



War-related stressors are associated with asthma risk among older Kuwaitis following the 1990 Iraqi invasion and occupation

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ABSTRACT

Background Contemporary warfare involving civilian populations is a growing public health concern. In addition to the psychological impact, war-related trauma may result in physiological alterations and even broader health effects. Associations were examined between war-related stressors and incident asthma in elderly Kuwaiti civilians following the Iraqi invasion.

Methods A random sample of all Kuwaiti nationals aged 50—69 years on the day prior to the invasion were identified. Among the 7873 meeting eligibility criteria, 5567 (71%) agreed to participate and 5028 completed the questionnaire (91% of those eligible). Of these, 3759 were in Kuwait during the invasion, of whom 2294 were alive at follow-up. After exclusions for prevalent asthma or missingness on covariates, 2066 were available for analysis. War-related experiences were summarised into a continuous score using Rasch modelling. Relative Cox proportional hazard rates (HR) were calculated for asthma adjusting for covariates.

Results Over 13 years of follow-up, physician-diagnosed asthma was reported by 66/996 (6.6%) men and 104/1070 (9.7%) women. In models adjusted for gender, socioeconomic status, smoking, BMI, and air pollution related to burning oil fires, those reporting highest stress exposure were more than twice as likely to report asthma (HR 2.3, 95% CI 1.3, 3.9) compared to civilians reporting no stressors. Experiences were more salient when anchored to fear for loss of life.

Conclusions War-related trauma is associated with increased asthma risk in these elderly civilians. Although prior research has documented the significant and persistent psychological toll of war, these findings implicate even broader health effects.

Contemporary warfare and political violence increasingly involves civilian populations and is a growing public health concern. Historically, epidemiological studies linking war-related exposures to long-term morbidity have been challenging. Often it is difficult to examine direct linkages of the original war-related experiences, given that most populations in this circumstance experience prolonged periods of armed conflict. Moreover, after the conflict, most endure prolonged economic crisis, lack of stable social systems and ongoing deprivation or displacement, which may continue to impact health. ²

The Kuwait experience provides a unique research opportunity for many reasons. The invasion of Kuwait in 1990 was followed by a 7-month-long Iraqi occupation. Although the period between

the invasion and liberation was relatively short compared to most conflicts, and many Kuwaiti civilians were out of the country at the time of the invasion and occupation, many remained and endured harsh conditions and fear of harm.³ At least 15 000 were arrested and approximately 1000 were killed. After liberation, Kuwaitis did not live in exile for long periods and the physical and social infrastructure was restored in a comparatively short time.⁴

Although epidemiologists are beginning to examine the long-term health effects of such experiences in population-based studies, the literature has largely focused on persistent psychological co-morbidities including depression, anxiety, and post-traumatic stress disorder (PTSD).⁵ Although such emotional states may represent adaptive functioning in the face of trauma, they may exact a toll on mental as well as physical health for years to come.

In addition to having significant psychological impact, war-related trauma may result in physiological alterations that lead to even broader health effects, a notion grounded in traumatic stress theory. ¹⁷ In general, stressors are thought to influence the pathogenesis of disease by causing negative affective states (eg anxiety, depression, and PTSD), which in turn exert effects on biological processes or behavioural patterns that influence disease risk. ⁸ When environmental demands are perceived to be threatening and exceed one's abilities to cope, or when one feels that events are unpredictable or that one has no control, individuals experience stress

Many aspects of the Iraqi invasion and subsequent occupation fit this paradigm. Traumatic experiences, including living in fear, separation exposure family, shelling to combat, witnessing or being a victim of violence or torture, displacement, witnessing arrests of others, and sexual assault have been documented.^{4 6} Arbitrary acts of oppression or violence by Iraqi soldiers had Kuwaiti civilians living in constant fear of potential harm, torture, sexual assault and other atrocities.³ In the context of a natural disaster, the persistence of psychological distress symptoms has been related to collapse of community infrastructure. Destruction of the physical infrastructure may have further predisposed this population to chronic psychological trauma. 4 Thus, the traumatic stress paradigm linking psychological stress and affective states to disruption of key physiological mechanisms, and ultimately to disease expression, provides a framework for plausible mechanisms linking war-related trauma, psychological functioning and physical health.⁷

Asthma provides a disease paradigm for examining this posited link. Efforts to understand the role of stress in asthma expression have expanded in the context of an increased understanding of the overlap between the neurobiology of stress and asthma pathophysiology. 10 Psychological stressors and negative emotional responses disturb the regulation of the sympathetic and adrenomedullary (SAM) system and the hypothalamic-pituitary-adrenocortical (HPA) axis - physiological systems may operate at higher or lower levels than during normal homeostasis. The detrimental cost of such accommodation has been conceptualised as allostatic load (ie chronic underor over-activity of the allostatic system). 11 Hormones and neuropeptides released into the circulation when individuals experience stress regulate both immune-mediated and neurogenic inflammatory processes. Altered neuroimmune responses may influence the expression of immune-mediated or inflammatory disorders in particular, including asthma. Moreover, the authors' group has previously demonstrated a link between asthma and caregiving stress¹² and other traumatic stressors (ie domestic violence). 13 14

Although an individual's reaction to trauma is multifaceted and associated with a range of psychological sequelae, in the context of extreme events (eg war) distress symptoms associated with PTSD (eg chronic re-experiencing of the trauma) have been most widely studied.⁵ Because war-related experiences are outside the range of usual human experience and almost universally distressful, they are more likely to lead to PTSD. In turn, PTSD has been linked to disruption of stress pathways even several years following the experienced trauma, 15 as well as being linked prospectively to increased mortality 16 and chronic disease risk¹⁷ in veterans. Amongst Cambodian¹⁸ and Bosnian¹⁹ refugees, researchers have shown a dose-response (cumulative) relationship in which incremental exposure to a variety of adverse events was associated with increasing risk of PTSD.¹⁸ Moreover, the significance and meanings underlying trauma experiences may be more important than discrete events in determining psychological functioning and other adverse health risk. Specifically, fear of loss of life in the context of trauma has emerged as a particularly salient precipitant of subsequent morbidity. 20 21

The relationship between war-related trauma and physical health has not been examined in non-refugee civilian populations experiencing such conflict. Here, associations were examined between war-related stressors and incident asthma in older adult Kuwaitis following the invasion and occupation. It was hypothesised that among Kuwaiti civilians who remained in the country, those experiencing greater cumulative stressors would be at increased risk for asthma compared to those with lesser exposure. Finally, fear for loss of life anchored to war experiences was considered as a modifier.

METHODS

The Public Authority for Civil Information provided the base population of all Kuwaiti nationals (ages 50–69 years) alive on the day before invasion (1 August 1990). In order to get a random sample who were either in or out of Kuwait during the invasion, subjects were matched to Ministry of Finance records of Kuwaitis receiving a stipend (~US \$1500) for remaining in Kuwait during the entire occupation. A random representative sample of 7873 Kuwaitis was selected. Subjects were initially contacted by phone and then written consent was obtained

during face-to-face interviews conducted in homes between December 2003 and January 2005. The questionnaire confirmed location (in or out of Kuwait) and ascertained information on war-related experiences, sociodemographics, smoking, height and weight, and asthma. Of 7873 subjects, 5567 (71%) agreed to participate; 5028 successfully completed the survey. Analyses were restricted to those in Kuwait during the occupation (3759/5028) and therefore at risk for exposure to trauma; of these, 2294 were alive at the time of the survey. Finally, 228 were excluded because of prevalent asthma or missing values on covariates, leaving 2066 for analyses.

War-related stressors

Assessing war-related trauma in epidemiological research is challenging. At the time of this survey, there were no empirically developed measures of war-related events in civilian, non-refugee populations. Determining which events are most relevant to risk of PTSD and other health sequelae is complex, given that civilians exposed to war typically experience multiple traumas, some of which are relatively universal and some that may be unique to a particular setting. The present survey was based on the Harvard Trauma Questionnaire, developed from refugees in clinical samples. This was supplemented through interviews with individuals from the relevant population to capture events specific to the Kuwaiti experience.

Another challenge is creating a predictor that is not directly measured, but is ascertained through multiple items related to the exposure (eg war-related traumatic events).²⁴ An approach previously applied to quantifying exposure to community violence was adapted.²⁵ Rasch modelling, a form of latent variable analysis, summarises dichotomous responses to the multi-item questionnaire into a continuous score on the logit scale. A continuous measure is obtained by modelling the conditional probability of responding "yes" to each item given the extremity or severity of each question. For example, when a subject with low exposure is asked whether a severe form of trauma was experienced, the expected probability of endorsement is low. Latent variable models also allow us to account for information that may modify the impact of the event [eg frequency of events and characteristics of the perpetrator and victim (stranger vs non-stranger)].²⁵ Subjects reported whether they directly experienced and/or had witnessed various events. The Rasch model assigns weights for the severity of the events experienced assuming events endorsed less frequently are more severe. In such a scale with both witness and victimisation items, witnessed events would be considered less severe as they were reported more frequently than victimisation. To accommodate this, a model using both witness and victimization items was implemented, creating two scales but taking into account the existing correlation between the scales. 26 A War-Related Stressor Score (WRSS) was defined as the average of the two continuous scales, that is, the latent variable of exposure to both witnessed events and victimization. These models were implemented using logistic non-linear mixed models (proc NLMIXED) in SAS 9.0 (SAS Institute, Cary, NC, USA).

Asthma

Asthma was determined by the question "Have you ever been told by a doctor that you have asthma?", with a follow-up to determine the year they received the diagnosis, a standard approach with demonstrated reliability.²⁷ Incident asthma was defined as being diagnosed following the date of liberation (27 February 1991).

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Covariates

Analyses were adjusted for age, gender, and social status including education and literacy. Age on the date of invasion was categorised as 50-54, 55-59, 60-64, and 65-69 years. Challenges to defining social status in this older age cohort²⁸ and in the Kuwaiti culture were considered.²⁹ For example, Arab elders often rely on assistance solely from family members and would not be expected to report income of their own. 30 Data were available on education level and literacy, which have been associated with differential health risks in the Gulf region.³¹ Education was dichotomised as never attending school or attending school at any level. Smoking was defined as "never", "former", or "current". Body mass index (BMI) was calculated based on self-reported weight and height (weight in kilograms divided by the square of height in metres) and entered as a continuous measure. Self-reported weight and height are acceptably correlated with measured weight and height in the elderly. 32 Air pollution from burning oil fires, an environmental contaminant from the war potentially related to asthma, 33 34 was also controlled. Daily exposure to oil fire fine particulate matter (2.5 μ m, PM_{2.5}) (modelled to account for area of residence, climate factors, etc, using CalPUFF dispersion models³⁵) was averaged over the time subjects reported being in Kuwait during the invasion up to 6 November 1991 when the last fire was extinguished, 36 and entered as a continuous indicator [mean oil fire PM_{2.5} (µg/m³)]. And finally, it was considered whether an individual reported fear for loss of life in the context of reported traumas as a modifier of the relationship between war-related experiences and asthma.

Analysis

Crude morbidity rates were calculated as the number of subjects reporting doctor-diagnosed asthma divided by the number within each WRSS exposure level. To assess for a dose—response, the trauma score was categorised as no trauma compared to the Rasch scale divided into tertiles [low (–1.0 to –0.2), intermediate (–0.2 to 0.7) and high (0.7 to 3.8)]. Time to incidence was calculated as the reported year of diagnosis (or year of interview for those without asthma) minus date of liberation. Relative Cox proportional hazard rates (HR) were calculated adjusting for covariates. Asthma risk was considered based on war-related stressor level and whether subjects experienced fear for loss of life related to events during the Iraqi occupation, which may potentiate post-trauma morbidity. Analyses were conducted using SAS version 9 (SAS Institute; Cary, NC, USA).

RESULTS

Mean age (58.7±5.6 and 58.2±5.5, men and women respectively) and BMI (26.7±4.7 and 28.8±5.8, men and women respectively) were similar across gender. Significant differences were seen for literacy (72.2% men; 29.8% women), any education (54.1% men; 16.0% women) and smoking status (non-smoker 41.6% and 84.3%; former smoker 37.2% and 9.0%; and current smoker 21.3% and 6.7%, men and women respectively). Education and literacy levels were low in this older sample, having historically had less opportunity for education. 37 Table 1 details the distribution of covariates relative to asthma diagnosis. In addition, BMI was higher among those with asthma (28.7 ± 6.3) compared to non-asthmatics (27.7 ± 5.3) (p = 0.02); mean oil fire PM_{2.5} was not significantly different between asthmatics and non-asthmatics [55.8±15.8 vs 50.4 ± 15.9 µg/m³ respectively (p=0.8)]. Table 2 depicts frequencies of war-related stressors included in the Rasch model; 368 reported no trauma [154 men (16%), 214 women (20%)]. More than half of the men (68%) and women (59%) feared for loss of life. The WRSS (figure 1) can be interpreted in a relative sense (eg -0.5 represents lower exposure than a score of 2.0). Among 413 with a score <-0.75, only one reported assault, three were arrested and 28 reported having their house searched; none witnessed torture, rape or executions. In contrast, among 517 with a score >0.85, 12% were arrested, 16% assaulted and 57% witnessed torture.

Incident asthma was reported by 66/996 (6.6%) men and 104/1070 (9.7%) women. In univariate analyses, there was a direct relationship between higher level exposure on the WRSS (p for trend 0.0003) and asthma risk (data not shown), with the highest tertile predicting a doubling of asthma risk (HR 2.0; 95% CI 1.2 to 3.2). Adjusting for covariates did not significantly change this relationship (HR 2.3; 95% CI 1.3 to 3.9) (table 3).

The elevated risk for asthma was most evident in the subgroup reporting fear for loss of life associated with the highest WRSS level (table 4), albeit a formal test of an interaction was not statistically significant (p = 0.12), likely in part due to small cell sizes.

DISCUSSION

War-related stressors were associated with elevated risk of incident asthma in elderly Kuwaiti civilians exposed to the 1990 Iraqi invasion adjusting for other war-related environmental contaminants (air pollution from the burning oil fires), social status, BMI and smoking. Although prior research has documented the significant and persistent psychological toll of such experiences, these findings implicate even broader health effects. The suggested dose—response relationship parallels studies demonstrating cumulative effects of war-related trauma on psychological health, eg incremental increases in exposure to war-related events among Bosnian refugees was associated with increasing risk of PTSD. 19

There was also a suggestion that trauma exposure was more salient when it was anchored to fear for loss of life. The significance and meanings underlying trauma experiences may be more important than discrete events in determining PTSD and other adverse health risk. These studies show that life threat is salient to triggering PTSD in Western cultures, a finding recently corroborated in populations with more diverse religious and cultural backgrounds (eg Bosnian Muslim refugees). Dahl and colleagues found a higher rate of PTSD among Bosnian women who had been threatened with death than those with general war experiences. In the present data, the suggestion that experiences were more salient when anchored to fear for loss of life is consistent with the higher rates of psychological morbidity reported among these Bosnian women threatened with death.

Future studies examining the long-term effects of war-related stressors on civilians should be designed to more formally test whether these relationships are mediated through adverse psychological sequelae (eg PTSD) and consequential physiological disruption. Research incorporating such multimodal assessments of the response to war-related experiences will be most informative. For example, prospective studies should examine alterations of stress axes (eg HPA axis, SAM system) linked to PTSD and other psychological sequelae resulting from war experiences that, in turn, play a role in disease expression. Future research should also consider differential gender effects — female gender is a risk factor for PTSD in general and studies of war-related trauma report higher levels of distress in women

 Table 1
 Distribution of covariates based on asthma diagnosis

		New asthma	
Covariates	Total	N (%)	p Value*
War-Related Trauma Score			
No reported trauma	368	23 (6.3)	0.001
Low	573	35 (6.1)	
Medium	571	45 (7.9)	
High	554	67 (12.1)	
Sex			
Male	996	66 (6.6)	0.01
Female	1070	104 (9.7)	
Age			
50-54	679	51 (7.5)	0.73
55—59	574	53 (9.2)	
60-64	467	37 (7.9)	
65-69	346	29 (8.4)	
Education			
No	1356	116 (8.6)	0.46
Yes	710	54 (7.6)	
Literacy			
No	1028	101 (9.8)	0.009
Yes	1038	69 (6.7)	
Smoking			
Non-smoker	1316	107 (8.1)	0.86
Ex-smoker	466	41 (8.8)	
Current	284	22 (7.8)	
Fear loss of life			
No	778	56 (7.2)	0.18
Yes	1284	114 (8.9)	

^{*}Denotes differences based on χ^2 p value.

compared to men, as well as differences related to contextual factors (eg the impact of fear for loss of life). $^{15\ 19}$

Although the study has strengths given the large sample and the unique circumstances related to the invasion and occupation, some limitations are also noted. Self-reported physiciandiagnosed asthma was used, which has been validated in other research albeit not in Kuwaiti populations.⁴² The prevalence of asthma in Kuwait as well as other Persian Gulf states appears to be similar to that in industrialised or Western countries.⁴³ ⁴⁴ A number of factors related to asthma diagnosis in this older

Table 2 War-related experiences*

	AII (N = 2066)	Men $(n = 996)$	Women (n = 1070) N (%)
War-related experiences	n (%)	n (%)	
Witnessed events			
Health threat due to lack of financial/medical support	471 (22.8)	250 (25.1)	221 (20.6)
Family arrested	661 (32)	313 (31.4)	348 (32.5)
Family rape	21 (1.0)	9 (0.9)	12 (1.1)
Family shot	244 (11.8)	106 (10.6)	138 (12.9)
Family torture	217 (10.5)	97 (9.7)	120 (11.2)
Other rape*	51 (2.5)	32 (3.2)	19 (1.8)
Other torture*	283 (13.7)	171 (17.2)	112 (11)
Other assault*	309 (15.0)	196 (19.7)	113 (11)
Execution*	241 (11.7)	157 (15.8)	84 (7.9)
House searched*	1242 (60.1)	661 (66.4)	581 (54.3)
House loss/destroyed*	532 (25.8)	279 (28)	253 (23.6)
Direct victimisation			
Assaulted at checkpoint*	57 (2.8)	42 (4.2)	15 (1.4)
Assaulted at demonstration	14 (0.7)	10 (1.0)	4 (0.4)
Assaulted somewhere else*	64 (3.1)	49 (5.0)	15 (1.4)
Tortured*	32 (1.5)	27 (0.7)	5 (0.5)
Sexually assaulted or raped	9 (0.4)	4 (0.4)	5 (0.5)
House searched	1142 (55.3)	564 (56.6)	578 (54)
House loss/destroyed*	328 (15.9)	141 (14.2)	187 (17.5)
Arrested*	86 (4.2)	81 (8.1)	5 (0.5)
In hiding*	288 (13.9)	121 (12.1)	167 (15.6)

^{*}Denotes significant differences between men and women based on $\chi^2\,p$ value $<\!0.05.$

Research report

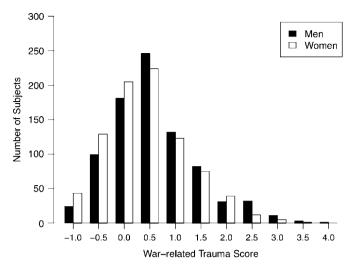


Figure 1 Rasch scores on the logit scale for those reporting any warrelated stressors summarised in table 2.

Kuwaiti adult population are corroborated by studies from the West^{45 46} as well as the region. ⁴⁷ It was found that the number of new cases of asthma was higher among women consistent with existing prevalence and incidence studies, ⁴⁷ as well as a suggestion of increased risk for former smokers. ⁴⁸ In this population of older adults there is also the possibility of overreporting of asthma due to confusion with reporting chronic obstructive pulmonary disease (eg emphysema or chronic bronchitis). This is only biased to the extent that such overreporting is correlated with our main exposure, war-related trauma, which seems unlikely. It also seems unlikely that associations might represent trauma leading to acute exacerbations of existing undiagnosed subclinical asthma, as the majority (81%) of incident cases were diagnosed a year or more

Table 3 War-Related Stressor Score and asthma incidence: adjusted model

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	HR (95% CI)*			
War-Related Stressor Score				
No reported stressors	Ref			
Low	1.0 (0.6 to 1.8)			
Intermediate	1.4 (0.8 to 2.4)			
High	2.3 (1.3 to 3.9)			
Gender				
Male	Ref			
Female	1.4 (1.0 to 2.4)			
Age at invasion				
50-54	Ref			
55-59	1.3 (0.9 to 2.1)			
60-64	1.1 (0.7 to 1.8)			
65-69	1.4 (0.9 to 2.4)			
Education				
No	Ref			
Yes	1.9 (1.1 to 3.3)			
Literacy				
Yes	Ref			
No	1.9 (1.1 to 3.2)			
Smoking				
Non-smoker	Ref			
Ex-smoker	1.3 (0.9 to 2.0)			
Current	1.1 (0.7 to 1.9)			
BMI (per SD change)	1.0 (0.9 to 1.2)			
Mean oil fire PM _{2.5}	1.0 (0.99 to 1.0)			

^{*}Hazard rate (95% confidence interval).

Table 4 Asthma risk by stressor level and fear for loss of life*

	N = 2066 HR (95% CI)†
Fear loss of life	
No reported stressors	0.7 (0.3 to 1.7)
Low	0.7 (0.4 to 1.5)
Intermediate	1.4 (0.7 to 2.7
High	2.0 (1.1 to 3.6
No fear loss of life	
No reported stressors	Ref (—)
Low	1.1 (0.5 to 2.2
Intermediate	0.8 (0.4 to 1.7
High	1.3 (0.6 to 2.8
	·

^{*}Adjusted for age, gender, education, literacy, smoking, BMI and mean oil fire PM_{2.5}. †Hazard rate and 95% confidence interval.

after the invasion. Also, socioeconomic status may be related to access to healthcare, differential exposure to war trauma, type of housing and other environmental factors that also may contribute to asthma risk. Although adjusting for education and literacy did not change the observed associations, characterisation of socioeconomic status in this sample is notably complex, 28 29 and residual confounding may remain. Another possible explanation for the findings is that individuals with psychological morbidity related to trauma experiences may be more likely to seek medical attention or over-report medical complaints. 49 Follow-up studies should systematically assess psychological co-morbidities and incorporate objective measures of the asthma outcome (eg airway responsiveness, lung function, asthma medication use, immunophenotyping). The study may also have been strengthened through the inclusion of personal monitoring of air pollution exposures rather than prediction modelling, albeit this information was not available in this dataset.

What is already known on this subject

- ► Contemporary warfare increasingly involving civilians is a growing public health concern.
- Although epidemiologists are beginning to examine the longterm health effects of war-related experiences, research to date has largely focused on persistent psychological sequelae or on cardiovascular disease.
- War-related trauma may result in physiological disruptions that may result in even broader health effects.

What this study adds

- ► This is the first study to find associations between war-related trauma and chronic respiratory disease, namely asthma risk among elderly civilians exposed to war.
- Research linking war-related stressors to long-term physical health expands our understanding of the full scope of the health effects of war.
- Recognising and intervening around psychological difficulties related to war experiences early in the course may diminish the risk of developing non-psychiatric sequelae in the long run.

In summary, evidence was found that increased war-related trauma is associated with elevated risk of new-onset asthma in elderly Kuwaiti civilians exposed to the Iraqi invasion and occupation. Although prior research has documented the significant and persistent psychological toll of such trauma, these findings implicate even broader health effects. Recognising and intervening around psychological co-morbidities related to trauma early in the course may diminish the risk of developing non-psychiatric sequelae related to such experiences, thus further research in this area is warranted.⁴⁹

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Dedication We dedicate this paper to the memory of our dear colleague, Professor Jaafar Behbehani of the Kuwait University Faculty of Medicine, who was the inspiration for this work. Jaafar's vision; his personal and professional understanding of the impacts of war on the physical and mental well-being of civilian populations; his untiring dedication and devotion to develop the scientific evidence needed to inform this matter; and his personal elegance and warmth made this work possible. We owe him a great debt and hope that the publication of this paper will in some small way ensure that his contributions to science are appreciated.

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Competing interests None.

Ethics approval Study procedures were approved by the Kuwait University Faculty of Medicine and Harvard School of Public Human Studies Committees.

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