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Original research

Cancer incidence in World Trade Center-exposed and non-exposed male firefighters, as compared with the US adult male population: 2001–2016

Mayris P Webber ^{1,2}, Ankura Singh ^{2,3}, Rachel Zeig-Owens ^{1,2,3},
Joke Salako ^{2,3}, Molly Skerker ^{2,3}, Charles B Hall ⁴, David G Goldfarb ^{2,3},
Nadia Jaber,² Robert D Daniels ⁵, David J Prezant ^{1,2,3}

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For numbered affiliations see end of article.

Correspondence to

Dr Mayris P Webber, Bureau of Health Services, Fire Department of the City of New York, Brooklyn, NY 11201, USA; mayris.webber@fdny.nyc.gov

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ABSTRACT

Objective To compare cancer incidence in Fire Department of the City of New York (FDNY) firefighters who worked at the World Trade Center (WTC) site to incidence in a population of non-WTC-exposed firefighters, the Career Firefighter Health Study (CFHS) cohort, and to compare rates from each firefighter cohort to rates in demographically similar US males.

Methods FDNY (N=10 786) and CFHS (N=8813) cohorts included male firefighters who were active on 11 September 2001 (9/11) and were followed until death or 31 December 2016. Cases were identified from 15 state cancer registries. Poisson regression models assessed cancers in each group (FDNY and CFHS) versus US males, and associations between group and cancer rates; these models estimated standardised incidence ratios (SIRs) and adjusted relative rates (RRs), respectively. Secondary analyses assessed surveillance bias and smoking history.

Results We identified 915 cancer cases in 841 FDNY firefighters and 1002 cases in 909 CFHS firefighters. FDNY had: higher rates for all cancers (RR=1.13; 95% CI 1.02 to 1.25), prostate (RR=1.39; 95% CI 1.19 to 1.63) and thyroid cancer (RR=2.53; 95% CI 1.37 to 4.70); younger median ages at diagnosis (55.6 vs 59.4; $p<0.001$, all cancers); and more cases with localised disease when compared with CFHS. Compared with US males, both firefighter cohorts had elevated SIRs for prostate cancer and melanoma. Control for surveillance bias in FDNY reduced most differences.

Conclusions Excess cancers occurred in WTC-exposed firefighters relative to each comparison group, which may partially be explained by heightened surveillance. Two decades post-9/11, clearer understanding of WTC-related risk requires extended follow-up and modelling studies (laboratory or animal based) to identify workplace exposures in all firefighters.

INTRODUCTION

Firefighters are repeatedly exposed to occupational hazards, including known carcinogens.^{1 2} There have been over 200 peer-reviewed studies of firefighting and cancer, shown in PubMed,^{3–9} most published since 2000. Despite this proliferation of studies, the degree to which firefighting is associated with cancer incidence remains uncertain. A 2014 National Institute for Occupational Safety and Health (NIOSH) study found that a cohort of

Key messages

What is already known about this subject?

► Firefighters are routinely exposed to known carcinogens. Evidence of an increased risk of cancer in firefighters remains mixed. Work exposures for firefighters at the World Trade Center (WTC) site, an especially toxic environment, further complicate risk assessment.

What are the new findings?

► WTC-exposed male firefighters had 13% higher all-cancer rates, and younger median age (55.6 vs. 59.4 years) and more localised disease at diagnosis compared with a cohort of non-WTC-exposed male firefighters. Both firefighter cohorts had elevated prostate cancer and melanoma rates compared with demographically similar US males.

How might this impact on policy or clinical practice in the foreseeable future?

► Surveillance bias may account for some of the excess cancers identified in both firefighter cohorts compared with similar US males. Two decades post-9/11, clearer understanding of the WTC-related cancer risk requires additional years of follow-up to allow for the reported long latency of some solid tumours. Laboratory-based or animal-based modelling studies should be encouraged to identify and track workplace exposures in WTC-exposed and non-WTC-exposed firefighters.

nearly 30 000 professional firefighters had a modest elevation in the rate of all-cancers combined (standardised incidence ratio (SIR)=1.09; 95% CI 1.06 to 1.12) compared with the general US population.⁴ A recent meta-analysis of 25 cohort studies, including the aforementioned NIOSH cohort, however, reported that the all-cancer risk for firefighters was similar to that of the general population (meta-relative risk=1.0; 95% CI 0.93 to 1.07).¹⁰

The World Trade Center (WTC) attacks on 11 September 2001 (9/11) exposed over 15 000 Fire Department of the City of New York (FDNY) firefighters and other rescue/recovery workers to



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carcinogenic substances^{11–13} including polychlorinated biphenyls (PCBs), polycyclic aromatic hydrocarbons (PAHs), asbestos, sulfuric acid, benzene and arsenic.^{2,13,14} Studies of WTC-exposed rescue/recovery workers have generally shown modestly elevated cancer rates compared with general populations.^{15–19}

The current study was undertaken to assess if work as an FDNY firefighter at the WTC site conferred a cancer risk above that attributed to firefighting under non-WTC conditions. Our 2016 study²⁰ compared post-9/11 cancer incidence in the FDNY WTC-exposed firefighter cohort to incidence in the aforementioned NIOSH cohort, hereafter called the Career Firefighter Health Study (CFHS), and found no difference between the cohorts in the rate of all cancers, although rates of some site-specific cancers were significantly elevated (eg, thyroid and prostate cancer) in FDNY. The current study extended follow-up to allow for the detection of cancers up to 15 years post-WTC exposure. We also used smoking data, when available, and we provide perspective for findings in firefighters (WTC-exposed and non-WTC-exposed) by comparing characteristics of these cases to those in the general population. This study is of importance to WTC research as it controls for both occupational exposures using a firefighter comparison group and US secular trends. Given the long latency of some cancers, the relationship between WTC exposure, firefighting and cancer is particularly worthy of close examination during this 20th anniversary year of the WTC attacks.

METHODS

Study population

Inclusion required that FDNY and CFHS members be actively employed by their respective departments on 11 September 2001. This study followed the Strengthening the Reporting of Observational Studies in Epidemiology guidelines.

FDNY cohort

The FDNY sample consisted of male firefighters who worked at the WTC site at any time between 11 September 2001 and 25 July 2002 (N=10 786). Male firefighters who did not report any WTC exposure (N=82) were excluded, as were WTC-exposed female firefighters (N=28) due to low numbers. Characteristics of the excluded female firefighters are shown in online supplemental table S1.

CFHS cohort

The CFHS cohort included 29 992 firefighters from the Philadelphia, Chicago, and San Francisco Fire Departments, originally identified from department rosters, as previously described.^{4,21} We chose this referent group because they were subject to similar pre-hire fitness requirements, worked in urban environments, and worked at departments with retirement policies similar to FDNY's. Study inclusion was limited to males who were actively employed on 11 September 2001 (N=8813); 585 females were excluded (online supplemental table S1). Since we were missing information on the WTC exposure history of CFHS members, we considered this cohort to be non-WTC-exposed.

Demographic/background information

Data were obtained from employee databases and FDNY medical records, and death information from the Social Security Death Master File and the National Death Index. One additional FDNY death was identified from employee records. Smoking status (current, former or never smoker) and prior military service were ascertained from health surveys completed

by FDNY firefighters during medical monitoring examinations. WTC-exposed FDNY members provided their initial arrival time at the WTC site, hereafter referred to as WTC exposure level, on their first post-9/11 health questionnaire.¹⁹ A similar health survey collected smoking status, occupational history and other background information from a sample of CFHS firefighters (N=2856, 32.4%).²²

Cancer cases

Information on cancer diagnoses was obtained by probabilistic matching to state cancer registries, as described elsewhere.^{4,19,20} States were selected for linkage based on the residence information of active and retired FDNY and CFHS firefighters. The FDNY population was linked to Arizona, Connecticut, Florida, North Carolina, New Jersey, New York, Pennsylvania, South Carolina and Virginia state cancer registries. The CFHS population was linked to Arizona, California, Florida, Illinois, Indiana, Michigan, New Jersey, Oregon, Pennsylvania, and Washington state. Ninety-eight per cent of FDNY firefighters and 97% of CFHS firefighters resided in one of the states selected for linkage. Cancer cases defined as primary malignant tumours or in situ bladder cancers²³ and diagnosed between 11 September 2001 and 31 December 2016 were analysed.

US cancer rates

We used the National Cancer Institute's Surveillance, Epidemiology and End Results Program (SEER-21) data to obtain rates of all-cancers combined and some site-specific cancers in the US male population.²⁴ Data were grouped by calendar year (2001 to 2016), race/ethnicity (non-Hispanic white, non-Hispanic Black, Hispanic, Asian and Native American) and 5-year age groups (15–19 to 85–89 years), and incidence rates calculated in these strata. We also obtained information on cancer stage at diagnosis²⁴ and median age at diagnosis.²⁵

Statistical analyses

Selected characteristics of each cohort were compared by calculating means (\pm SD), medians (IQR) or proportions (%). T-tests, Wilcoxon Mann-Whitney U tests or χ^2 tests were performed, as appropriate. Follow-up time started on 11 September 2001 and ended on 31 December 2016 or, if earlier, date of death.

To examine trends over time, we graphed cancer incidence rates by calendar year in FDNY, CFHS and US males (SEER-21) using output from Poisson regression models of year and group predicting cancer cases, with the log of person-years as an offset, controlling for race/ethnicity and 5-year age group. To provide context for the interpretation of cancer risk in firefighters, with and without WTC exposure, our primary analyses involved estimating SIRs and 95% CIs for all-cancer sites combined (all cancers) and some site-specific cancers²³ between 11 September 2001 and 31 December 2016, comparing each firefighter cohort with SEER-21.²⁴ We used Poisson regression models with observed numbers of FDNY cancer cases as the outcome variable and the log of the expected case count in each demographic stratum, calculated by multiplying the stratum-specific SEER-21 rate with the number of person-years belonging to that stratum in the FDNY cohort, as an offset. We then reran these models using CFHS in place of FDNY data. Site-specific cancers were chosen based on our previous work examining post-9/11 cancer incidence in FDNY firefighters.^{19,20}

Additional primary analyses fit Poisson regression models assessing the associations between firefighter cohort and cancer rates, controlling for age on 9/11 and race/ethnicity. Adjusted

relative rates (RRs) and 95% CIs were estimated using cohort as the independent variable, cancer cases as the outcome, and log of person-years as the offset. The RR for melanoma was estimated from non-Hispanic white males (N=16 238) due to limited cases in non-whites. A sensitivity analysis assessed differences in FDNY and CFHS cohort cancer rates when only first primary cancer cases were included as the outcome; this analysis excluded individuals whose first cancers were diagnosed prior to 9/11 (N=229) and ended follow-up at the first diagnosis date.

Secondary analyses addressed possible surveillance bias, as previously described.^{19 20} Briefly, WTC-exposed FDNY firefighters undergo free health monitoring exams without copays, including for blood testing, and chest CT scans, as indicated, even after retirement. FDNY cancers could therefore be diagnosed earlier due to screenings that are not widely available to others. Accordingly, we categorised cases of lung, liver, thyroid or kidney cancer or Hodgkin or non-Hodgkin lymphoma diagnosed ≤ 6 months after chest CT scans and cases of prostate or haematological cancers diagnosed ≤ 6 months after routine blood tests as cases identified by screening (N=204), repeating the primary analyses after delaying the diagnosis dates of these cancers by 2 years. This time delay was chosen based on the 2-year average lead time for prostate cancers diagnosed in participants of a randomised cancer screening study who received annual screening.^{20 26} We also performed sensitivity analyses delaying diagnosis dates by 5 years.

We conducted two additional secondary analyses: we analysed the subset of 10 723 FDNY and 2856 CFHS firefighters for whom we had smoking information, repeating the RR analyses comparing FDNY and CFHS cancer rates with smoking status (ever vs never) included as a covariate in the Poisson regression models. We then explored a possible dose-response relationship between WTC exposure level and cancer within the WTC-exposed FDNY cohort only (N=10 786), redoing the RR analyses with an ordinal WTC exposure level variable ranging from lowest to highest-level WTC exposure group¹⁹ (table 1) in the models.

All analyses were performed in SAS (V9.4, SAS, Institute Inc., Cary, NC, <http://www.sas.com>). Yearly incidence rate graphs were created via PROC SGPLOT using a locally weighted smoothing (LOESS) function for estimates generated from the first Poisson regression models described above.

RESULTS

Selected characteristics of 10 786 WTC-exposed FDNY and 8813 CFHS firefighters are shown in table 1. FDNY firefighters were younger, on average, more likely to be white and never-smokers, and less likely to be combat veterans compared with CFHS firefighters ($p < 0.001$ for all). The FDNY cohort was also consistently younger at diagnosis compared with CFHS: for all cancers (median (IQR)=55.6 (50.2–60.2) vs 60.0 (54.4–64.8) years) and for every cancer subtype examined. Since 9/11, 92.7% of WTC-exposed male FDNY firefighters have had at least 1 prostate-specific antigen (PSA) test, 98.8% at least 1 complete blood count and 47.7% at least 1 chest CT. While we lack comparable data for the CFHS, 96% of those who completed the CFHS health survey reported a visit to a medical doctor within the previous 2 years.

Between 11 September 2001 and 31 December 2016, we identified 915 and 1002 incident cancer cases in 841 FDNY and 909 CFHS firefighters, respectively. Prostate cancer was the most common site-specific cancer in both groups (N=332 and N=358 in FDNY and CFHS, respectively), followed by melanoma of the

Table 1 Population characteristics

	WTC-exposed male FDNY firefighters active on 9/11	Male CFHS firefighters actively employed on 9/11
Total N	10 786	8813
Age on 9/11, mean \pm SD*	40.4 \pm 7.5	43.9 \pm 9.2
Race/ethnicity, N (%)*		
Non-Hispanic white	10 121 (93.8)	6117 (69.4)
Non-Hispanic Black	282 (2.6)	1589 (18.0)
Hispanic	353 (3.3)	736 (8.3)
Other†	30 (0.3)	371 (4.2)
Smoking status, N (%)*		
Current	373 (3.5)‡	189 (6.6)§
Former	3233 (30.2)‡	1056 (37.0)§
Never	7117 (66.4)‡	1611 (56.4)§
Military combat experience, N (%)*	429 (4.1)¶	283 (9.9)§
WTC site arrival time, N (%)		
Exposure group 1 (highest): Morning of 9/11	1741 (16.1)	
Exposure group 2: Afternoon of 9/11	5683 (52.7)	
Exposure group 3: 12/9/2001	1873 (17.4)	
Exposure group 4: 13/9–24/9/2001	1315 (12.2)	
Exposure group 5 (lowest): After 24 September 2001	174 (1.6)	
Age at cancer diagnosis, median (IQR)		
All cancers*	55.6 (50.2–60.2)	60.0 (54.4–64.8)
Prostate cancer*	57.9 (53.3–62.4)	60.4 (55.2–65.4)
Lung cancer	60.4 (55.6–65.5)	62.8 (57.9–67.2)
Kidney cancer	56.0 (49.0–63.0)	58.1 (54.7–63.2)
Non-Hodgkin lymphoma**	53.6 (48.9–59.4)	60.1 (50.1–65.3)
Melanoma (skin)*	51.9 (45.5–57.8)	61.2 (54.4–66.5)
Thyroid cancer**	51.2 (44.0–56.5)	59.0 (49.2–64.5)
Alive at end of follow-up, N (%)*	10 525 (97.6)	8208 (93.1)
Follow-up time (years), mean \pm SD*	15.2 \pm 1.1	14.9 \pm 2.0

* $P < 0.001$, ** $P < 0.05$.

†Includes Asian and Native American race categories.

‡N=10 723 who self-reported smoking status.

§N=2856 who completed CFHS survey.

¶N=10 429 who self-reported occupational history.

CFHS, Career Firefighter Health Study; FDNY, Fire Department of the City of New York; WTC, World Trade Center.

skin (N=96) and non-Hodgkin lymphoma (N=55) in FDNY and lung cancer (N=83) and melanoma (N=70) in CFHS.

Cancer rates in firefighters (FDNY and CFHS) compared with US males (SEER-21)

Comparing median age at diagnosis across the three groups, we found that generally, FDNY firefighters had the youngest median age and SEER-21 the oldest.²⁵ Similarly, there were differences in cancer stage at diagnosis: FDNY firefighters were usually diagnosed at an earlier, more localised disease stage (table 2).

Figure 1 displays the race/ethnicity-adjusted and age group-adjusted cancer incidence rates by calendar year in the FDNY, CFHS and SEER-21 populations. Rates of prostate cancer (figure 1A), non-Hodgkin lymphoma (figure 1B) and melanoma of the skin (figure 1C) were consistently elevated in the FDNY cohort compared with US males; this was especially evident for prostate cancer rates after 2007. Prostate cancer and melanoma rates also appeared to be elevated in CFHS versus US males. In contrast, rates of lung cancer were lower in both firefighter cohorts than in US males (figure 1D).

Table 3A displays results of SIR analyses comparing cancer incidence in the FDNY and CFHS cohorts with expected

Table 2 Proportion of cancers* in localised, regional, distant or unknown stage at time of diagnosis (%)

Site		FDNY	CFHS	SEER-21
Prostate	Localised	78.9	77.7	62.0
	Regional	11.7	14.0	9.0
	Distant	2.1	1.7	4.0
	Unknown	7.2	6.7	24.0
Lung	Localised	45.5	15.7	14.2
	Regional	20.5	26.5	18.0
	Distant	29.6	51.8	43.5
Kidney	Localised	4.5	6.0	24.4
	Regional	79.5	67.3	55.3
	Distant	17.9	12.7	14.3
Non-Hodgkin lymphoma	Localised	2.6	16.4	12.7
	Regional	0	3.6	17.7
	Distant	32.7	32.6	22.3
	Unknown	9.1	13.9	12.0
Melanoma (skin)	Localised	49.1	41.9	41.9
	Regional	9.1	11.6	23.7
	Distant	71.2	78.6	68.5
	Unknown	4.2	2.9	8.0
Thyroid	Localised	4.2	7.1	4.1
	Regional	19.8	11.4	19.3
	Distant	65.2	53.3	51.6
	Unknown	32.6	16.7	28.8
Thyroid	Distant	0	0	5.7
	Unknown	2.2	0	13.8

*All malignant cancers (multiple primaries).

CFHS, Career Firefighter Health Study; FDNY, Fire Department of the City of New York; SEER, Surveillance, Epidemiology, and End Results Program.²⁴

numbers based on SEER-21 rates. In the FDNY cohort, SIRs for all cancers, prostate cancer, thyroid cancer, melanoma of the skin, and non-Hodgkin lymphoma were significantly elevated, whereas among CFHS firefighters, prostate cancer and melanoma SIRs were significantly elevated. Lung cancer incidence was significantly lower than expectation in both cohorts.

After correction for possible surveillance bias by delaying the diagnosis dates of certain cancer cases by 2 years, the FDNY all-cancer incidence remained higher than expected compared with the US male population (SIR=1.09; 95% CI 1.02 to 1.16), as did the incidence of prostate cancer, thyroid cancer and melanoma (table 3B). In a sensitivity analysis that delayed diagnosis dates by 5 years, all-cancer incidence was no longer significantly elevated in the FDNY cohort versus the US male population. SIRs for prostate and thyroid cancer were attenuated, but continued to show modestly higher rates in the FDNY cohort (data not shown).

Cancer rates in FDNY compared with CFHS

Comparing the FDNY and CFHS cohorts directly, the FDNY cohort had significantly higher rates of all cancers, prostate and thyroid cancer (table 4). A sensitivity analysis that included only first primary cancer cases when estimating RRs showed similar associations between firefighter cohort and all cancer, prostate and thyroid cancer rates (data not shown). Race/ethnicity was generally not associated with cancer in the above models, although non-Hispanic Black race was significantly associated with prostate cancer (RR vs non-Hispanic white=1.89, 95% CI 1.55 to 2.31).

Table 4 also shows results corrected for potential surveillance bias in FDNY using a 2-year lag in diagnosis dates of certain cancers: after correction, the RRs were slightly attenuated, but prostate and thyroid cancer rates remained significantly elevated in FDNY firefighters. In sensitivity analyses which used a 5-year lag in diagnosis dates, the RRs were further attenuated for all cancers and for prostate, lung and thyroid cancer (data not shown). Rates of prostate and thyroid cancer in the FDNY cohort, while still elevated, were no longer significantly different from the CFHS rates.

In the analysis restricted to individuals for whom we had smoking information (N=13 579), the all-cancer RR was 15% higher in ever smokers versus never smokers, controlling for demographics and cohort membership (95% CI 1.02 to 1.30). Overall, >80% of lung cancer cases in this subpopulation (43/52) were reported in ever smokers. After controlling for smoking status and demographics, FDNY members had higher rates of all cancer and of prostate cancer compared with CFHS (RR=1.26, 95% CI 1.09 to 1.45 and RR=1.29, 95% CI 1.05 to 1.59, respectively.) Finally, when restricting our primary RR analyses to the FDNY population (N=10 786) we did not observe an association between increasing WTC exposure level and cancer rate (all-cancer RR=1.00, 95% CI 0.94 to 1.07).

DISCUSSION

On this, the 20th anniversary of 9/11, we set out to compare cancer rates in two firefighter cohorts, the WTC-exposed FDNY cohort and the non-WTC-exposed CFHS cohort, to each other and to rates among demographically similar men in the US population (SEER-21). The current investigation extended our previous study, which used CFHS data to estimate excess risk in WTC-exposed firefighters through 2009.²⁰ Consistent with the previous study, we continue to report evidence of excess thyroid and prostate cancer risk in the FDNY firefighters compared with CFHS firefighters. Unlike in the previous study, however, we now also report excess risk for all-cancers combined.

We documented modest excesses of cancer risk in the WTC-exposed FDNY cohort in relation to each comparison group. First, comparing FDNY to the CFHS cohort, we found a 13% excess risk for all cancers, largely driven by prostate and thyroid cancer. After correction for possible surveillance bias in FDNY by delaying diagnosis dates of certain cancer cases by 2 and then 5 years, excess risks were substantially reduced. Some proportion of the excess prostate cancer risk may be due to WTC exposure on top of usual firefighting risks, as some chemicals, like PCBs, commonly found at building sites including the WTC, are known endocrine disruptors, interfering with androgen metabolism.²⁷ This may have elevated the bioavailability of androgen, which could have been a factor in prostate cancer initiation. However, non-WTC-exposed firefighters may also be at risk of PCB exposure through usual firefighting exposures.² There is little biological evidence for increased thyroid cancer risk from exposures to environmental risk factors other than ionising radiation,²⁸ which was not detected at the WTC site. There is some evidence regarding thyroid cancer in pesticide-exposed workers²⁹⁻³¹; pesticides, however, were also not reported at the site.¹¹⁻¹³ Alternatively, high rates of some cancers, including thyroid and prostate cancers, could have resulted from non-biological factors like enrollment in screening programmes,³² especially WTC-related health programmes.^{17-20 33-35} Evidence for non-biological factors due to screening include younger median age at diagnosis among the FDNY cohort for all cancers and cancer subtypes compared with CFHS and SEER-21.²⁵

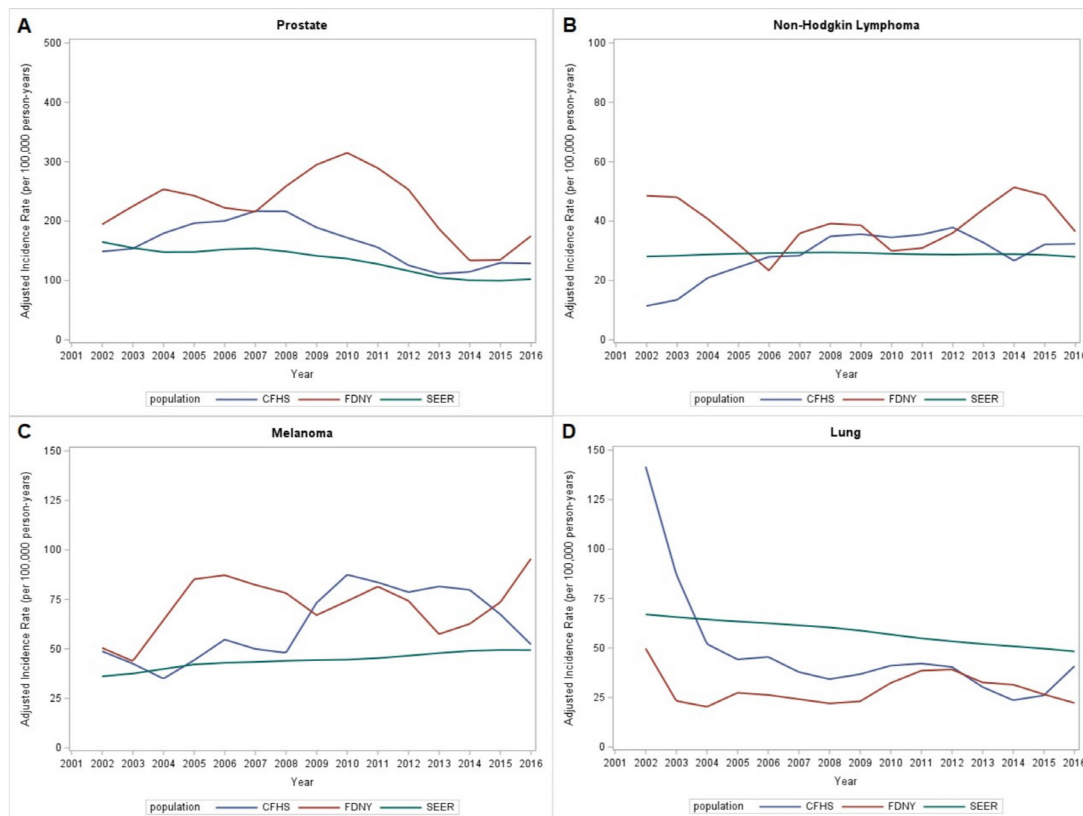


Figure 1 Incidence of site-specific cancers by calendar year in World Trade Center (WTC)-exposed firefighters, non-WTC-exposed firefighters and US males. Shown are the adjusted incidence rates over time of selected site-specific cancers, estimated by applying a locally weighted smoothing function to output from Poisson regression models of calendar year and cohort predicting cancer cases. (A) Shows estimated incidence rates of prostate cancer by calendar year in WTC-exposed Fire Department of the City of New York (red) and non-WTC-exposed Career Firefighter Health Study male firefighters (blue) versus in US males based on Surveillance, Epidemiology and End Results Program data (green). (B–D) Show estimated yearly incidence rates for non-Hodgkin lymphoma, melanoma and lung cancer, respectively, in the above three populations. Rates were adjusted for race/ethnicity and 5-year age group. CFHS, Career Firefighter Health Study; FDNY, Fire Department of the City of New York; SEER, Surveillance, Epidemiology, and End Results Program.

While the median ages at diagnosis were lowest for FDNY, CFHS median ages were also generally lower than SEER-21. Additional support for non-biological factors can be found in the staging data: cancers reported in SEER-21 were less likely to be classified as localised at time of diagnosis, although data from SEER-21 contained more unknown stages than either FDNY or CFHS. Occupational and WTC-related health programmes are designed for the early detection of cancer to minimise harm and improve survival. They clearly succeed in this mission, but may also overdiagnose occult, asymptomatic cancers.^{17–19 34 36}

Previous evidence from five non-WTC-exposed firefighter studies consistently supports excess risks of prostate cancer

and melanoma in firefighters.^{4 8–10 37} In our analyses, both firefighter cohorts had elevated rates of prostate cancer and melanoma relative to US males, although the SIRs for FDNY were higher. We proposed possible reasons for excess prostate cancer in firefighters above. As for melanoma, beyond UV exposure, melanoma has also been associated with PAH, PCB, aromatic hydrocarbons and other chemicals identified as present at the WTC^{11 13} and at non-WTC-related fires.³⁸

Compared with US males, the FDNY cohort also had significantly elevated SIRs for all cancer, non-Hodgkin lymphoma and thyroid cancer. The elevated all-cancer risk was driven by prostate cancer. After correcting for possible surveillance bias, we

Table 3A Standardised incidence ratios (SIRs) of cancers in male FDNY and CFHS firefighters versus US males,²⁴ 11 September 2001–31 December 2016

Site	FDNY case count	FDNY SIR	95% CI	CFHS case count	CFHS SIR	95% CI
All-cancer sites*	915	1.15	1.08 to 1.23	1002	1.05	0.98 to 1.12
Prostate	332	1.70	1.53 to 1.88	358	1.22	1.11 to 1.35
Lung	44	0.53	0.39 to 0.72	83	0.71	0.57 to 0.89
Kidney	39	0.93	0.67 to 1.28	55	1.19	0.90 to 1.56
Non-Hodgkin lymphoma	55	1.39	1.06 to 1.83	43	1.04	0.77 to 1.41
Melanoma (skin)	96	1.59	1.30 to 1.96	70	1.39	1.07 to 1.79
Thyroid	46	2.37	1.78 to 3.17	15	1.01	0.61 to 1.67

*All malignant cancers (multiple primaries), and in situ bladder cancers. CFHS, Career Firefighter Health Study; FDNY, Fire Department of the City of New York.

Table 3B Standardised incidence ratios (SIRs) of cancers in male FDNY firefighters versus US males.²⁴ 11 September 2001–31 December 2016, after 2-year adjustment for potential surveillance bias

Site	SIR*	95% CI
All-cancer sites†	1.09	1.02 to 1.16
Prostate	1.55	1.39 to 1.73
Lung	0.47	0.34 to 0.65
Kidney	0.85	0.61 to 1.19
Non-Hodgkin lymphoma	1.29	0.97 to 1.71
Melanoma (skin)	1.59	1.30 to 1.96
Thyroid	2.01	1.47 to 2.75

*Diagnosis dates of certain FDNY cancer cases (n=204) delayed by 2 years to account for potential surveillance bias.

†All malignant cancers (multiple primaries), and in situ bladder cancers. FDNY, Fire Department of the City of New York.

similarly observed elevated, although attenuated, SIRs. Despite our control efforts, surveillance bias concerns remain, as the US general population lacks access to comprehensive, no-copay healthcare like the WTC Health Program.³⁴

The comparison group of CFHS firefighters is based on a subset of the original NIOSH cohort of 29 992 career firefighters.⁴ Cancer data from the original cohort study accrued from 1985 to 2009, whereas our study began and ended later, covering the time period between 9/2001 and 12/2016. Thus, the current CFHS cohort markedly differs from the original study cohort, which likely explains differences observed. For example, compared with the US population, the earlier study found an all-cancer SIR of 1.09 (1.06–1.12) and lung cancer SIR of 1.12 (1.04–1.21),⁴ whereas we report CFHS SIRs of 1.05 (0.98–1.12) and 0.71 (0.57–0.89), respectively. We attribute these differences to our use of the truncated cohort (those active on 9/11), which resulted in a younger and more recently employed group.

Notably, both firefighter groups had lower than expected rates of lung cancer when compared with US males, probably due to lower current smoking rates: 3.5%, 6.6% and 15.0% in the FDNY, CFHS, and US male populations, respectively.³⁹ Among firefighters with smoking data, ever smokers demonstrated higher overall cancer rates than never smokers. Differences in incident cancers may also be attributed to a previously described secular trend of declining cancer rates in firefighters hired since

Table 4 Adjusted relative rates (RR) and 2-year surveillance bias-adjusted RR of cancers in WTC-exposed male FDNY firefighters versus male CFHS firefighters, 11 September 2001–31 December 2016

Site	Adj. RR (95% CI)*	Surveillance bias Adj. RR (95% CI)*†
All-cancer sites‡	1.13 (1.02 to 1.25)	1.07 (0.96 to 1.18)
Prostate	1.39 (1.19 to 1.63)	1.28 (1.09 to 1.51)
Lung	0.87 (0.57 to 1.33)	0.77 (0.50 to 1.19)
Kidney	0.82 (0.52 to 1.30)	0.75 (0.47 to 1.20)
Non-Hodgkin lymphoma	1.26 (0.80 to 2.00)	1.21 (0.75 to 1.94)
Melanoma (skin)§	1.12 (0.80 to 1.57)	N/A
Thyroid	2.53 (1.37 to 4.70)	2.11 (1.14 to 3.90)

*Regression models adjusted for age on 11 September 2001 and race/ethnicity.

†Diagnosis dates of some FDNY cancer cases (N=204) delayed by 2 years to account for potential surveillance bias.

‡All malignant cancers (multiple primaries), and in situ bladder cancers.

§Analysis restricted to white males (N=16 238) due to low case counts in other race/ethnic groups.

CFHS, Career Firefighter Health Study; FDNY, Fire Department of the City of New York; NA, not available; WTC, World Trade Center.

1970, generally attributed to better personal protective equipment (PPE).¹⁰

Regular assessment of cancer risk in firefighters remains imperative because firefighting continues to be a common career and volunteer activity. Building contents (synthetics and plastics), fire suppression materials and PPE change over time and over geographic regions—facts which may explain inconsistent cancer results from previous studies.^{3–10 37} For example, The International Agency for Research on Cancer has classified perfluoroalkyl, a chemical used during suppression activities, as possibly carcinogenic to humans,⁴⁰ but has not yet fully evaluated this possibility. Differences in rates should also be considered in the context of observed behavioural changes (eg, diet, smoking, alcohol use and sunblock use) and in diagnostic testing (access and technology).

Finally, assessment of cancer risk in FDNY firefighters who worked at the WTC site remains complex; these firefighters were subject to carcinogenic exposures, while also enduring enormous physical and mental burdens related to the attacks. Examination of longitudinal FDNY, CFHS and SEER-21 data by calendar year shows elevated cancer rates in the FDNY cohort, particularly prostate cancer between 2008 and 2012. While these increased rates may be driven by higher participation in medical monitoring after 2007, they could also reflect prostate cancer latency.⁴¹ Evidence is slowly accruing about cancer and other long latency illnesses in relation to WTC exposure, although much remains to be determined. Molecular epidemiological studies of biomarkers may provide better understanding of chronic disease development in firefighters, both WTC-exposed and non-WTC-exposed.

Strengths of this study include the use of two different comparison groups to assess excess cancer risk in the WTC-exposed FDNY cohort, an achievement that no other group that we know of has been able to claim. Thus, we were able to report WTC-exposed firefighter cancer risk versus risk in other firefighters and WTC-exposed risk versus risk in demographically similar US males. In addition, we restricted analyses to firefighters actively employed on and after 9/11 not only to allow for WTC exposure comparison, but also so that findings would be relevant to firefighting in modern structures and PPE. Additional strengths include the lengthy follow-up time for each cohort (~15 years), and that smoking history was accounted for in those with available information.

Weaknesses include limitations common to all observational studies, that confounding may be insufficiently controlled in analyses, especially as we had more data from FDNY than from CFHS. For example, smoking history was available for nearly all FDNY firefighters but only for the 32% of CFHS firefighters who completed the CFHS survey. Similarly, we had incomplete information on WTC exposure in the CFHS cohort. If CFHS firefighters who volunteered at the WTC site had exposures similar to those of active duty FDNY firefighters, misclassifying these CFHS members as non-WTC-exposed would have biased our estimated cancer RRs towards the null. Another limitation was that correcting for surveillance bias by lagging FDNY diagnosis dates of certain cancers was an imperfect way to mitigate screening effects, as the 2-year lag time may be insufficient. We found that applying a 5-year delay to the diagnosis dates mostly removed the excess cancer incidence observed in WTC-exposed FDNY firefighters. Asymptomatic cases in the US population, and to a lesser extent in the CFHS cohort, may go undiagnosed without regular physical exams or adherence to screening programmes. However, as cancer screening guidelines such as those from the US Preventive Service Task Force (USPSTF)⁴² and

the American Cancer Society⁴³ become more widely accepted, these differences should become less pronounced. For firefighters, these cancer screening guidelines have been recommended since 1997 by the International Association of Firefighters.⁴⁴ Although the fire departments represented in the CFHS cohort have had far less funding for cancer screening than FDNY, adjustment for surveillance bias only in the FDNY cohort could have led to an overestimation of this bias if the other departments' screening practices are similar. Alternatively, if the fire departments in CFHS followed the 2012 USPSTF prostate cancer screening guidelines recommending against PSA screening between 2012 and 2016,⁴⁵ while FDNY continued to carry out PSA screening throughout the follow-up period, the estimated RR for prostate cancer may have been biased away from the null. In a future study, we plan to further address potential surveillance bias by examining cause-specific mortality in the FDNY and CFHS populations, particularly among those diagnosed with cancer, as our current study found that the proportion of firefighters alive at the end of follow-up was greater in FDNY than in CFHS.

Clearer understanding of the WTC-related cancer risk for firefighters requires progress in at least two directions: additional years of follow-up to allow for the suspected long latency of some solid tumours; and, modelling studies (laboratory or animal based) to identify and track workplace exposures in WTC-exposed and non-WTC-exposed firefighters. These steps may aid our understanding of the complex relationships between WTC exposure, firefighting and cancer.

Author affiliations

¹Division of Epidemiology, Department of Epidemiology and Population Health, Montefiore Medical Center and Albert Einstein College of Medicine, Bronx, New York, USA

²Bureau of Health Services, Fire Department of the City of New York, Brooklyn, New York, USA

³Pulmonary Medicine Division, Department of Medicine, Montefiore Medical Center and Albert Einstein College of Medicine, Bronx, New York, USA

⁴Division of Biostatistics, Department of Epidemiology and Population Health, Montefiore Medical Center and Albert Einstein College of Medicine, Bronx, New York, USA

⁵World Trade Center Health Program, Centers for Disease Control and Prevention (CDC), National Institute for Occupational Safety and Health (NIOSH), Cincinnati, Ohio, USA

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ORCID iDs

Mayris P Webber <http://orcid.org/0000-0002-8322-4573>

Ankura Singh <http://orcid.org/0000-0002-8495-4850>

Rachel Zeig-Owens <http://orcid.org/0000-0002-8679-2306>

Joke Salako <http://orcid.org/0000-0003-2417-9810>

Molly Skerker <http://orcid.org/0000-0002-7511-610X>

Charles B Hall <http://orcid.org/0000-0001-9982-8120>

David G Goldfarb <http://orcid.org/0000-0002-8679-2306>
 Robert D Daniels <http://orcid.org/0000-0002-3915-7991>
 David J Prezant <http://orcid.org/0000-0001-9562-0330>

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