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Elena Salmoirago Blotcher
University of Massachusetts Medical School

George Fitchett
Rush University

Kathleen M. Hovey
University at Buffalo

See next page for additional authors

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Frequency of Private Spiritual Activity and Cardiovascular Risk in Postmenopausal Women: The Women's Health Initiative

Authors

Elena Salmoirago Blotcher, George Fitchett, Kathleen M. Hovey, Eliezer Schnall, Cynthia Thomson, Christopher A. Andrews, Sybil Crawford, Mary Jo O'Sullivan, Stephen Post, Rowan T. Chlebowski, and Judith K. Ockene

Keywords

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Title: Frequency of private spiritual activity and cardiovascular risk in post-menopausal women:
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Short title: Spirituality and cardiovascular risk in women

Elena Salmoirago-Blotcher, MD¹); George Fitchett, PhD²); Kathleen M. Hovey, MS³); Eliezer Schnall, PhD⁴); Cynthia Thomson, PhD, ⁵); Christopher A. Andrews, PhD^{3,6}); Sybil Crawford, PhD¹); Mary Jo O'Sullivan, MD⁷); Stephen Post, PhD⁸); Rowan T. Chlebowski, MD⁹); Judith Ockene, PhD¹)

¹) University of Massachusetts Medical School; ²) Rush University; ³) University at Buffalo;
⁴) Yeshiva University; ⁵) University of Arizona; ⁶) University of Michigan; ⁷) University of
Miami; ⁸) Stony Brook University; ⁸) University of California-Los Angeles

Corresponding author:

Elena Salmoirago-Blotcher
Division of Cardiovascular Medicine
University of Massachusetts Medical School
55 Lake Avenue North, room S3-855
Worcester, MA 01655
Phone: 508-856-4413; Fax: 508-856-4571
Email: Elena.Salmoirago-Blotcher@umassmed.edu

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ABSTRACT

Purpose Spirituality has been associated with better cardiac autonomic balance, but its association with cardiovascular risk is not well studied. We examined whether more frequent private spiritual activity was associated with reduced cardiovascular risk in postmenopausal women enrolled in the Women's Health Initiative Observational Study.

Methods Frequency of private spiritual activity (prayer, Bible reading, and meditation) was self-reported at year 5 of follow-up. Cardiovascular outcomes were centrally adjudicated, and cardiovascular risk was estimated from proportional hazards models.

Results Final models included 43,708 women (mean age: 68.9 ± 7.3 ; median follow-up: 7.0 years) free of cardiac disease through year 5 of follow-up. In age-adjusted models private spiritual activity was associated with increased cardiovascular risk (HR: 1.16; CI 1.02, 1.31, weekly vs. never; 1.25; CI 1.11, 1.40, daily vs. never). In multivariate models adjusted for demographics, lifestyle, risk factors, and psychosocial factors, such association remained significant only in the group with daily activity (HR 1.16; CI: 1.03, 1.30). Subgroup analyses indicate this association may be driven by the presence of severe chronic diseases.

Conclusion In aging women, higher frequency of private spiritual activity was associated with increased cardiovascular risk, likely reflecting a mobilization of spiritual resources in order to cope with aging and illness.

Key words: women's health, cardiovascular diseases, spirituality

List of abbreviations and acronyms

R/S = Religiosity/spirituality

MESA=Multi Ethnic Study of Atherosclerosis

SNS= Sympathetic Nervous System

WHI= Women's Health Initiative

CVD= cardiovascular disease

OS= Observational Study

YR=year

MET = Metabolic Equivalent of Task

STROBE = STrengthening the Reporting of OBServational studies in Epidemiology

Introduction

There is increasing interest in the study of the relationship between religiosity/spirituality (R/S) and health, fostered by the finding that individuals who regularly attend religious services [1-9] have a reduction in overall mortality of almost 20%. The evidence for the possible association between R/S and reduced cardiovascular risk, however, is less consistent. Multiple prospective cohort and case-control studies [3, 9-12] have linked R/S with reduced cardiovascular mortality and morbidity. A cross-sectional study conducted in a rural population in India has shown that men participating in prayer and yoga had a lower prevalence of coronary heart disease after adjustment for coronary risk factors. [13] A composite R/S measure (frequency of service attendance, prayer and fasting) was inversely associated with acute coronary events in a recent case-control study [14] conducted in a predominantly Moslem and Christian population. On the other hand, other studies somewhat contradicted these results. Schnall [8] reported a lack of association between several self-reported measures of R/S (religious affiliation, frequency of service attendance, and religious coping) and cardiovascular mortality in a large population of postmenopausal women. More frequent service attendance, prayer or meditation, and higher daily spiritual experience (DSE) scores were not associated with reduced cardiovascular risk in a prospective cohort study involving more than five thousand healthy men and women enrolled in the Multi-Ethnic Study of Atherosclerosis (MESA). [15] In a study of non-institutionalized older adults, Colantonio and colleagues reported that more frequent worship attendance was associated with a reduced risk of stroke in unadjusted models [16]; in adjusted models, however, this association was no longer significant. Another study conducted in myocardial infarction survivors with depression and low social support did not show an association between DSE scores, church attendance, or frequency of private spiritual activity and non-fatal cardiac events. [17]

The link between frequency of individual private spiritual practices such as prayer, meditation, and devotional reading and cardiovascular health has been less frequently investigated at the population level compared with the more public aspects of R/S, namely, religious services attendance. Practices such as prayer or meditation are particularly interesting because their association with cardiovascular risk is less likely to be affected by variables – such as social support – that are known confounders of the association between service attendance and cardiovascular outcomes. Furthermore, unlike service attendance, the physiological changes occurring during prayer and meditation have been extensively studied and consist in a decrease in sympathetic nervous system (SNS) activity, oxygen consumption, respiratory rate, and minute ventilation. [18] Prayer or mantra recitation have been linked with higher levels of cardiac autonomic control [19] and increased baro-reflex sensitivity, [20] conditions which in turn have been associated with a reduced risk of cardiac arrhythmias and cardiac mortality. [21-23] A reduction in the activity of the SNS may be beneficial for patients with coronary heart disease, as shown by findings of increased exercise tolerance and delayed onset of electrocardiographic signs of ischemia in patients enrolled in a transcendental meditation program. [24] Similar results were shown by Benson, who reported that meditation reduced the number of premature ventricular contractions in these patients. [25]

The primary aim of this study was to evaluate whether private spiritual activity (prayer, meditation and reading of religious literature) is associated with a reduced incidence of fatal and non-fatal cardiovascular events in a large population of community-dwelling postmenopausal women enrolled in the Women's Health Initiative (WHI) Observational Study (OS). The WHI affords a unique opportunity to evaluate this association given its large sample size as well as the adjudication of cardiovascular outcomes. Our study hypothesis was that women who report to

engage in private spiritual activity, compared to women reporting no spiritual activity, would have a reduced cardiovascular risk.

The secondary aim was to assess whether the association may differ in women with a history of severe chronic diseases prior to the assessment of private spiritual activity compared to women without such history. This analysis was undertaken in light of the important role of R/S, including private spiritual activity, in coping with serious illness. This is illustrated by the fact that the proportion of people with serious illnesses (e.g., cancer) who report praying for their health (77%) [26] may be double the proportion in the general population (43%) [27]. Further, there is evidence [28] that the association between private spiritual activity and cardiovascular risk may differ depending on whether such activity is a longstanding, lifelong behavior or rather an episodic phenomenon elicited in response to stressful events or illness.

Despite the concerns raised in regard to the methodological limitations and lack of direct clinical applications of the research in this field, [29] the study of the role of R/S in the prevention of cardiovascular disease (CVD) may have important public health consequences if these practices are associated with a lower cardiovascular risk. [30]

Methods

The WHI included a set of randomized clinical trials and a separate OS. [29] The OS, the focus of this analysis, was a large prospective cohort study conducted in 93,676 postmenopausal women ineligible or unwilling to participate in the clinical trials. To be eligible women had to be 50 to 79 years old, postmenopausal, and reside in the study recruitment area for at least 3 years following enrollment. Exclusion criteria were medical conditions predictive of a survival time of less than 3 years; conditions inconsistent with study participation such as alcoholism, drug dependency, mental illness, and dementia; and enrollment in another study. All participants

provided written informed consent according to the human subjects' protection oversight at the 40 participating sites. Recruitment (1994-1998) was conducted through mailings to eligible women from large mailing lists. Women were followed through 2005 as part of the original study and were offered the opportunity to continue for an additional 5 years in the WHI extension study. After obtaining informed consent, yearly questionnaires were administered by mail between 2005 and 2010.

All OS participants underwent a baseline visit including physical measurements (height, weight, blood pressure, heart rate, waist and hip circumferences), blood specimen collection, a medication/supplement inventory, and completion of medical, family/ reproductive history, lifestyle/behavioral factors, and quality of life questionnaires. Follow-up included an on-site clinic visit three years after enrollment and annual mailings (a medical history update and questionnaires about lifestyle and dietary habits, demographics, hormone therapy, and psychosocial variables). The WHI OS outcomes were coronary heart disease, stroke, breast and colorectal cancer, osteoporotic fractures, diabetes, and total mortality.

Study population

Inclusion criteria for this secondary analysis were completion of the follow-up year 5 (YR5) assessments, and absence of self-reported CVD and any cardiovascular events through YR5. Women who did not consent to the WHI extension study were censored at the date of last follow up.

Exposure

Time spent in private spiritual activity was assessed only at YR5 using a self-administered questionnaire completed approximately 5 years (1999-2003) after enrollment in the original study asking how often the participant spent time in the following activities: prayer,

meditation, or reading religious literature during the prior year. The possible answers were never, a few times per year, a few times per month, about once a week, a few times per week, every day.

Outcome

The outcome for this analysis was fatal and non-fatal incident cardiovascular events, a cumulative endpoint defined as the first occurrence of non-fatal angina, myocardial infarction, congestive heart failure, coronary and carotid revascularization procedures, stroke, transient ischemic attack, peripheral arterial disease, and any adjudicated fatal cardiovascular event. Outcomes were identified by self-report on the medical history update or by reporting directly to clinic staff in the intervals between questionnaires. Centrally trained physicians adjudicated cardiovascular outcomes. [32]

Covariates and other variables of interest

Covariates were selected based on the literature and/or whether they were associated with frequency of private spiritual activity, cardiovascular events, or both. Whenever possible, we used information collected at the same time (YR5) as frequency of private spiritual activity. When variables had not been collected at YR5 we used covariates collected at YR3 or at baseline, whichever was closest in time. Except for body mass index (BMI) all covariates were collected by means of self-administered questionnaires and included: age; race/ethnicity (American Indian or Alaskan Native, Asian or Pacific Islander, Black or African-American, Hispanic/Latino, White, other); marital status (never married, married or in marriage-like relationship, widowed, divorced/separated); total years of education; and income (<\$10,000; \$10,000-19,999; \$20,000-34,999; \$35,000-49,999; \$50,000-74,999; \$75,000-99,999; \$100,000-149,000; >\$150,000; does not know). Risk factors for CVD included history of hypertension and

diabetes; family history; smoking (ever, never, current); and BMI (weight in kilograms/ (height in meters)² calculated from direct measurements of height and weight performed at YR3).

Cholesterol levels were not measured in the entire OS cohort so history of elevated cholesterol requiring pharmacological treatment was used as a proxy. Physical activity was calculated as energy expenditure from recreational physical activity (total MET-hours per week). Alcohol intake was measured as the number of servings of alcohol per week. Psychosocial variables included social support (from the Medical Outcomes Study); [33] physical functioning, emotional wellbeing and satisfaction with quality of life (Medical Outcome Study 36-Item Health Survey); [34] and depression (assessed using the shortened version of the Center for Epidemiological Studies Depression Scale). [35]

History of severe chronic diseases was defined as a self-reported history of severe non-cardiovascular co-morbidities before YR5 including diabetes, liver disease, dialysis, asthma, emphysema, lupus, Alzheimer disease, multiple sclerosis, Parkinson disease, amyotrophic lateral sclerosis as well as all adjudicated cancer outcomes (except non-melanoma skin cancers).

Statistical analysis

We used descriptive statistics to describe the baseline characteristics of the study population across categories of frequency of private spiritual activity. ANOVA F and independence χ^2 tests were used to evaluate differences. The association between private spiritual activity (categorized as daily, once or more per week, less than once a week, and never) and cardiovascular events was evaluated using univariate and multivariate Cox proportional hazards regression models. We tested the assumption of the proportional hazard model by entering time dependent covariates into the model, and no violations were observed. Several models were fitted: unadjusted; adjusted for age only (model 1); adjusted also for demographic

variables (model 2); adjusted for all variables included in model 2 plus health behaviors (model 3); adjusted for model 3 variables plus cardiovascular risk factors (model 4); adjusted for model 4 variables and physical functioning, satisfaction with QOL, and social support (model 5). The final model included model 5 variables plus depression and emotional wellbeing (model 6). The need for a separate model including psychological variables arose because such variables may be on the causal pathway linking spirituality and survival. In addition, we planned to conduct a subgroup analysis by presence/absence of history of severe chronic diseases prior to YR5.

Results are presented as unadjusted and adjusted hazard ratios (HR) with 95% confidence intervals. P values <0.05 were considered significant. All statistical analyses were performed using SAS/STAT® version 9.2, SAS Institute, Cary, NC.

Results

Figure 1 (STROBE diagram) [36] shows the flow of patients through this study. Of the original cohort of 93,676 women enrolled in the OS, 61,370 met the eligibility criteria for this study. Due to missing information on covariates included in multivariate models, 17,662 women were excluded from the final analysis. These women tended to be more frequently African American (9% vs. 5.4%) or Hispanic (5% vs. 2.8%), less educated (post-graduate education, 28.3% vs. 34.6%) and more frequently current smokers (5% vs. 3.7%) compared to subjects included in the analysis, but had otherwise similar characteristics with regard to spiritual practices and cardiovascular events (no spiritual practice, excluded vs. included: 17.2% vs. 17.9); cardiovascular events: 6.3% vs. 5.8%).

The baseline characteristics of the study population are shown in Table 1. Women with higher frequency of private spiritual activity tended to be older, less educated, of African American ancestry, widowed or never married, and of lower socio-economic status; they also

reported lower functional status scores. The prevalence of a diagnosis of a severe chronic disease prior to YR5 as well as of most risk factors for CVD was also higher in women reporting more frequent private spiritual activity. Specifically, these women were more frequently obese, used cholesterol-lowering drugs more often, had a history of hypertension or diabetes and reported lower levels of physical activity. The only exception among coronary risk factors was smoking, as women frequently engaging in private spiritual activity were less likely to be current or former smokers. Alcohol intake also was significantly lower in those with higher frequency of private spiritual activity. Finally, women with more frequent spiritual practice had slightly better emotional wellbeing, higher social support and a slightly lower prevalence of depression and antidepressants use.

The median duration of follow-up was 7 years, during which 2,554 cardiovascular events were observed (402 fatal). Age-adjusted models (Table 2, Model 1) showed that women who reported weekly and daily spiritual practice had a higher risk of cardiovascular events compared to women who never engaged in such activity (HR: 1.16; CI 1.02, 1.31, weekly vs. never; 1.25; CI 1.11, 1.40, daily vs. never). In a series of models that further adjusted for other demographic factors (model 2), lifestyle habits (model 3), and cardiovascular risk factors (model 4), women who reported daily private spiritual activity had a significantly increased risk for cardiovascular events (HR in the fully adjusted model 1.16, CI 1.03, 1.30) compared to women who did not reported such activity. Further adjustment for psychosocial measures (model 5) and depression and emotional wellbeing (model 6) did not modify these associations. We also fitted a model adjusting for presence/absence of severe chronic diseases in addition to the variables included in model 2 (age, ethnicity, marital status, income, and education), but the association between cardiovascular events and spiritual practice was not altered (HR: 1.08; CI 0.95, 1.23, weekly vs.

never; 1.13; CI 1.01, 1.27, daily vs. never). Finally, trend analysis showed that the risk of cardiovascular events increased with rising frequency of private spiritual activity (p for trend = 0.010).

Subgroup analysis

Table 3 and Figure 2 show the results of the subgroup analysis by presence/absence of self-reported severe chronic diseases before YR5. Results indicate that the presence of severe chronic diseases was not an effect modifier (p= 0.6449).

Discussion

In previous population-based studies more frequent worship participation [3, 10] or, among Jews, adherence to orthodox practices and teachings [11, 12] were found to be associated with a reduced cardiovascular risk. In our study of a large population of community-dwelling postmenopausal women, however, when we focused our attention on frequency of meditation, prayer or reading of religious texts, we found that this dimension of R/S was not associated with a reduced cardiovascular risk and in fact women reporting daily private spiritual activity had a higher risk of cardiovascular events compared to women who reported no private spiritual activity.

The present study was designed to follow-up on the findings of a previous investigation conducted in the WHI-OS, [8] which did not detect an association between baseline measures of R/S and cardiovascular risk; these measures however did not include private spiritual activity. The relationship between multiple measures of R/S (worship attendance, prayer, and DSE scores) and cardiovascular risk factors and events has been studied among more than 5,000 participants aged 45-84 in the MESA study. [15] As in the present study, higher levels of R/S, including prayer, were associated with greater risk of obesity and decreased risk of drinking and

smoking. Unlike the results reported here, among the MESA participants there was no significant association between frequency of prayer and cardiovascular risk. Our study, however, had a larger sample size and a longer duration of follow-up; in addition, even in the MESA analysis, the point estimate in the group with daily private spiritual activity, although not significant, indicated an increased risk of cardiovascular events in the more religiously involved.

A possible explanation for our findings would be confounding by the presence of severe chronic co-morbidities; in fact, the prevalence of severe chronic co-morbidities prior to YR5 was associated with a higher frequency of private spiritual activity (Table 1). In multivariate models however, the presence of severe chronic disease was not a confounder. Although the interaction term was not significant, the results of the stratified analysis nevertheless suggest that women with a prior diagnosis of severe chronic diseases likely drive the association between private spiritual activity and increased cardiovascular risk. Similar findings have been reported in another study conducted in an elderly population [28] in which the protective association between private spiritual activity and reduced overall mortality was detected only in individuals without impairment in activities of daily living, while it was not detected in the cohort reporting such condition.

Our results are not consistent with some of the previously published studies indicating that R/S is associated with a lower cardiovascular risk. [9] It appears, however, that our knowledge about the link between R/S and cardiovascular health is far from being conclusive for several reasons. For example, studies that have detected a protective association between R/S and cardiovascular risk measured very different aspects of the R/S experience. Some researchers measured worship attendance only [3, 10] while others measured self-reported religiosity within specific religious groups. [1, 11, 12] Studies also differed as to whether they adjusted for

coronary risk factors (i.e. diabetes, high cholesterol) and other variables that may confound the association between R/S and cardiovascular mortality. The only studies extensively adjusting for coronary risk factors and reporting an association between R/S (defined as being orthodox vs. secular) and reduced risk of cardiovascular mortality or acute myocardial infarction were conducted in a Jewish population in Israel. [1, 11, 12] Orthodox Judaism, however, is somewhat unique in that it regulates every aspect of life, [37, 38] thus raising the question of whether these results are generalizable to other religious affiliations. The populations studied in other investigations were also heterogeneous with regard to their age and health status, with protective associations between R/S and cardiovascular risk usually shown in younger and healthier populations, [3, 10, 12, 13] but not in older persons with concomitant diseases. [8, 9, 16, 17] The direction and the magnitude of the association between R/S and cardiovascular risk may change depending on whether the exposure began in healthy individuals and early in life as opposed to a later stage, when individuals may turn to religion as a coping mechanism in the face of declining health and advancing age. [28, 39] Finally, the duration of follow-up differed greatly among studies; studies reporting a protective effect usually had a longer follow-up duration. [3, 10, 11]

This study has several strengths: first, a relatively long duration of follow-up; second, extensive adjustment for coronary risk factors, demographic and psychosocial variables; third, a relatively understudied population (women); and fourth, the validation of cardiovascular outcomes. The limitations of this analysis include the assessment of private spiritual activity as an aggregate only, with the resulting inability to assess associations between specific activities (i.e. meditation and/or prayer vs. reading of religious texts) and cardiovascular events; and the collection of private spiritual activity data only at YR5. Furthermore, some covariates were not collected at the same time point (YR5) as the spirituality activity assessment, and we had to use

baseline or YR3 covariates. Residual confounding (in particular regarding coronary risk factors such as blood pressure and serum lipid levels that were not directly measured) as well as selection bias (due to the large number of women excluded) are additional limitations of this analysis. Also, we cannot rule out the possible contribution of survivor bias (e.g., women with lower levels of private spirituality activity not surviving long enough to be included in the current analysis) to our findings. Finally, results are not generalizable to younger or male populations, and to women that were excluded from participation in the parent study (i.e. women with alcoholism, drug dependency, mental illness, and dementia).

In conclusion, our results confirm recent indications that higher R/S does not prevent fatal and non-fatal cardiovascular events, [8, 15] at least in older female populations. Further research involving younger and healthier populations is needed to achieve a better understanding of the role of R/S in the prevention of cardiovascular risk.

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\$10,000-\$19,999	3,675 (8.4)	496 (6.3)	598 (6.6)	847 (7.8)	1,734 (10.9)	
\$20,000-\$34,999	9,265 (21.2)	1,288 (16.5)	1,743 (19.1)	2,337 (21.6)	3,897 (24.4)	
\$35,000-\$49,999	8,795 (20.1)	1,392 (17.8)	1,761 (19.3)	2,248 (20.8)	3,394 (21.3)	
\$50,000-\$74,999	9,530 (21.8)	1,741 (22.2)	2,045 (22.5)	2,462 (22.8)	3,282 (20.6)	
\$75,000-\$99,999	4,723 (10.8)	1,051 (13.4)	1,133 (12.4)	1,140 (10.6)	1,399 (8.8)	
\$100,000-\$149,999	3,625 (8.3)	912 (11.7)	936 (10.3)	848 (7.8)	929 (5.8)	
≥\$150,000	2,055 (4.7)	641 (8.2)	537 (5.9)	470 (4.4)	407 (2.6)	
Don't Know	1,050 (2.4)	148 (1.9)	209 (2.3)	241 (2.2)	452 (2.8)	
<hr/>						
BMI, year 3 ¹						
Normal <25	18,258 (41.8)	3,858 (49.3)	3,951 (43.4)	4,107 (38.0)	6,342 (39.7)	<0.0001 ^{b)}
Overweight 25-<30	15,174 (34.7)	2,568 (32.8)	3,159 (34.7)	3,926 (36.3)	5,521 (34.6)	
Obese ≥30	10,276 (23.5)	1,403 (17.9)	1,999 (22.0)	2,777 (25.7)	4,097 (25.7)	
<hr/>						
Depression, year 3 ¹	3,653 (8.4)	653 (8.3)	757 (8.3)	991 (9.2)	1,252 (7.8)	0.0020 ^{b)}
Antidepressant use, year 3 ²	3,842 (8.8)	692 (8.9)	810 (8.9)	1,022 (9.5)	1,318 (8.3)	0.0090 ^{b)}
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Physical activity, year 5 (MET-hours/week; mean (SD))	14.25 (14.01)	15.92 (14.99)	14.88 (14.09)	13.62 (13.43)	13.50 (13.75)	<0.0001 ^{c)}
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High cholesterol requiring pills ever, baseline	5,095 (11.7)	845 (10.8)	1,043 (11.5)	1,279 (11.8)	1,928 (12.1)	0.0274 ^{b)}
<hr/>						
History of hypertension, year 5	15,621 (35.7)	2,416 (30.9)	3,143 (34.5)	3,952 (36.6)	6,110 (38.3)	<0.0001 ^{b)}
<hr/>						
Diabetes, year 5	1,996 (4.6)	289 (3.7)	368 (4.0)	516 (4.8)	823 (5.2)	<0.0001 ^{b)}
<hr/>						
Relative had heart attack, baseline	21,882 (50.1)	3,736 (47.7)	4,515 (49.6)	5,527 (51.1)	8,104 (50.8)	<0.0001 ^{b)}
<\$10,000	990 (2.3)	160 (2.0)	147 (1.6)	217 (2.0)	466 (2.9)	<0.0001 ^{d)}

Social Support, baseline (mean (SD))	36.67 (7.37)	36.26 (7.75)	36.23 (7.40)	36.59 (7.15)	37.18 (7.26)	<0.0001 ^{c)}
Physical Functioning, year 3 (mean (SD))	82.13 (19.62)	84.21 (18.76)	83.14 (18.94)	81.76 (19.42)	80.78 (20.41)	<0.0001 ^{c)}
Emotional Well-being, year 3 (mean (SD))	81.37 (13.91)	81.03 (14.50)	80.72 (14.08)	80.94 (13.82)	82.20 (13.52)	<0.0001 ^{c)}
Satisfaction with quality of life, year 3 (mean (SD))	8.31 (1.78)	8.22 (1.84)	8.21 (1.80)	8.28 (1.77)	8.44 (1.73)	<0.0001 ^{c)}
Smoking status (year 5)						
Never smoked	22,748 (52.1)	3,430 (43.8)	4,396 (48.3)	5,722 (52.9)	9,200 (57.6)	<0.0001 ^{b)}
Past smoker	19,336 (44.2)	4,017 (51.3)	4,357 (47.8)	4,700 (43.5)	6,262 (39.2)	
Current Smoker	1,624 (3.7)	382 (4.9)	356 (3.9)	388 (3.6)	498 (3.1)	
Alcohol servings per week, year 3 (mean (SD))	2.67 (5.23)	3.84 (6.65)	3.08 (5.50)	2.48 (4.71)	1.99 (4.45)	<0.0001 ^{c)}
Diagnosis of severe chronic diseases prior to year 5	14,708 (33.7)	2,472 (16.8)	2,937 (20.0)	3,597 (24.5)	5,702 (38.8)	<0.0001 ^{b)}

Values are n (%) unless otherwise specified

All variables collected at year 5 of follow-up unless otherwise specified

^{a)} ANOVA, ^{b)} Chi-square, ^{c)} Kruskal-Wallis

Depression = Center of Epidemiological Studies (CES-D) score >.06

BMI=body mass index (kg/m²); MET=metabolic equivalent tasks

Table 2 – Unadjusted and adjusted hazard ratios of fatal and non-fatal cardiovascular events by frequency of private spiritual activity

	Private spiritual activity				P for trend
	Never (REF.)	< Once per Week	Once or more per Week	Daily	
Cardiovascular events n (%)	408 (5.2)	499 (5.5)	612 (5.7)	1035 (6.5)	
Unadjusted HR	1.00	1.06 (0.93-1.21)	1.12 (0.99-1.27)	1.31 (1.17-1.47)	<.001
Model 1 HR	1.00	1.10 (0.96-1.25)	1.16 (1.02-1.31)	1.25 (1.11-1.40)	<.001
Model 2 HR	1.00	1.06 (0.93-1.21)	1.08 (0.96-1.23)	1.14 (1.01-1.28)	0.025
Model 3 HR	1.00	1.07 (0.94-1.22)	1.10 (0.97-1.25)	1.17 (1.04-1.31)	0.008
Model 4 HR	1.00	1.05 (0.92-1.19)	1.06 (0.94-1.21)	1.14 (1.01-1.28)	0.022
Model 5 HR	1.00	1.05 (0.92-1.19)	1.07 (0.94-1.21)	1.15 (1.03-1.30)	0.011
Model 6 HR	1.00	1.05 (0.92-1.19)	1.07 (0.94-1.21)	1.16 (1.03-1.30)	0.010

The analysis included women with complete data, cardiovascular mortality and cardiovascular events (n=43,708)

Hazard ratios (HR) are reported with 95% confidence intervals.

Model 1 – age-adjusted

Model 2 – adjusted for age, ethnicity, marital status, income, and education

Model 3 – model 2 plus physical activity (MET-hours), smoking status, alcohol servings/week

Model 4 – model 3 plus BMI, high cholesterol, hypertension, diabetes, and family history of myocardial infarction

Model 5 – model 4 plus physical functioning, satisfaction with quality of life, social support

Model 6 – model 5 plus CES-D and emotional well-being scores

Cardiovascular events = cumulative endpoint (angina, coronary and carotid revascularization, stroke, transient ischemic attack, congestive heart failure, peripheral arterial disease, and any adjudicated cardiovascular death)

CES-D= Center for Epidemiologic Studies depression scale

Figure legends

Figure 1. STROBE diagram.

CV=cardiovascular; WHI-OS=Women's Health Initiative Observational Study

Figure 2.

Adjusted associations between private spiritual activity and fatal and non-fatal cardiovascular events, overall and by presence/absence of severe chronic diseases prior to YR5. Severe chronic diseases included any cancer (except non-melanoma skin cancer), diabetes, liver disease, dialysis, asthma, emphysema, lupus, Alzheimer disease, multiple sclerosis, Parkinson disease, ALS

Table 3. Adjusted hazard ratios of cardiovascular events by frequency of spiritual practice, results of the subgroup analysis by presence/absence of chronic disease prior to YR5

	n	CV events n (%)	Private spiritual activity			
			Never	< Once per Week	Once or more per week	Daily
			REF	HR (95% CI)	HR (95% CI)	HR (95% CI)
No CDH	29,000	1,469 (5.1)	1.00	1.06 (0.90-1.26)	1.05 (0.89-1.24)	1.08 (0.93-1.26)
CDH	14,708	1,085 (7.4)	1.00	1.06 (0.86-1.31)	1.12 (0.92-1.37)	1.21 (1.01-1.45)

P value for differences in CV events by presence/absence of CDH= <0.0001

P value for interaction term added to model 6= 0.6449

The analysis included women with complete data (n=43,708)

Hazard ratios (HR) are reported with 95% confidence intervals.

Chronic disease history is defined as history of chronic disease reported before year 5 including cancer (except non-melanoma skin cancer), diabetes, liver disease, kidney dialysis, asthma, emphysema, lupus, Alzheimer disease, MS, Parkinson disease, ALS.

CV – cardiovascular

CDH – chronic disease history

Figure 1

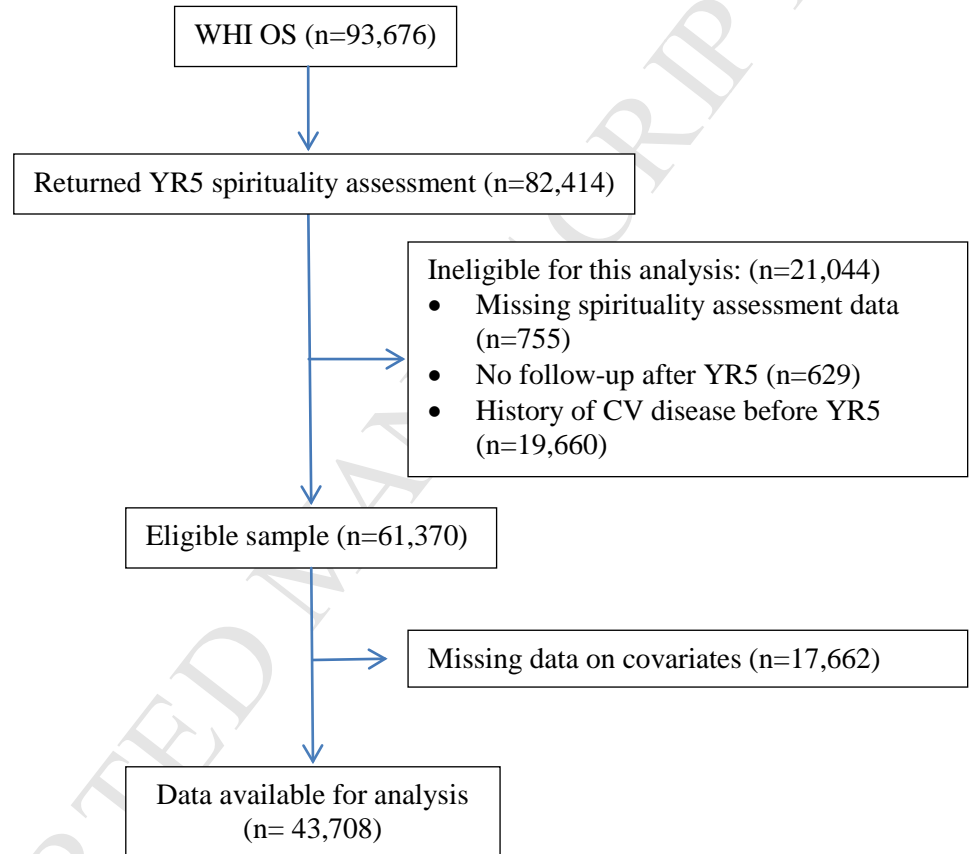


Figure 2

