

Risk Factors for Depression Among Civilians After the 9/11 World Trade Center Terrorist Attacks: A Systematic Review and Meta-Analysis

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Abstract

Introduction: The development of depressive symptoms among the population of civilians who were not directly involved in recovery or rescue efforts following the 9/11 World Trade Center (WTC) terrorist attacks is not comprehensively understood. We performed a meta-analysis that examined the associations between multiple risk factors and depressive symptoms after the 9/11 WTC terrorist attacks in New York City among civilians including survivors, residents, and passersby.

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Methods: PubMed, Google Scholar, and the Cochrane Library were searched from September, 2001 through July, 2016. Reviewers identified eligible studies and synthesized odds ratios (ORs) using a random-effects model.

Results: The meta-analysis included findings from 7 studies (29,930 total subjects). After adjusting for multiple comparisons, depressive symptoms were significantly associated with minority race/ethnicity (OR, 1.40; 99.5% Confidence Interval [CI], 1.04 to 1.88), lower income level (OR, 1.25; 99.5% CI, 1.09 to 1.43), post-9/11 social isolation (OR, 1.68; 99.5% CI, 1.13 to 2.49), post-9/11 change in employment (OR, 2.06; 99.5% CI, 1.30 to 3.26), not being married post-9/11 (OR, 1.59; 99.5% CI, 1.18 to 2.15), and knowing someone injured or killed (OR, 2.02; 99.5% CI, 1.42 to 2.89). Depressive symptoms were not significantly associated with greater age (OR, 0.86; 99.5% CI, 0.70 to 1.05), no college degree (OR, 1.32; 99.5% CI, 0.96 to 1.83), female sex (OR, 1.24; 99.5% CI, 0.98 to 1.59), or direct exposure to WTC related traumatic events (OR, 1.26; 99.5% CI, 0.69 to 2.30).

Discussion: Findings from this study suggest that lack of post-disaster social capital was most strongly associated with depressive symptoms among the civilian population after the 9/11 WTC terrorist attacks, followed by bereavement and lower socioeconomic status. These risk factors should be identified among civilians in future disaster response efforts.

Introduction

The mental health consequences of the September 11, 2001 (9/11) World Trade Center (WTC) terrorist attacks in New York City (NYC) have been the focus of a substantial number of research endeavors over the past 15 years.¹ A majority of this research has documented the etiology, prevalence, treatment, and risk factors for posttraumatic stress disorder (PTSD) in particular, as PTSD is reported to be the most common post-disaster associated condition.² According to the Fifth Edition of the Diagnostic and Statistical Manual of Mental Disorders (DSM-5), PTSD is a conditional disorder that can develop among individuals after a qualifying trauma exposure.³ Qualifying trauma exposure must result from directly experiencing a traumatic event, being an eyewitness to trauma as it occurred to others, learning that a close associate suffered from a traumatic event, or experiencing repeated or extreme exposure to aversive details of a traumatic event.³

Probable depression is another commonly studied post-disaster mental health outcome. Probable depression refers to a positive screen on a depression symptom screening instrument, such as the Patient Health Questionnaire.^{4,5} PTSD and probable depression among trauma-exposed groups of the WTC terrorist attacks have been thoroughly described.^{6,7,8,9,10} Examples of frequently studied trauma exposed groups include firefighters, police, emergency medical technicians, first responders, and recovery and cleanup workers.

In contrast to trauma-exposed groups, the general civilian population of NYC, comprised of survivors, NYC residents, people working in the area, and passers-by on the day of the attacks, is an example of a mixed-exposure group. After the WTC terrorist attacks, members of the civilian population suffered from varying levels of exposure to trauma. Many of these exposures may have not met DSM-5 trauma exposure criteria.^{11,12} The general civilian population is not always included in studies of prevalence estimates and risk factors for post-disaster PTSD and probable depression because disaster mental health research is typically interested in understanding mental disorders in relation to trauma exposure.¹³ In particular, comprehensive analyses of post-disaster probable depression among civilians are lacking in the literature.

The unpredictable nature of terrorist attacks have introduced new definitions of affected populations in disaster mental health research, as the purpose of terrorism is to invoke fear and anxiety among civilians in general.¹⁴ Moreover, probable depression is not dependent on qualifying trauma exposure, so the population susceptible to probable depression after a disaster such as a terrorist attack is larger than the population susceptible to PTSD.⁵ Carefully identifying risk factors for probable depression among the civilian population that were associated with the 9/11 WTC terrorist attacks may better inform future disaster preparation efforts.

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Over the past 15 years, a number of studies have screened for probable depression among affected populations and stratified their analysis of trauma-exposed groups and civilians, or sampled cohorts of civilians specifically.^{15,16,17,18,19,20,21} The aim of this study was to synthesize the results from research on 9/11-affected survivors, residents, and passers-by, summarize the influence of probable depression among these cohorts, and evaluate the associations between probable depression and various risk factors.

Methods

Search Strategy

We conducted a meta-analysis following the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) Statement.²² PubMed, Google Scholar, and the Cochrane Library were searched without language restriction from September, 2001 through July, 2016. Search terms included "World Trade Center", "WTC", "World Trade Center Disaster", "WTCD", "September 11", "9/11", "Depression", "Major Depression", "Probable Depression", and "Mental Health" in various relevant combinations. Both published and unpublished sources of data were considered. References of studies and review articles were also manually searched to yield additional studies not found through the original search.

Study Inclusion/Exclusion Criteria

Two reviewers (A.C., and S.B.) screened the search results for eligibility. Disagreements between reviewers regarding the inclusion or exclusion of a study were resolved by a third reviewer (J.P.). Studies were eligible based on the following inclusion criteria: 1) original research article; 2) focused on the effects of the WTC terrorist attacks; 3) focused on adult populations based on age at the time of interview; 4) focused on civilians or reported separate analysis of civilians; 5) screened for depression using validated diagnostic criteria; 6) documented probable depression prevalence (number of patients in the cohort screening positive for depressive symptoms) or odds ratio (OR) among civilians in response to pre-defined risk factors; and 7) conducted within the NYC metropolitan area.

Data Extraction

Reported risk factors for probable depression were extracted and compared across studies. Our main exposure variables consisted of only risk factors that were similarly defined among at least two studies included in the metaanalysis. Probable depression prevalence, ORs, and standard errors corresponding to each risk factor were then extracted from each study. Our primary outcome was the OR for the association between the development of depression symptomatology and each risk factor. All ordinal and categorical risk factor variables were dichotomized to simplify outcome synthesis; the Cochran-Mantel-Haenszel method was used to evaluate ORs based on probable depression prevalence among combined groups and Woolf's method was used to combine weighted averages of log ORs.^{23,24} Other data elements that were extracted from each article were author surname, publication year, inclusion criteria, method and time period of cohort recruitment, sample size, baseline demographic characteristics of the sample, and overall prevalence of depressive symptoms.

Statistical Analysis

We used the DerSimonian and Laird (DL) random-effects model²⁵ to calculate pooled ORs and corresponding standard errors for each included risk factor. These pooled estimates were interpreted as summary effect sizes that expressed the common odds of screening positive for probable depression with the presence of a risk factor, versus absence of a risk factor. Weights were calculated by the inverse variance method.

Heterogeneity across studies was investigated by the Cochran Q test and measured using I² and H² statistics. We interpreted I² values of 0-25%, 26-50%, 51-75%, and 76-100% as unimportant, moderate, substantial, and considerable heterogeneity, respectively.²⁶ Publication bias was evaluated visually by examining symmetry in funnel plots. Egger regression²⁷ and the Begg-Mazumdar test²⁸ were used to quantify asymmetry by providing an

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estimate of correlations between effect sizes and corresponding variances. All analyses were performed using R software version 3.2.3 with the metafor package.²⁹

We conducted sensitivity analysis by stratifying studies by the type of diagnostic criteria used and the recruitment period midpoint. This allowed us to evaluate whether structured diagnostic interviews and symptom screening instruments yielded different findings, and whether more recent event exposure had a stronger influence on depressive symptoms. Prevalence estimates were weighted for the sample size of each study within a particular strata and averaged. Variance weighted least squares regression was used to test for temporal trend. Sensitivity analysis was also conducted by assessing how responsive our summary measures were to individual data. We repeatedly fit the DL model for each risk factor while censoring one study at a time.³⁰

To account for multiple tests, corresponding to the number of risk factors identified, we applied the Bonferroni correction³¹ to adjust the width of our confidence intervals (CIs) while testing for significance. Associations were considered significant at an α = 0.05/10 = 0.005 level.

Results

Electronic Search and Selection of Studies

We screened 2,103 search results and excluded 2,058 articles on the basis of their titles or abstracts for not being WTC related, not being an original research article, or not investigating depression symptomatology (Figure 1). We then identified 5 additional studies after screening references of reviews. Of the remaining 50 studies, 43 were excluded for using self-reported prior diagnosis or unclear screening measures of depression (n = 7), not reporting sufficient data that could be pooled for analysis (n = 11), focusing solely on responders or not separating civilians in their analysis (n = 19), using a sample that overlapped with another study and reported the same risk factors (n = 5), or not being conducted within the NYC metro area (n = 1). The remaining seven studies met the inclusion criteria for our meta-analysis.



Study Characteristics

The 7 studies included were prospective cohort studies of civilians with sample sizes ranging from n = 149 to n = 22,026 (Table 1). Each study cohort was recruited independently from the civilian population, making the possibility of subjects participating in multiple studies, or overlap in the samples, unlikely. Study cohorts were also recruited at different times, and the mid-point of their recruitment periods were on average 2.94 years post 9/11.

Characteristic	Period of Enrollment	Data Source	Screening Criteria	Probable Depression Prevalence, (%)	Sample Size	Age*, mean	Female Sex, (%)	Non-Hispanic White, (96)
Article								
Boscarino et al, 2011	October, 2002 – February, 2004	Prospective cohort of residents from the 5 boroughs in NYC	BSI -18	12.0	1,681	45.0	53.8	43.0
Caramanica et al., 2014	September, 2003 – November, 2004	Residents and passersby from Wave 3 of the World Trade Center Health Registry dataset	PHQ-8	16.7	22,026	44.7	53.8	62.9
Galea et al. 2002	October, 2001 – November, 2001	Sample of Manhattan residents living south of 110 ⁵ Street	SOD	9.7	988	42.0	53.0	71.6
Person et al. 2006	March, 2002 – June, 2002	Cross-sectional survey of residents from NIC metropolitan area	SOD	9.4	2,700	43,4	53.9	53.2
Neria et al., 2008	April, 2002 – January, 2003	Sample of adults seeking primary care in northern Manhattan community	PRIME -MD	21.5	929	47.2	69.5	÷
North et al. 2015	September, 2004 - September, 2005	Survivors working in the World Trade Center towers or buildings affected	SOD	26.0	373	45.1	53.0	10
Shear et al., 2006	January, 2003 – April, 2003	NVC residents utilizing Project Liberty crisis counseling services	SOD	31.0	149	45.8	69.0	58.0

Abbreviations: BSI, Brief Symptom Inventory: PHQ, Patient Health Questionnaire; SQD, Structured Clinical Interview for Diagnostic and Statistical Manual of Mental Disorders; PRIME -MD, Primary Care Evaluation of Mental Disorders.

*Age in years of subjects at the time they interviewed

Table 1: Baseline Characteristics of Studies that Satisfied the Inclusion Criteria

Across studies, the average age of subjects at the time they were interviewed was 44.5 years, 61% were non-Hispanic white, and 54% were female. Probable depression prevalence ranged from 9.4% to 31.0%, and the overall prevalence was 15.9%. Of the 7 studies, 4 used a full Structured Clinical Interview for DSM-IV (SCID), which is considered the gold standard for diagnosing Major Depressive Disorder.³² The remaining studies used screening instruments to identify depression symptomatology, which is why we refer to our outcome as "probable depression" rather than as "clinically diagnosed depression". The three studies reporting probable depression all used different screening instruments. Results from Caramanica et al. were adapted using Wave 3 of the WTC Health Registry, ^{16,20} which used the 8-item Patient Health Questionnaire (PHQ-8), applying a cutoff at scores \geq 10. This criteria has been shown to have sensitivity = 0.99 and specificity = 0.92 relative to SCID.³³ Boscarino et al.¹⁷ used the Brief Symptom Inventory-18 (BSI-18), which has been shown to have sensitivity = 0.71 and specificity = 0.87,³⁴ and Neria et al.¹⁸ used the Primary Care Evaluation of Mental Disorders (PRIME MD) Questionnaire, which has been shown to have sensitivity = 0.85 and specificity = 0.75,³⁵ all relative to structured clinical interviews.

Identification of Risk Factors

There were 10 risk factors that were reported in more than one study. These 10 risk factors were divided into 3 categories: baseline demographic characteristics, post-disaster attributes, and exposure characteristics. Demographic characteristics included greater age, female sex, minority race/ethnicity (non-white or Hispanic/Latino ethnicity), no college degree, and lower income level. Post-disaster attributes included post-9/11 social isolation, post-9/11 change in employment status, and not being married post-9/11 at the time of interview. Exposure characteristics included knowing someone injured or killed during the attacks and direct exposure to 9/11 traumatic events.

Not all risk factors were defined identically in each study. No college degree, minority race/ethnicity, post 9/11 social isolation, not being married post-9/11, and direct exposure were reported both as multilevel categorical variables and as dichotomous variables comparing the presence of a risk factor to its absence. To facilitate pooling together these results, multi-level variables were dichotomized.

Association between Probable Depression and Risk Factors

Among the baseline demographic risk factors, only minority race/ethnicity (OR, 1.40; 99.5% CI, 1.04 to 1.88) and lower income level (OR, 1.25; 99.5% CI, 1.09 to 1.43) were significantly associated with elevated odds of probable depression (Figure 2). The pooled ORs for greater age (OR, 0.86; 99.5% CI, 0.70 to 1.05), female sex (OR, 1.24; 99.5% CI, 0.98 to 1.59), and no college degree (OR, 1.32; 99.5% CI, 0.96 to 1.83) indicated that these risk factors were not significantly associated with odds of probable depression.

	OR [99.3% CI]
Baseline Demographic Characteristics	
Greater Age	
Caramanica et al., 2014	1.04 [0.76 to 1.44]
Boscarino et al., 2011	0.94 [0.50 to 1.77]
Person et al., 2006	0.79[0.01 to 1.04]
DL Model Subgroup OR	0.86[0.70101.05]
Female Sex	No. 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 1997 - 19
Caramanica et al., 2014 North et al., 2015	1.17 [0.99 to 1.38]
Boscarino et al., 2011	0.90[0.50 to 1.61]
Person et al., 2006	1.27 [0.89 to 1.79]
Galea et al., 2002	1.73 [0.93 to 3.24]
DL Model Subgroup OR	 1.24 [0.98 to 1.59]
Minority Race/Ethnicity	
Caramanica et al., 2014	Here 1.31 [1.18 to 1.53]
North et al., 2015 Borcarino et al., 2011	1.32 [0.53 to 3.27] 148 [0 92 to 2 40]
Person et al., 2006	1.05 [0.74 to 1.49]
Galea et al., 2002	2.29 [1.29 to 4.07]
DL Model Subgroup OR	◆ 1.40 [1.04 to 1.88]
No College Degree	
Caramanica et al., 2014	1.28 [1.08 to 1.51]
Boscarino et al. 2011	1.18 (0.65 to 2.14)
Person et al., 2006	1.15 [0.81 to 1.62]
Galea et al., 2002	2.54 [1.35 to 4.77]
DL Model Subgroup OR	• 1.32 [0.96 to 1.83]
Lower Income Level	127/1004-1401
Caramanica et al., 2014 Galea et al., 2003	1.2/[1.08 to 1.49]
Person et al., 2002	1.19 [0.89 to 1.59]
DL Model Subgroup OR	◆ 1.25 [1.09 to 1.43]
Post-Disaster Attributes	
Social Isolation	
Caramanica et al., 2014	H■H 2.23 [1.83 to 2.72]
Boscarino et al., 2011	1.34 [0.84 to 2.15]
Galea et al., 2002 Person et al., 2006	1.56 [0.83 to 2.95]
DL Model Subgroup OR	1.68 [1.13 to 2.49]
Change in Employment	
Caramanica et al., 2014	H■H 1.89 [1.58 to 2.27]
Galea et al., 2002	2.80 [1.20 to 6.51]
DL Model Subgroup OR	2.06 [1.30 to 3.26]
Not Married	
Caramanica et al., 2014	■ 1.71 [1.53 to 1.90]
Person et al. 2006	1.08 [0.60 to 1.92]
DL Model Subgroup OR	
Exposure Characteristics	
Knew Someone	
North et al., 2015	2.72[0.91 to 8.10]
Neria et al., 2006	2.38 [0.85 to 0.05]
Galea et al., 2002	2.27 [1.04 to 4.97]
DL Model Subgroup OR	 2.02 [1.42 to 2.89]
Direct Exposure	
Caramanica et al., 2014 North et al. 2015	1.11 [0.87 to 1.42]
Boscarino et al., 2011	2.28 [1.37 to 3.80]
Galea et al., 2002	1.19 [0.65 to 2.21]
Di Madal Subaraun OD	1 26 [0 69 to 2 30]

Fig. 2: Forest Plot of Odds Ratios Stratified by Risk Factor.

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All three post-disaster attributes (social isolation, change in employment, and not being married) were significantly associated with elevated odds of probable depression, and their pooled ORs were among the largest summary effect sizes of the ten evaluated. The largest effect size in this category was the pooled OR for post-9/11 change in employment status (OR, 2.06; 99.5% CI, 1.30 to 3.26). The second largest effect size in this category was the pooled OR for social isolation (OR, 1.68; 99.5% CI, 1.13 to 2.49), and the smallest effect size was the pooled OR for not being married post-9/11 (OR, 1.59; 99.5% CI, 1.18 to 2.15).

Knowing someone injured or killed by the attacks and direct exposure to traumatic events were classified as exposure characteristics. The former significantly increased odds of probable depression (OR, 2.02; 99.5% CI, 1.42 to 2.89) while the latter was not significantly associated with probable depression (OR, 1.26; 99.5% CI, 0.69 to 2.30).

Sensitivity Analysis for Diagnostic Criteria, Time from Disaster, and Removal of Individual Data

Out of the 7 studies included, 4 used full structured diagnostic interviews and 3 used symptom screening instruments. Studies using symptom screeners detected higher rates of probable depression prevalence (Table 2). All associations for risk factors that were statistically significant in the primary analysis remained statistically significant in both sub-groups of diagnostic criteria. Additionally, while the magnitude of the estimates remained comparable, in the sensitivity analysis female sex was a significant risk factor based on structured interviews, and lack of college degree and direct exposure were significant risk factors among studies that used symptom screeners.

	Structured Diagnostic Interview	Symptom Screening Instrument			
Studies	Shear et al., North et al., Person et al., & Galea et al.	Caramanica et al, Neria et al., Boscarino et al.			
Probable Depression Prevalence, (% [95% CI])	11.7 (10.7 to 12.7)	16.6 (16.1 to 17.0)			
Greater Age, (OR [99.5% CI])	0.79 (0.66 to 0.96)	1.02 (0.77 to 1.36)			
Female Sex, (OR [99.5% CI])	1.40 (1.06 to 1.86)	1.15 (0.98 to 1.35)			
No College Degree, (OR [99.5% Cl])	1.32 (0.99 to 1.76)	1.27 (1.08 to 1.49)			
Lower Income Level, (OR [99.5% CI])	1.21 (1.01 to 1.44)	1.27 (1.13 to 1.42)			
Minority Race/Ethnicity, (OR [99.5% CI])	1.30 (1.07 to 1.59)	1.35 (1.19 to 1.53)			
Social Isolation, (OR [99.5% CI])	1.54 (1.13 to 2.11)	2.06 (1.72 to 2.48)			
Change in Employment (OR [99.5% CI])	2.80 (1.55 to 5.04)	1.89 (1.66 to 2.15)			
Not Married, (OR [99.5% Cl])	1.78 (1.38 to 2.31)	1.68 (1.51 to 1.87)			
Knew Someone, (OR [99.5% CI])	2.41 (1.40 to 4.13)	1.77 (1.27 to 2.47)			
Direct Exposure, (OR [99.5% CI])	1.03 (0.61 to 1.76)	1.27 (1.02 to 1.58)			

Table 2: Study Outcomes Stratified by Type of Diagnostic Instruments Used

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We evaluated the prevalence of probable depression among all civilians included in studies that conducted interviews (midpoint of interview period) within a particular year (Supplementary Table). Although there were few studies included for each year, there was no significant temporal trend in depression prevalence (P = 0.25). We also assessed how stable our results were by evaluating the DL model for each risk factor after removing one study at a time (Table 3). This was only conducted for risk factors where results from more than two studies were synthesized. The female sex, minority race/ethnicity, no college degree, lower income level, and not being married post 9/11 risk factors lost or gained significance after removal of an individual study, however, all point estimates remained within the 95% confidence intervals of the original pooled estimates.

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Table 3. Leave-One-Out Sensitivity Analysis for Each Risk Factor.

Test Statistic	Pooled OR	99.5% CI	P-value for association	Q	τ²	12, (96)	Hž
Risk Factors and Study Remo	oved						
Greater Age	0.86	(0.70 to 1.05)	0.59	3.50	0.01	42.91	1.75
Person et al.	1.02	(0.78 to 1.35)	0.82	0.16	0	0	1
Caramanica et al.	0.81	(0.64 to 1.04)	0.02	0.51	0	0	1
Boscarino et al.	0.90	(0.62 to 1.31)	0.46	3.44	0.03	70.92	3.44
Female Sex	1.24	(0.98 to 1.59)	0.02	6.58	0.01	39.26	1.65
North et al.	1.21	(0.96 to 1.52)	0.02	4.98	0.01	39.77	1.66
Person et al.	1.26	(0.89 to 1.80)	0.07	6.39	0.03	53.03	2.13
Galea et al.	1.19	(0.98 to 1.44)	0.01	3.79	0	20.94	1.26
Caramanica et al.	1.31	(0.90 to 1.92)	0.05	5.81	0.04	48.36	1.94
Boscarino et al.	1.29	(1.03 to 1.62)	< 0.001	4.49	0.01	33.14	1.50
Lower Income Level	1.25	(1.09 to 1.43)	< 0.001	0.31	0	0	1
Person et al.	1.27	(1.09 to 1.48)	< 0.001	0	0	0	1
Galea et al.	1.25	(1.09 to 1.43)	< 0.001	0.29	0	0	1
Caramanica et al.	1.21	(0.95 to 1.55)	0.03	0.13	0	0	1
Minority Race/Ethnicity	1.40	(1.04 to 1.88)	<0.001	11.01	0.03	63.67	2.75
North et al	1.41	(1.02 to 1.96)	< 0.001	11.01	0.04	72.75	3.67
Person et al	1.53	(1.10 to 2.12)	< 0.001	6.68	0.03	55.11	2.23
Galea et al	1.29	(1.08 to 1.55)	< 0.001	3.92	0.01	23.49	1.31
Caramanica et al.	1.46	(0.88 to 2.40)	0.04	11.01	0.09	72.75	3.67
Boscarino et al.	1.39	(0.97 to 2.01)	0.01	10.66	0.05	71.85	3.55
No College Degree	1.32	(0.96 to 1.83)	0.02	11.04	0.04	63.77	2.76
North et al.	1.38	(0.97 to 1.94)	0.01	10.16	0.04	70.47	3.39
Person et al.	1.40	(0.88 to 2.22)	0.05	10.02	0.08	70.06	3.34
Galea et al.	1.24	(1.08 to 1.42)	< 0.001	1.41	0	0	1
Caramanica et al.	1.35	(0.79 to 2.33)	0.13	11.03	0.11	72.80	3.68
Boscarino et al.	1.36	(0.93 to 2.00)	0.03	10.87	0.05	72.39	3.62
Social Isolation	1.68	(1.13 to 2.49)	<0.001	12.92	0.06	76.77	4.31
Person et al.	1.72	(1.04 to 2.86)	< 0.001	9.2	0.08	78.26	4.60
Galea et al.	1.70	(1.08 to 2.69)	< 0.001	12.04	0.07	83.39	6.02
Caramanica et al.	1.48	(1.15 to 1.90)	< 0.001	0.48	0	0	1
Boscarino et al.	1.81	(1.21 to 2.71)	< 0.001	7.83	0.05	74.44	3.91
Not Married	1.59	(1.18 to 2.15)	0.002	5.03	0.02	60.25	2.52
Person et al.	1.42	(0.76 to 2.64)	0.12	4.84	0.09	79.35	4.84
Caramanica et al.	1.42	(0.71 to 2.83)	0.16	4.28	0.10	76.61	4.28
Boscarino et al.	1.71	(1.55 to 1.90)	< 0.001	0.1	0	0	1
Knew Someone	2.02	(1.42 to 2.89)	< 0.001	2.45	0	0	1
Galea et al.	1.96	(1.33 to 2.90)	< 0.001	1.36	0	0	1
Neria et al.	2.41	(1.42 to 4.07)	< 0.001	0.14	0	0	1
North et al.	1.98	(1.37 to 2.87)	< 0.001	1.35	0	0	1
Shear et al.	1.95	(1.35 to 2.82)	< 0.001	0.93	0	0	1
Direct Exposure	1.26	(0.69 to 2.30)	0.31	15.41	0.14	80.53	5.14
North et al.	1.43	(0.76 to 2.69)	0.12	12.70	0.13	84.26	6.35
Galea et al.	1.26	(0.55 to 2.88)	0.44	15.39	0.22	87.00	7.69
Caramanica et al.	1.30	(0.52 to 3.20)	0.43	10.77	0.26	81.43	5.39
Boscarino et al	1.10	(0.89 to 1.36)	0.23	1.92	0	0	1

Abbreviations: OR, odds ratio; O, confidence interval

Table 3: Leave-One-Out Sensitivity Analysis for Each Risk Factor

Publication Bias

Visual inspection of funnel plots for each risk factor and for all studies combined did not reveal any notable publication bias (Supplementary Figures 1 & 2). Additionally, neither the Begg-Mazumdar rank correlation test (Kendall's τ = 0.19, P = 0.10) nor the Egger regression test (z = 1.06, P = 0.29) detected statistically significant publication bias, although it is possible that the small number of included studies limits the power of this test.

Discussion

This study draws attention to probable depression after the 9/11 WTC terrorist attacks among civilians in the NYC metro area. We meta-analyzed effect sizes for various factors associated with probable depression from studies conducted over the past 15 years. Our analysis identified baseline demographic characteristics, exposure types to the WTC terrorist attacks, and post-disaster attributes, and quantified their association with post-9/11 probable depression.

The prevalence of probable depression varied considerably across studies included in our meta-analysis. This can partly be attributed to differences in the characteristics of each cohort and the type of diagnostic criteria employed. We observed disparities between results yielded by symptom screening instruments and full structured diagnostic interviews that could have implications for future studies of post-disaster probable depression. Symptom screening instruments, which generally have high sensitivity for diagnosis of major depression,^{33,34,35} detected higher rates of overall probable depression prevalence (16.6%) compared to full structured diagnostic interviews (11.7%). Moreover, although all risk factors that were significantly associated with probable depression in the primary analysis remained significant in the sensitivity analysis, the associations for some risk factors, including female sex, no college degree, and direct exposure, were not consistent across diagnostic criteria. The magnitude of these differences were small, however, and can partially be explained by high heterogeneity or lack of power since just two studies were present in subgroups that were inconsistent with the primary analysis.

Of the baseline demographic characteristics that we evaluated, only being of a minority race/ethnicity and earning lower income were significantly associated with elevated odds of probable depression. Since race/ethnicity and income are known to correlate with socioeconomic status,³⁶ our results may indicate an important association between post-disaster socioeconomic status and elevated risk of probable depression after a terrorist attack among civilians. Minority race/ethnicity has also been reported to be a risk factor for probable PTSD and depression among responders.^{6,7,8} Taken together, lower socioeconomic status could be used to identify target populations of mental health interventions so that the effect of intervention is maximized.

We also evaluated the association between WTC terrorist attack-related experiences and probable depression by focusing on two types of exposures: direct exposure to trauma, or knowing someone killed or injured by the attacks. Suffering the latter did not require being physically present in the vicinity of the attacks while they occurred, however, it was still significantly associated with elevated odds of probable depression. This finding strengthens evidence that probable depression may not be geographically constrained to the area affected after a disaster,¹³ and that bereavement may play an important role in driving the development of probable depression associated with terrorist attacks.¹⁸ Furthermore, the finding that direct exposure to 9/11 trauma was not significantly associated odds of probable depression suggests that terrorist attacks that expose few civilians directly to trauma may still lead to meaningful community-wide depressive symptoms.

An important limitation surrounding our discussion of the association between trauma exposure and probable depression is the definition of traumatic exposure that each study used. The studies included in this meta-analysis mainly found that traumatic exposure was not significantly associated with elevated odds of probable depression, but studies that used lenient definitions of what constituted trauma may have underestimated this association. We found, however, that the North et al.¹³ study used the most stringent definition of exposure to trauma, using "careful categorization of 9/11 trauma exposures based on the DSM-IV-TR definition," and still did not find a

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significant association between exposure to trauma and elevated odds of probable depression. As categorization of trauma became more careful and strict among the studies included in this meta-analysis, the effect size for its association with probable depression did not increase. Therefore, it is difficult to argue that inconsistent criteria for what constituted trauma in different studies explains the lack of association with elevated odds of probable depression observed.

We also classified social isolation, post-9/11 change in employment status, and not being married post-9/11 as post-disaster attributes and identified their association with probable depression. Social support is the perception that one belongs to a supportive social network and has access to a variety of social integration sources.³⁷ In the studies we encountered, social support was reported as the perceived number of social integration sources and was variably categorized. Our results suggest that a lack of social support, or social isolation, may play an important role in the development of probable depression after a terrorist attack; civilians without social support suffered 68% increased odds of probable depression after the WTC terrorist attacks. Similar results were observed for post-9/11 change in employment status and not being married post-9/11 as risk factors. The magnitudes of the pooled ORs for these three risk factors were among the four largest of the ten summary effect sizes evaluated in our study. Social integration resources, employment, and marriage are pivotal forms of social capital that influence how psychological stress affects civilians.³⁸ We found that access to these resources generally had a stronger association with probable depression than exposure or baseline demographics after the 9/11 WTC terrorist attacks. The causal direction of this association, however, remains uncertain as social capital and probable depression were both measured at the same time in the studies included.

These results must also be interpreted in the context of the study's limitations. Primarily, we could not include preexisting psychopathology as a risk factor due to large inconsistencies in how studies addressed it when reported. Pre-existing psychopathology has been strongly associated with post-disaster depression and other mental health outcomes.^{5,30} Its exclusion is not intended to undermine the need to consistently identify, screen, and triage patients with pre-existing mental disorders requiring treatment after a disaster such as a terrorist attack. Second, we could not consistently adjust for several confounding factors, such as prior trauma, when calculating ORs from reported data in the included studies. Some studies adjusted for covariates and provided adjusted ORs, but these adjustments may vary for the included studies. Third, there was between-study variability in the time of cohort recruitment and guestionnaire administration. Although we evaluated the influence of more recent event exposure in a sensitivity analysis and could not discern a significant trend in depression prevalence among cohorts with less recent event exposure, there may have been unaccounted confounding with cohort characteristics and specific risk factors. Fourth, we observed high heterogeneity in specific subgroups when calculating pooled ORs. This may stem from individual variation in the effect of the risk factor, but it also may be caused by variation in the original classification of risk factors among different studies. Being able to differentiate between the influence of being a direct eyewitness to trauma and being physically harmed during the events would have been useful in this context. Differentiating between changes in employment status that were related to the WTC terrorist attacks, versus changes in employment status that occurred for other reasons, is another example of how having more specific classification of risk factors in different studies may have strengthened our analysis.

Conclusions

This meta-analysis identified risk factors for probable depression associated with the WTC terrorist attacks among mixed-exposure groups from the civilian population in the NYC metro area that was not directly involved in rescue or recovery efforts. We found that social isolation, change in employment, and not being married after the WTC terrorist attacks were three of the four most strongly associated risk factors for probable depression. Furthermore, we found that direct exposure to WTC related traumatic events was not significantly associated with probable depression, while knowing someone injured or killed by the attacks was. Knowing someone injured or killed was the second most strongly associated risk factor for probable depression in this meta-analysis, suggesting that probable depression among civilians after a terrorist attack may primarily be bereavement driven. Finally, we

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identified minority race/ethnicity and lower income as the only baseline demographic characteristics that predicted elevated odds of probable depression. This association suggests that further efforts are necessary to understand and address the influence of socioeconomic status and probable depression among civilians after terrorist attacks.

Our analysis allows for a better understanding of the associations between probable depression and risk factors among civilians who were not involved in rescue or recovery efforts by providing quantitative estimates for each association. The strong association between lack of social capital and depressive symptoms suggests that monitoring employment status and the availability of support should be points of focus in future studies and intervention efforts. We further recommend that persons of lower socioeconomic status or difficulties coping with bereavement receive greater attention, irrespective of exposure to trauma, after terrorist attacks. These efforts could improve the efficiency at which high-risk persons from the civilian population are identified in future mental health interventions and disaster response efforts.

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Competing Interests

The authors have declared that no competing interests exist.

Data Availability

All relevant data are reported in the manuscript.

Ethics Statement

This analysis used only de-identified, publicly available, and previously published data sources. The need for informed patient consent was waived by the institutional review board and ethical committee at Weill Cornell Medicine.

Appendix

PRISMA Checklist

Supplementary Figure 1. Funnel plot of all effect sizes extracted from studies included



Supplemental Figure 1. Funnel plot of all effect sizes extracted from studies included.



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--0.50 0.08 0.58 1.08 1.58 2.08 --1.08 --0.58 8.08 0.58 1.80 Log Oddi Ratio

Supplemental Figure 2. Funnel plots of effect sizes for each risk factor.

Year Interviews Conducted	Studies	Prevalence (95% Cl) ^a 9.7 (7.9, 11.5)		
2001	Galea et al. ¹⁵			
2002	Neria et al. ¹⁸ & Person et al. ²¹	12.8 (11.7, 13.9)		
2003	Boscarino et al. ¹⁷ & Shear et al. ¹⁹	13.5 (120, 15.1)		
2005	North et al. ¹³	26.0 (21.5, 30.5)		
2011	Caramanica et al. ²⁰	16.5 (16.0, 17.0)		

Supplemental Table. Temporal Trend in Prevalence of Probable Depression.

References

1. North CS. Disaster Mental Health Epidemiology: Methodological Review and Interpretation of Research Findings. Psychiatry. 2016;79(2):130-146.

2. Neria Y, DiGrande L, Adams BG. Posttraumatic stress disorder following the September 11, 2001, terrorist attacks: a review of the literature among highly exposed populations. Am Psychol. Sep 2011;66(6):429-446.

3. American, Association P. Diagnostic and statistical manual of mental disorders, (5th ed.). Washington, DC2013.

4. Smith DJ, Nicholl BI, Cullen B, et al. Prevalence and characteristics of probable major depression and bipolar disorder within UK biobank: cross-sectional study of 172,751 participants. PLoS One. 2013;8(11):e75362.

5. North CS, Pfefferbaum B. Mental health response to community disasters: a systematic review. JAMA. Aug 2013;310(5):507-518.

6. Bromet EJ, Hobbs MJ, Clouston SA, Gonzalez A, Kotov R, Luft BJ. DSM-IV post-traumatic stress disorder among World Trade Center responders 11-13 years after the disaster of 11 September 2001 (9/11). Psychological medicine. Mar 2016;46(4):771-783.

7. Stellman JM, Smith RP, Katz CL, et al. Enduring mental health morbidity and social function impairment in world trade center rescue, recovery, and cleanup workers: the psychological dimension of an environmental health disaster. Environmental health perspectives. Sep 2008;116(9):1248-1253.

8. Zvolensky MJ, Kotov R, Schechter CB, et al. Post-disaster stressful life events and WTC-related posttraumatic stress, depressive symptoms, and overall functioning among responders to the World Trade Center disaster. Journal of psychiatric research. Feb 2015;61:97-105.

9. Berger W, Coutinho ES, Figueira I, et al. Rescuers at risk: a systematic review and meta-regression analysis of the worldwide current prevalence and correlates of PTSD in rescue workers. Soc Psychiatry Psychiatr Epidemiol. Jun 2011;47(6):1001-1011.

10. Stein CR, Wallenstein S, Shapiro M, et al. Mortality among World Trade Center rescue and recovery workers, 2002-2011. American journal of industrial medicine. Feb 2016;59(2):87-95.

11. Silver RC, Holman EA, McIntosh DN, Poulin M, Gil-Rivas V. Nationwide longitudinal study of psychological responses to September 11. JAMA. Sep 2002;288(10):1235-1244.

12. Schlenger WE, Caddell JM, Ebert L, et al. Psychological reactions to terrorist attacks: findings from the National Study of Americans' Reactions to September 11. JAMA. Aug 2002;288(5):581-588.

13. North CS, Pollio DE, Hong BA, Pandya A, Smith RP, Pfefferbaum B. The postdisaster prevalence of major depression relative to PTSD in survivors of the 9/11 attacks on the World Trade Center selected from affected workplaces. Compr Psychiatry. Jul 2015;60:119-125.

14. Neria Y, Gross R, Marshall RD, Susser ES. 9/11: Mental Health in the Wake of Terrorist Attacks. Cambridge University Press; 2006.

15. Galea S, Ahern J, Resnick H, et al. Psychological sequelae of the September 11 terrorist attacks in New York City. N Engl J Med. Mar 2002;346(13):982-987.

16. Murphy J, Brackbill RM, Thalji L, Dolan M, Pulliam P, Walker DJ. Measuring and maximizing coverage in the World Trade Center Health Registry. Stat Med. Apr 2007;26(8):1688-1701.

17. Boscarino JA, Adams RE, Figley CR. A prospective cohort study of the effectiveness of employer-sponsored crisis interventions after a major disaster. Int J Emerg Ment Health. 2005;7(1):9-22.

Risk Factors for Depression Among Civilians After the 9/11 World Trade Center Terrorist Attacks: A Systematic Review and Meta-Analysis - PLOS C...

18. Neria Y, Olfson M, Gameroff MJ, et al. The mental health consequences of disaster-related loss: findings from primary care one year after the 9/11 terrorist attacks. Psychiatry. 2008;71(4):339-348.

19. Shear KM, Jackson CT, Essock SM, Donahue SA, Felton CJ. Screening for complicated grief among Project Liberty service recipients 18 months after September 11, 2001. Psychiatr Serv. Sep 2006;57(9):1291-1297.

20. Caramanica K, Brackbill RM, Liao T, Stellman SD. Comorbidity of 9/11-related PTSD and depression in the World Trade Center Health Registry 10-11 years postdisaster. J Trauma Stress. Dec 2014;27(6):680-688.

21. Person C, Tracy M, Galea S. Risk factors for depression after a disaster. The Journal of nervous and mental disease. Sep 2006;194(9):659-666.

22. Liberati A, Altman DG, Tetzlaff J, et al. The PRISMA statement for reporting systematic reviews and metaanalyses of studies that evaluate healthcare interventions: explanation and elaboration. BMJ. Jul 2009;339:b2700.

23. Mantel N, Haenszel W. Statistical aspects of the analysis of data from retrospective studies of disease. J Natl Cancer Inst. Apr 1959;22(4):719-748.

24. Woolf B. On estimating the relationship between blood group and disease. Annals of Human Genetics. 1955;19:2.

25. DerSimonian R, Laird N. Meta-analysis in clinical trials. Control Clin Trials. Sep 1986;7(3):177-188.

26. Higgins JP, Thompson SG. Quantifying heterogeneity in a meta-analysis. Stat Med. Jun 2002;21(11):1539-1558.

27. Egger M, Davey Smith G, Schneider M, Minder C. Bias in meta-analysis detected by a simple, graphical test. BMJ. Sep 1997;315(7109):629-634.

28. Begg CB, Mazumdar M. Operating characteristics of a rank correlation test for publication bias. Biometrics. Dec 1994;50(4):1088-1101.

29. Viechtbauer W. Conducting Meta-Analyses in R with the metafor Package. Journal of Statistical Software. 2010;36:48.

30. Liu B, Tarigan LH, Bromet EJ, Kim H. World Trade Center disaster exposure-related probable posttraumatic stress disorder among responders and civilians: a meta-analysis. PLoS One. 2014;9(7):e101491.

31. Bender R, Lange S. Adjusting for multiple testing--when and how? J Clin Epidemiol. Apr 2001;54(4):343-349.

32. Sheehan DV, Lecrubier Y, Sheehan KH, et al. The Mini-International Neuropsychiatric Interview (M.I.N.I.): the development and validation of a structured diagnostic psychiatric interview for DSM-IV and ICD-10. J Clin Psychiatry. 1998;59 Suppl 20:22-33;quiz 34-57.

33. Kroenke K, Spitzer RL, Williams JB. The PHQ-9: validity of a brief depression severity measure. J Gen Intern Med. Sep 2001;16(9):606-613.

34. Petkus AJ, Gum AM, Small B, Malcarne VL, Stein MB, Wetherell JL. Evaluation of the factor structure and psychometric properties of the Brief Symptom Inventory-18 with homebound older adults. Int J Geriatr Psychiatry. Jun 2010;25(6):578-587.

35. Spitzer RL, Kroenke K, Williams JB. Validation and utility of a self-report version of PRIME-MD: the PHQ primary care study. Primary Care Evaluation of Mental Disorders. Patient Health Questionnaire. JAMA. Nov 1999;282(18):1737-1744.

36. Ostler K, Thompson C, Kinmonth AL, Peveler RC, Stevens L, Stevens A. Influence of socio-economic deprivation on the prevalence and outcome of depression in primary care: the Hampshire Depression Project. The British journal of psychiatry : the journal of mental science. Jan 2001;178(1):12-17.

Risk Factors for Depression Among Civilians After the 9/11 World Trade Center Terrorist Attacks: A Systematic Review and Meta-Analysis – PLOS C ...

37. Peirce RS, Frone MR, Russell M, Cooper ML, Mudar P. A longitudinal model of social contact, social support, depression, and alcohol use. Health Psychol. Jan 2000;19(1):28-38.

38. Hobfoll SE. Conservation of resources. A new attempt at conceptualizing stress. Am Psychol. Mar 1989;44(3):513-524.