E=1.6, SMR=641, 95% CI 307-1179) and refinery J (O=9, E=3.4, SMR=262, 95% CI 120-498). These deaths

30-39				40-49				≥ 50			
O	E	SMR	(95% CI)	O	E	SMR	(95% CI)	O	E	SMR	(95% CI)
5081	3242.6	95	(92-98)	1288	1388.3	93	(88-99)	746	824.8	90	(84-97)
826	895.1	92	(86-99)	320	354.6	90	(80-100)	173	180.9	96	(82-111)
74	83.0	89	(70-112)	33	34.4	96	(66-135)	16	16.9	94	(54-153)
70	58.8	119	(93-150)	20	23.8	84	(51-130)	12	13.3	90	(47-158)
37	40.3	92	(65-127)	16	16.2	99	(56-160)	10	8.6	116	(56-213)
305	351.8	87	(77-97)	100	138.0	72	(59-88)	62	64.6	96	(74-123)
55	60.0	92	(69-119)	38	29.0	131	(93-180)	21	20.2	104	(64-159)
4	4.2	95	(26-243)	1	1.2	81	(2-452)	0	0.5	0	(0-738)
19	18.5	103	(62-161)	5	7.2	69	(23-162)	2	3.9	51	(6-159)
266	291.7	91	(81-103)	116	143.8	81	(67-97)	83	101.2	82	(65-102)
1059	1091.0	97	(91-103)	423	444.1	95	(86-105)	227	242.6	94	(82-107)
107	82.5	130	(106-157)	54	39.7	136	(102-177)	35	28.2	124	(86-172)
123	128.0	96	(80-115)	57	69.2	82	(62-107)	48	63.3	76	(56-100)
112	148.5	75	(62-91)	60	76.2	79	(60-101)	25	43.3	58	(37-85)

Table 6 Observed (O) and expected deaths, SMRs, and 95% CIs for labourers for selected disease and refinery subgroups

Disease	Refinery	O	E	SMR	(95% CI)
All causes	A	83	99.9	83	(66-103)
	B	288	320.4	90	(80-101)
	C	269	256.4	105	(93-118)
	D	232	208.0	112	(98-127)
	F	644	536.0	120	(111-130)
	G	16	14.3	112	(64-182)
	H	174	162.8	107	(92-124)
	J	708	620.3	114	(106-123)
	Total	2414	2217.9	109	(105-113)
	All neoplasms	D	63	54.5	116
F		139	129.3	108	(90-127)
J		200	158.6	126	(109-145)
Total		609	557.8	109	(101-118)
Malignant neoplasm of stomach	D	12	5.6	215	(111-378)
	F	26	15.0	173	(113-254)
	J	34	17.0	201	(139-280)
Total	88	59.9	147	(118-181)	
Malignant neoplasm of lung	Total	254	221.3	115	(101-130)
	Total	24	16.8	143	(92-213)
Malignant neoplasm of oesophagus	Total	24	16.8	143	(92-213)
	Total	24	16.8	143	(92-213)
Cerebrovascular disease:	J	73	57.3	127	(100-160)
	Total	221	214.9	103	(90-117)
Ischaemic heart disease	F	186	162.4	115	(99-132)
	Total	221	214.9	103	(90-117)
Diseases of the arteries	F	27	13.7	198	(130-288)
	Total	64	52.0	123	(95-157)
Other circulatory diseases	Total	22	14.5	152	(95-230)
	Total	22	14.5	152	(95-230)
Pneumonia	J	51	28.0	182	(136-240)
	Total	121	101.8	119	(99-142)
Bronchitis and emphysema	F	61	36.9	166	(127-213)
	Total	166	131.9	126	(107-147)

occurred mainly in operators overall (O=11, E=3.0, SMR=364, 95% CI 182-652), and, at refinery F, in other craftsmen (O=3, E=0.15, SMR=2037, 95% CI 420-5954) and pipefitters (O=3, E=0.4, SMR=731, 95% CI 151-2135).

The only large non-malignant disease group to show consistently raised mortality across several refineries and many of the other subgroups analysed was diseases of the arteries (ICD-9th revision 440-448). Table 7 summarises the results for this disease group. The mortality increased in the second follow up period and was much increased for every five year age group after 60 with the exception of the over 85 group. Three refineries and several job groups showed increased mortality from diseases of the arteries and there was a pronounced trend for increasing mortality with increasing time since first employment (see table 5).

Of the 298 deaths from the broad disease category, diseases of the arteries, 73 were from atherosclerosis (E=70.6, SMR=103, 95% CI 81-130) and 187 were from aortic aneurysm. Table 8 summarises the results for aortic aneurysm.

Table 7 Observed (O) and expected (E) deaths, SMRs, and 95% CIs for diseases of the arteries by refinery and other subgroups

Subgroup	O	E	SMR	(95% CI)
Refinery:				
A	9	11.1	81	(37-154)
B	45	41.5	108	(79-145)
C	14	13.8	102	(56-170)
D	40	27.7	145	(103-197)
F	65	43.8	148	(115-189)
G	3	2.9	105	(22-306)
H	43	31.7	136	(98-183)
J	79	75.6	104	(83-130)
Age:				
< 60	31	37.8	82	(56-116)
60-64	40	31.2	128	(92-175)
65-69	54	43.8	123	(93-161)
70-74	69	50.0	138	(107-175)
75-79	55	42.6	129	(97-168)
80-84	34	27.1	125	(87-175)
> 85	15	15.6	96	(54-159)
Job:				
Operator	58	56.5	103	(78-113)
Fitter	22	12.2	181	(113-274)
Pipefitter	7	9.9	71	(28-146)
Rigger	6	5.1	117	(43-255)
Other craftsman	20	19.7	102	(62-157)
Labourer	64	52.0	123	(95-157)
Storeman	8	5.5	147	(63-289)
Driver	13	7.4	175	(93-299)
Fire and safety	16	12.7	126	(72-206)
Foreman	32	24.5	131	(89-184)
Scientific and technical	5	6.8	73	(24-171)
Administrator/clerical/managerial	34	26.8	127	(88-178)
Engineer	13	8.9	146	(78-250)

Mortality from aortic aneurysm was high in four of the eight refineries and accounted for the raised mortality from diseases of the arteries. The same job groups that showed raised mortality from diseases of the arteries showed raised mortality from aortic aneurysm. Only one refinery, F, showed raised mortality from atherosclerosis (O=27, E=16.9, SMR=159, 95% CI 105-232).

CAUSES OF DEATH FROM MALIGNANT DISEASES

Table 9 gives the observed and expected deaths, the SMRs, and the 95% CIs for the SMRs by malignant causes of death for the total refinery study population. These are presented separately for the first follow up, the second follow up, and the total follow up periods.

The SMRs for the two follow up periods for all neoplasms were identical and many of the malignant subgroups showed a decreased SMR in the second period—namely, malignant neoplasms of the oesophagus, stomach, intestine, rectum, liver and gall bladder, pancreas, and larynx, and melanoma. The opposite trend was seen for malignant neoplasms of the lung and pleura, bladder, and brain, whereas the SMRs for malignant neoplasm of the prostate were identical for the two periods and those for leukaemia similar. There

Table 8 Observed (O) and expected (E) deaths, SMRs, and 95% CIs for aortic aneurysm by refinery and other subgroups

Subgroup	O	E	SMR	(95% CI)
Refinery:				
A	7	6.6	106	(43-218)
B	27	23.0	117	(77-171)
C	5	8.1	61	(20-143)
D	26	16.8	154	(101-226)
F	29	20.4	143	(95-205)
G	0	1.7	0	(0-220)
H	34	17.7	192	(133-268)
J	59	41.6	142	(108-159)
Total	187	135.9	138	(119-159)
Age:				
< 60	22	22.2	99	(62-150)
60-64	31	20.4	152	(103-216)
65-69	35	29.2	120	(83-167)
70-74	51	29.5	173	(129-227)
75-79	34	21.8	156	(108-218)
> 80	14	14.1	99	(54-166)
Job:				
Operator	34	32.8	104	(72-145)
Fitter	15	6.7	226	(126-372)
Pipefitter	4	5.6	71	(19-183)
Rigger	3	2.9	103	(21-302)
Other craftsman	14	11.2	125	(68-210)
Labourer	30	25.7	117	(79-167)
Storeman	6	2.8	214	(79-466)
Driver	9	4.3	210	(96-399)
Fire and safety	11	6.2	177	(89-317)
Foreman	21	13.6	154	(95-235)
Scientific and technical	5	5.0	100	(32-233)
Administrator/clerical/ managerial	24	14.9	162	(103-240)
Engineer	11	4.9	224	(112-402)

were no further deaths from malignant neoplasms of the nasal cavities and sinus in the second follow up period.

Examination of the results in table 9 for the total

follow up period showed large deficits of observed deaths compared with expected deaths for the total study population for all neoplasms, malignant neoplasms of the buccal cavity and pharynx, and malignant neoplasms of the trachea, bronchus, and lung. There were a few SMRs greater than 100 for the total follow up period, but of these only melanoma was clearly increased.

Although the total number of observed deaths from all neoplasms was much fewer than those expected, three refineries (A, D, and J) had nearly the same observed deaths as expected. There was also a clear increase of about 10% in mortality from all neoplasms for labourers (see table 6).

Malignant neoplasms of the digestive tract

All these disease groups showed a tendency for a lower mortality in the second follow up period, (see table 9). Slightly increased mortality was found for malignant neoplasms of the oesophagus, especially at refinery D (O = 18, E = 11.4, SMR = 158, 95% CI 93-249), and for malignant neoplasms of the stomach, especially at refineries F (O = 56, E = 48.7, SMR = 115, 95% CI 87-149) and J (O = 111, E = 91.2, SMR = 122, 95% CI 100-147). There was raised mortality of over 40% overall for labourers in these two disease groups (see table 6), with excesses from malignant neoplasms of the stomach for labourers at refineries D, F, and J. Operators also showed raised mortality from malignant neoplasms of the oesophagus (O = 32, E = 23.0, SMR = 139, 95% CI 95-197) and stomach (O = 82, E = 70.5, SMR = 116, 95% CI 93-144).

Table 9 Observed (O) and expected (E) deaths, SMRs, and 95% CIs for malignant causes of death

Disease	Follow up 1951-75				Follow up 1976-89				Follow up 1951-89			
	O	E	SMR	(95% CI)	O	E	SMR	(95% CI)	O	E	SMR	(95% CI)
All neoplasms	1163	1234.6	94	(89-100)	1610	1715.2	94	(89-99)	2773	2949.8	94	(91-98)
Malignant neoplasm of buccal cavity and pharynx	9	18.4	49	(22-93)	15	26.0	58	(32-95)	24	44.4	54	(35-80)
Oesophagus	37	30.2	123	(86-169)	67	62.3	108	(83-137)	104	92.5	112	(92-136)
Stomach	169	150.8	112	(96-130)	133	145.6	91	(76-108)	302	296.4	102	(91-104)
Intestine	84	77.0	109	(87-135)	102	113.3	90	(73-109)	186	190.3	98	(84-113)
Rectum	59	55.9	106	(80-136)	67	76.4	88	(68-111)	126	132.3	95	(76-113)
Gall bladder	25	22.5	111	(72-164)	22	27.0	81	(51-123)	47	49.5	95	(70-126)
Pancreas	50	49.9	100	(74-132)	60	73.2	82	(63-106)	110	123.1	89	(73-108)
Larynx	13	12.1	107	(57-184)	13	16.1	81	(43-138)	26	28.2	92	(60-135)
Lung and pleura	421	508.3	83	(75-91)	611	658.0	93	(86-101)	1032	1166.3	88	(83-94)
Prostate	48	45.8	105	(77-139)	123	117.1	105	(87-125)	171	162.9	105	(90-122)
Bladder	34	42.9	79	(55-111)	64	67.5	95	(73-121)	98	110.4	89	(72-108)
Kidney	23	21.5	107	(68-161)	33	33.8	98	(67-137)	56	55.3	101	(77-132)
Brain	36	44.5	81	(57-112)	38	39.5	96	(68-132)	74	84.0	88	(69-111)
Melanoma	14	6.4	219	(120-367)	16	10.5	152	(87-247)	30	16.9	178	(120-254)
Reticulum cell sarcoma	5	7.8	64	(21-150)	1	1.6	63	(2-348)	6	9.4	64	(23-139)
Lymphosarcoma	11	8.2	134	(67-240)	7	4.4	159	(64-328)	18	12.6	143	(85-226)
Hodgkin's disease	13	16.4	79	(42-136)	4	7.0	57	(16-146)	17	23.4	73	(42-116)
Other neoplasms of the lymphoid tissue	4	4.3	93	(25-238)	23	27.4	84	(53-126)	27	31.7	85	(56-124)
Multiple myeloma	11	10.2	108	(54-193)	15	21.8	69	(39-113)	26	32.0	81	(53-119)
Leukaemia	30	31.2	96	(65-137)	38	38.6	98	(70-135)	68	69.8	97	(76-124)

Table 10 Observed (O) and expected (E) deaths, SMRs, and 95% CIs for melanoma, by refinery and other subgroups

Subgroup	O	E	SMR	(95% CI)
Refinery:				
A	2	0.88	228	(28-825)
B	5	2.5	202	(66-471)
C	3	1.0	287	(59-839)
D	1	2.2	45	(1-252)
F	2	2.1	94	(11-341)
G	0	0.2	0	(0-2052)
H	11	2.5	444	(222-795)
J	6	5.5	109	(40-238)
Total	30	16.9	178	(120-254)
Job:				
Operator	10	4.4	227	(109-418)
Fitter	1	0.9	110	(3-613)
Pipefitter	1	0.7	150	(4-837)
Rigger	0	0.4	0	(0-922)
Other craftsman	4	1.6	245	(67-627)
Labourer	3	2.6	117	(24-343)
Storeman	0	0.3	0	(0-1230)
Driver	1	0.53	188	(5-1048)
Fire and safety	0	0.5	0	(0-738)
Foreman	1	1.5	68	(2-379)
Scientific and technical	0	0.9	0	(0-410)
Administrative/clerical/ managerial	7	1.8	390	(157-803)
Engineer	2	0.8	253	(31-914)

Malignant neoplasms of the intestine, rectum, liver and gall bladder, and pancreas showed slightly lowered mortality overall although there was raised mortality at refinery F from malignant neoplasms of the rectum (O = 32, E = 21.3, SMR = 150, 95% CI 115-189); this excess was spread across many of the job groups. Thirteen scientists or technicians died from malignant neoplasm of the intestine (E = 6.4, SMR = 203, 95% CI 108-347).

There was a tendency for mortality from all the digestive neoplasms to decrease with increasing year of entry. Mortality from malignant neoplasms of the stomach tended to decrease with increasing time since first employment (table 5) but there was no clear pattern by time since first employment for the other disease groups.

Malignant neoplasms of the respiratory system

There were only 26 deaths from malignant neoplasms of the larynx; 15 of these were from refinery J (E = 8.7, SMR = 173, 95% CI 97-288) and they were spread across many job groups.

Although mortality from malignant neoplasms of the lung was again clearly decreased in the second follow up period the trend was for increasing mortality from this disease as the calendar period of follow up increased. Mortality was raised for riggers and storemen, and clearly raised for labourers (see table 6).

In the first follow up period there was an excess of observed deaths from malignant neoplasms of the nasal cavities and sinus compared with those expected, particularly at refinery J. There were no

further deaths from this cause in the second follow up period.

Melanoma

The raised mortality from melanoma found in the first follow up period continued in the second follow up period, although to a slightly lesser extent. Table 10 summarises some of the results for this disease. The observed deaths were more than those expected in five of the eight refineries but especially at refinery H. Excesses of observed deaths compared with those expected were found overall for operators, administrative, clerical and managerial staff, and other craftsmen. There was a tendency for mortality to increase with increasing year of entry, with a corresponding decrease by time since first employment (table 5).

Malignant neoplasms of the genitourinary system

In the second follow up period mortality from malignant neoplasm of the prostate was again slightly high, with that of malignant neoplasms of the bladder still low, although rather higher than in the first follow up period. The observed deaths from malignant neoplasm of the kidney were similar to those expected in both follow up periods.

There was clearly increased mortality from malignant neoplasms of the prostate in one refinery (B; O = 40, E = 27.7, SMR = 144, 95% CI 103-197). Eight of these men were foremen (E = 3.6, SMR = 222, 95% CI 96-437), and five were fitters (E = 1.1, SMR = 477, 95% CI 155-1114). There was also raised mortality overall from malignant neoplasms of the prostate in operators due mainly to increased mortality in operators at refinery J (O = 18, E = 11.9, SMR = 151, 95% CI 90-239).

Mortality from malignant neoplasms of the prostate and bladder tended to decrease with increasing year of entry whereas malignant neoplasm of the kidney showed the opposite pattern. There were no clear patterns of mortality by time since first employment.

Malignant neoplasms of the brain and central nervous system

The mortality from this disease group was higher in the second follow up period than in the first, but still lower than expected. There were no patterns of raised mortality for this disease group.

Malignant neoplasms of the lymphatic and haematopoietic tissue

In general the numbers of observed deaths in these disease groups were small, making interpretation of the results difficult. No raised mortality was found in three of the groups, namely reticulum cell sarcoma, Hodgkin's disease, and other neoplasms of the

lymphoid tissue. Mortality from lymphosarcoma increased in the second follow up period to give 18 deaths in total. Thirteen of these were from one company, at refinery H ($O=5$, $E=1.7$, $SMR=297$, 95% CI 97–694), and refinery J ($O=8$, $E=3.9$, $SMR=204$, 95% CI 88–401). Fifteen of the 18 deaths were day workers ($O=8.7$, $SMR=173$, 95% CI 97–286) but the deaths were spread across the job groups with several having slightly higher observed deaths than expected. Numbers were too small to show any patterns by year of entry or time since first employment. Mortality from multiple myeloma was lower than expected overall although one refinery (A) showed a pronounced excess based on only five deaths ($E=1.6$, $SMR=307$, 95% CI 100–717).

Details of the results for leukaemia have been reported elsewhere.³ Briefly, the observed and expected deaths were similar in both follow up periods and only two types showed raised mortality overall, other myeloid leukaemia ($O=7$, $E=1.0$, $SMR=678$, 95% CI 273–1397) and monocytic leukaemia ($O=7$, $E=3.2$, $SMR=221$, 95% CI 89–435). There were 14 observed deaths from leukaemia at refinery D ($E=8.6$, $SMR=164$, 95% CI 89–2751), 12 of which were myeloid leukaemia ($E=4.98$, $SMR=241$, 95% CI 124–421). This refinery showed excess mortality for acute myeloid leukaemia ($O=7$, $E=3.3$, $SMR=210$, 95% CI 84–433) and other myeloid leukaemia ($O=2$, $E=0.1$, $SMR=1621$, 95% CI 196–5854). There were also three deaths from leukaemia at refinery G ($E=0.8$, $SMR=393$, 95% CI 81–1149); two of these were chronic lymphatic leukaemias. Twenty one of the 68 deaths from leukaemia were in operators ($E=17.3$, $SMR=123$, 95% CI 75–186) and six of the seven deaths from other myeloid leukaemias were in operators.

Discussion

Before comment on the results of this cohort study it is appropriate to indicate some of the limitations. The study examines mortality only. It is therefore dependent on the completeness and accuracy of the information on cause of death given by the death certificates.

The SMR (and its confidence interval) has been used in this study to describe the patterns of mortality and to search for consistency across variables and subgroups examined. The interpretation of results in the light of an overall healthy worker effect and the impact of the reanalysis of the data after a further 14 years follow up, as presented in tables 4 and 9, must also be taken into account.

Differences in the mortality patterns between the eight refineries may be due to differences in the workforce or type of plant at the refineries. Details have been discussed previously^{1,2} but the refineries

are of varying age, size, and complexity. Two refineries, A and D, came into use in the 1950s and the others were all expanded at this time. The patterns of entry to and exit from the refineries were all similar with the populations expanding in the 1950s and decreasing in the 1960s. The distributions by birth year were also similar, with the exception of refinery F where 16% of the study population was born before 1900 compared with 5% in the other refineries. With the exception of refinery G, which was a bitumen manufacturing refinery, all had comparable crude oil processing plants. Refineries B and J also had some chemical plants but it was not possible to distinguish oil workers from chemical workers.

The large size of the cohort was the reason why only the last job title for leavers (or current title for those in employment at 31 December 1975) was collected to define the occupational group. Caution is thus necessary when interpreting the analyses by job defined in this way. No information was collected on job history either within or outside the oil industry; nor were data on exposures to potential hazards available. Methods of monitoring data on exposures to the many substances present were often unreliable, not regularly carried out, or not documented, until the late 1960s when the need for monitoring became more recognised.⁴⁴ It has been suggested that analyses of job title may detect associations that are not detectable by substance analysis, however, if the substance occurs in many occupations but is only a risk in the circumstances of a particular occupation.⁴⁵

Dates of leaving for those who were in post on 31 December 1975 were not obtained from the oil companies in this second follow up. For this reason, analyses by duration of employment could not be carried out. The mean duration of employment before 1976 was 13.2 years. Assuming those in post at 31 December 1975 worked an average of seven years in the second follow up the mean duration of employment of the cohort would be between 15 and 16 years by the end of 1989.

The person-years have increased by about 360 000 and the number of deaths has more than doubled from 4 406 in the first follow up to 1975 to 10 193 at 31 December 1989. The overall relative mortality of the study population, however, remained low, a continuation of the healthy worker effect. This may reflect both the initial selection of a healthy workforce and subsequent standard of medical care, industrial hygiene, and standard of living, and the comparison with the national population, which includes those in institutions, in unstable employment, or in ill health during their working life.

This lowered mortality was also reflected in many of the numerically large non-malignant causes of

death such as ischaemic heart disease and respiratory disease. Some of the individual job groups showed slightly raised mortality from specific non-malignant disease groups, in particular labourers. The expected deaths were not adjusted to take into account variations in mortality by social class. Many of the diseases for which raised mortality was found in labourers, such as cerebrovascular disease, bronchitis, and pneumonia, show clear trends in the population of England and Wales from low to high mortality for social classes I to V.⁴⁶ If adjustment had been made to the expected numbers of deaths for labourers with the data for social class V for these disease groups some of the raised mortality would be greatly diminished or disappear altogether.

Accidental fire and explosion, a plausible potential occupational hazard, continued to exhibit raised mortality although the number of deaths were small.

The consistently raised mortality from diseases of the arteries, and in particular from aortic aneurysm, is more difficult to interpret, because it is high across several refineries, age groups, and job groups. The patterns for this disease contrast with those of most of the heart disease groups, in which mortality is generally low. Increased mortality from aortic aneurysm has also been found in a study of Canadian oil distribution and marketing workers (AR Schnatter, unpublished data).

Mortality from all neoplasms was once again lower than expected, due mainly to pronounced deficits in malignant neoplasm of the lung. There was a trend, however, for increasing mortality from malignant neoplasms of the lung as the follow up period increased, with raised mortality for labourers. It has been suggested that the amount of smoking by refinery employees may be somewhat less than that of the general population.²⁸ It may be that smoking habits change after leaving and this may cause the mortality from lung cancer to increase as the cohort becomes older, although there are no data to substantiate this. Two small surveys (unpublished) carried out recently in two of the refineries in this study found that roughly 20% of those surveyed were current smokers compared with 33% of men aged 15 to 64 in England and Wales.⁴⁷ There was a trend for the proportion of smokers to increase as age increased in the refineries, similar to the trend in England and Wales. Only 15% of those aged under 40 smoked in the refinery survey, however, compared with 33% in the population of England and Wales. There was also a suggestion from one of the surveys, although based on small numbers, that those in the refinery smoked less than the general population, 40% in the refinery survey smoking less than 20 cigarettes a day compared with 18% of the men aged 15 to 64

in England and Wales who were current smokers.

There were some raised mortality patterns for malignant neoplasms of the digestive tract, in particular for malignant neoplasm of the stomach at refineries F and J. Two job groups were involved, operators and labourers, and the SMR for social class V in the general population was again high. No adjustment was made to the expected deaths to take into account known regional variations in mortality in England and Wales but both F and J refineries are situated in areas with high rates of stomach cancer (the SMR for the region in which refinery F is situated was 113 and that for the region in which refinery J is situated was 123)⁴⁸ and adjustment for both social class and regional variations would have reduced or removed the excess mortality.

The raised mortality from melanoma continued in the second follow up, in particular at refinery H, and was found in three very different job groups. Three other refinery studies have also found excesses from this disease^{16 27 29} but as yet no occupational cause linked to the oil industry has been suggested.

The slightly raised mortality from malignant neoplasm of the prostate continued in the second follow up period, with operators again having clearly increased mortality at refinery J. Overall mortality from this disease was also raised appreciably at refinery B. Several other studies have found raised mortality for malignant neoplasms of the prostate.^{21 30 31 35} Unlike other studies,^{18 49 50} the present study has not found any evidence of raised mortality for malignant neoplasms of the kidney or brain.

Particular interest has been paid to mortality from leukaemia and lymphosarcoma and their possible relation to exposure to benzene and other solvents.^{51 52} In this study there was increased mortality from lymphosarcoma at two refineries although there are only 18 deaths in total.

The total observed deaths from leukaemia were once again slightly lower than those expected and clearly increased mortality was found in only one refinery (D). This was from myeloid leukaemia in particular. Just under a third of the deaths from leukaemia were in operators.

Although the contributory causes of death on the death certificates were coded and entered into the database no detailed analyses of these have so far been carried out. The deaths with a malignant neoplasm as a contributory cause were investigated, however. For example, there were 17 contributory cases of malignant neoplasms of the stomach, 22 of the intestine, 42 of the lung, and 41 of the prostate. The ratio of the number of total mentions of each malignant neoplasm: the number of underlying causes of that disease group in the study population

was compared with the ratios from the population of England and Wales⁴⁶ and found to be similar.

In conclusion, the mortality of the oil refinery study population continues to be lower than that of the general population, both overall and from many of the numerically large disease groups. This current analysis has highlighted some issues that may warrant further investigation. The refinery population is now largely flagged at the NHSCRs and as further deaths occur in the cohort the death certificates will be provided. Further patterns of mortality can thus be monitored as required. Specific issues of particular concern—for example, the increased mortality from melanoma and aortic aneurysm—could be further investigated by carrying out nested case-control studies in which the deaths from the disease of interest are matched to controls selected from the rest of the study population. Further details such as job history and estimates of exposure to potential hazards might then be obtained and compared for the limited number of cases and controls.

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