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Is hyperlipidemia or its treatment associated with erectile dysfunction? Subtitle: Results from the Boston Area Community Health (BACH) Survey

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Abstract

Introduction—Studies and reports suggest that both hyperlipidemia and its pharmacologic treatment may lead to an increased risk of erectile dysfunction (ED).

Aims—Our objectives were to examine the association between 1) treated hyperlipidemia and ED; 2) untreated hyperlipidemia and ED.

Methods—Data from 1,899 men aged 30–79 were used from the Boston Area Community Health Survey of community-dwelling residents of Boston, MA, collected during 2002–2005 using an in-person interview, self-administered questionnaires, and a venous blood draw.

Main Outcome Measures—ED was measured using the Short Form International Index of Erectile Function. A case of treated hyperlipidemia was defined by use of anti-lipemics in the past month, while untreated hyperlipidemia was serum total cholesterol \geq 240 milligrams per deciliter with no anti-lipemic use. We estimated associations using odds ratios (ORs) and 95% confidence intervals (CIs) from multivariate logistic regression.

Results—Men with treated hyperlipidemia were older, had more comorbidities and used more medications compared to men with untreated hyperlipidemia or no hyperlipidemia. In multivariate models stratified by age and the presence of diabetes and/or cardiovascular disease (CVD), we saw no association between hyperlipidemia drug treatment and ED, except among younger men (<55) who had diabetes and/or CVD, where a strong association with an imprecise confidence interval was observed (OR=10.39, 95% CI: 3.25, 33.20). There was no significant positive association between untreated hyperlipidemia and ED in any multivariate model.

Conclusion—Lipid-lowering medications may be associated with ED among some men. The well-established benefits of lipid-lowering therapy should always be weighed against potential adverse effects.

Conflict of interest: Susan A. Hall is a former employee of GlaxoSmithKline and a former consultant to GlaxoSmithKline but has no equity interest in GlaxoSmithKline. Raymond Rosen is a consultant to, and investigator for Eli Lilly, Bayer Schering Pharma, and Pfizer. Martin Miner is a consultant to GlaxoSmithKline and has received funds for research from GlaxoSmithKline and Indevus. Peter Ganz has received speaker fees, fees for consulting and funds for research from Pfizer, and fees for consulting from GlaxoSmithKline. All other authors have no conflict of interest.

INTRODUCTION

Lipid regulators for hyperlipidemia and other indications were the most popular prescription drugs sold in the U.S. in 2007.¹ A statin, atorvastatin, was the top-selling medication of any class in the U.S. from 2000 to 2006², ³ and was used by 6.7% of adults in 2006.⁴ Because of their popularity as well as their indication for long-term use,⁵ the broader physiologic effects of statins are of special clinical and public health importance.

Both hyperlipidemia and erectile dysfunction (ED) are common problems in older men, and may be interrelated.⁶ As hyperlipidemia may contribute to ED by promoting endothelial dysfunction, it has been suggested that statins could be beneficial in alleviating ED through their pleiotropic effects in improving endothelial function.^{7, 8} The body of evidence for the complex interrelationships between hyperlipidemia, its treatment, and erectile dysfunction was recently reviewed.⁹ While the observation that hyperlipidemia is more prevalent among men with ED is largely consistent, existing evidence for a role for lipid-lowering treatments in the alleviation or promotion of ED is at present conflicting. In addition to case reports from drug safety monitoring and a systematic review,^{10, 11} one recent prospective clinical study of approximately 100 men with cardiovascular risk factors showed a marked worsening of erectile function six months following statin initiation.¹²

We examined these interrelationships in a population-based study of community-dwelling men. The objectives were 1) to describe the characteristics of those with treated or untreated hyperlipidemia and ED; 2) to estimate the association between treated and untreated hyperlipidemia and ED; and 3) to determine whether any observed association could be explained by potential confounding factors, such as cardiovascular risk factors.

METHODS

Design and data collection

The Boston Area Community Health (BACH) Survey is a population-based observational study of residents of Boston, Massachusetts. A two-stage, stratified cluster sampling design was used to recruit approximately equal numbers of participants to pre-specified age, race/ ethnic, and gender groups. Interviews were completed for 63.3% of eligible subjects, with a resulting sample of 2,301 men (age range 30–79) with 700 black, 766 Hispanic and 835 white participants. This analysis used baseline data collected between April 2002 and June 2005 after written informed consent. A non-fasting venous blood sample (20 ml) was also collected as close to awakening as possible. All protocols and informed consent procedures were approved by New England Research Institutes' Institutional Review Board. Further details regarding the study are available.¹³

Hyperlipidemia and ED definitions

A treated case of hyperlipidemia was defined as a user of any prescription anti-lipemic medications in the past month (statins and/or non-statin anti-lipemics), while an untreated case of hyperlipidemia was defined as no use of anti-lipemics and total serum cholesterol ≥240 milligrams per deciliter (in accordance with National Cholesterol Education Program guidelines to define high total cholesterol).¹⁴ Men who did not use anti-lipemics and had total serum cholesterol <240 were considered to have no hyperlipidemia. Serum cholesterol was measured enzymatically¹⁵ in a core laboratory certified by the National Heart, Lung, and Blood Institute/Centers for Disease Control and Prevention Lipid Standardization Program. The method combines the specificity of the enzymatic reaction with peroxidase/ phenol-4-aminophenazone indicator reaction, and was performed using the Hitachi 917 analyzer using reagents and calibrators from Roche Diagnostics (Indianapolis, IN). At cholesterol concentrations of 132.8 and 280.4 mg/dL, the day-to-day reproducibility

reflected by coefficient of variation (CV), was 1.7% (SD=2.4 mg/dL) and 1.6%, respectively.

ED was defined using the 5-item short form International Index of Erectile Function (IIEF-5), a self-administered, validated questionnaire.¹⁶ The score ranges from 5 to 25, with lower scores indicating poorer erectile function. In Tables 2⁻³, a case of treated ED was defined as any user of phosphodiesterase inhibitors, papaverine or prostaglandin, a case of untreated ED was defined as IIEF-5 score of ≤ 16 and no evidence of ED drug treatment, while no ED was defined as IIEF-5 score of ≥ 17 and no evidence of ED drug treatment. In Tables 5⁻⁶, a dichotomous definition of ED was created for use in bivariate analyses and logistic regression, using a cutoff of IIEF-5 ≤ 16 (combining mild to moderate, moderate, and severe categories to indicate the presence of ED regardless of treatment). In Figure 1, ED was displayed by severity as follows: severe (IIEF-5 score 5–7), moderate (8–11), mild to moderate (12–16), mild (17–21), and no ED (22–25).¹⁶.

Covariates

The choice of covariates was informed by prior analyses.¹⁷ Socioeconomic status (SES) was constructed as a function of standardized income and education variables for the Northeastern U.S.¹⁸ Body mass index (BMI) was calculated from interviewer-measured weight and height. Physical activity level was defined using the Physical Activity for the Elderly (PASE) scale.¹⁹ Persons reporting at least 5 of 8 symptoms on the abridged Center for Epidemiologic Studies Depression Scale were considered to have depression.²⁰ Cardiovascular disease (CVD) was a composite variable (Table 1). Other comorbidities were defined as a `yes' response to "Have you ever been told by a health care provider that you have or had....?".

Participants were asked to gather all medications used in the past four weeks for label recording, and were asked if they were taking drugs for specific indications such as asthma. Medications were coded using a modified form of the American Hospital Formulary Service (AHFS) Drug Pharmacologic Therapeutic Classification System.^{21 22} We considered use of medications thought to exacerbate symptoms of ED^{23–25} and created a count variable (0, 1, 2+) that included any use of beta blockers, selective serotonin reuptake inhibitors (SSRIs), tricyclic antidepressants, anti-psychotics, and diuretics. Similarly, we created a three-level count variable for use of other CVD medications.

Analytic sample and statistical analysis

Of 2,301 men in BACH, 402 men were excluded for missing blood, leaving 1899 in the analysis sample. To account for design effects and to allow generalizability to the Boston, MA, population, all analyses were weighted by the inverse of the probability of selection, and conducted using SUDAAN (version 9.0.1).^{26, 27} Missing data were replaced by plausible values using 25 multiple imputations; <1% were missing overall except for ED, where 13.3% of values were imputed. The distribution of covariates was examined within hyperlipidemia status (treated, untreated and none) and ED status (treated, untreated, and none) and tests for significant differences (p<0.05) were conducted using the chi-squared test and the Wald test. Mean IIEF-5 score and standard errors within subgroups were also examined; due to low frequencies, statistical testing was not emphasized in all groups.

We used multivariate logistic regression models to quantify the association between hyperlipidemia status and ED (defined as an IIEF-5 score of ≤16 regardless of ED treatment) using adjusted odds ratios (ORs) and 95% confidence intervals (CIs). We built parsimonious models that controlled confounding of the hyperlipidemia-ED relationship and full models containing all covariates of interest (regardless of statistical significance). All

models were backwards-selected and always adjusted for age. Confounders were identified using a change-in-estimate criterion²⁸ with a 10% as a threshold for confounding. Because we had very large differences by age between treated and untreated hyperlipidemia making it difficult to properly adjust for age, we also ran separate models among older (55+) and younger men (<55). In exploratory models, we identified a significant statistical interaction term (p<0.01) between the presence of comorbidity (diabetes and CVD) and hyperlipidemia status among men aged <55; consequently, we present all models stratified by both age and diabetes/CVD status.

Results

Men excluded from the analysis sample for missing blood were not different from included men on self-reported health, depression, smoking status, diabetes, body mass index, and SES (all overall p>0.12) but were more likely to be older, to be black and to have CVD (all overall p<0.03). Men with imputed IIEF-5 scores were more likely to be older, to have fair/ poor self-reported health, to be Hispanic, of low SES, and to have CVD (all overall p<0.04) but were not different on other variables above (all overall p<0.24). In the analysis sample, the overall prevalence of treated hyperlipidemia was 14.8%, while 13.0% of the sample had untreated hyperlipidemia and 72.2% had no hyperlipidemia. The prevalence of treated ED was 2.8%, while 18.8% had untreated ED (IIEF-5 score ≤16 and no use of ED medications) and 78.4% had no ED by our definition (IIEF-5 score of 17+). The characteristics of men by hyperlipidemia status and ED status are presented in Tables 1 and 2, respectively. There were substantial age differences by hyperlipidemia status (Table 1), while men with treated or untreated ED were older than men without ED (means of 52.6 and 55.3 vs. 45.5 years, respectively) (Table 2). Socioeconomic differences were stronger considering ED status compared to hyperlipidemia status, while the treated hyperlipidemia and treated ED groups had the highest proportion of white men. Men with treated or untreated hyperlipidemia were more likely to have a BMI \geq 30 compared to men without hyperlipidemia, while BMI category was not related to ED status.

Men with treated hyperlipidemia were more than twice as likely to report CVD, erectile dysfunction, hypertension, and diabetes compared to men with no hyperlipidemia or men with untreated hyperlipidemia (Table 1) and accordingly, were much more likely to use cardiovascular and antihypertensive medications (Table 3). Men with untreated hyperlipidemia reported a generally lower proportion of comorbidities and had lower use of medications compared to men without hyperlipidemia. Men with ED (treated or untreated) had higher proportions of all comorbidities in Table 2 and used more medications compared to men without ED (Table 3). In this comparison, differences were especially pronounced for statins, beta blockers, ACE inhibitors, and calcium channel blockers.

Table 4 shows mean erectile function scores (IIEF-5) within strata of age and hyperlipidemia status. IIEF-5 scores declined from 21.9 among those aged 30–39 to 14.9 among those aged 70–79 (p<0.001), indicating poorer function with increasing age. The mean for all ages among those treated for hyperlipidemia (17.0) was lower than those with untreated hyperlipidemia (21.6) and no hyperlipidemia (20.6) (p<0.001). Among the two youngest age groups, there were larger differences in mean score between men treated for hyperlipidemia and men without hyperlipidemia compared to the older age groups, with the most marked difference by treatment status among men 30–39 (13.7 among treated vs. 22.1 among those with no hyperlipidemia, p<0.001). At every age group, men with untreated hyperlipidemia had higher mean IIEF-5 scores than men without hyperlipidemia. Although there were few men treated with statins plus another class of anti-lipemics (n=14), these men had the lowest mean score (13.5) compared to 32 men treated with non-statin anti-lipemics only (19.5) (data not shown), or statins only (17.0).

We also examined severity of ED by hyperlipidemia treatment status (Figure 1). Overall, the prevalence and severity of ED increased with age; however, this pattern was not consistent by treatment status. Men aged 30–39 with treated hyperlipidemia had a higher prevalence of moderate or severe ED (58.8%) compared to men in the same age group who had untreated hyperlipidemia or no hyperlipidemia (1% and 2.7% had moderate or severe ED, respectively). Compared to men with untreated hyperlipidemia and no hyperlipidemia, men with treated hyperlipidemia had a greater prevalence of moderate or severe ED at every age group except 70–79.

Table 5 shows the prevalence of ED (combining mild to moderate, moderate and severe ED) by age and presence of CVD and/or diabetes by hyperlipidemia treatment status. Of all strata, the highest prevalence of ED was among men <55 who were being treated for hyperlipidemia (60.7%) and had CVD disease and/or diabetes. Among those 55+ who had CVD and/or diabetes, the prevalence of ED did not differ by hyperlipidemia status. Table 6 shows estimated ORs and 95% CIs for ED by hyperlipidemia status using logistic regression models. Among those aged <55 with diabetes and/or CVD, we observed a large OR for ED comparing treated hyperlipidemia to no hyperlipidemia (Model 1: OR=10.39, 95% CI: 3.25, 33.20). Although confidence intervals were imprecise, they excluded 1.00 and the association was persistent across modeling strategies. Considering those <55 without CVD or diabetes, there was no association for treated hyperlipidemia in any models but we observed an *inverse* association that persisted across modeling strategies between untreated hyperlipidemia and ED (Model 1: OR= 0.24, 95% CI: 0.10, 0.56). Considering those aged 55+ with diabetes and/or CVD, we saw no association between treated or untreated hyperlipidemia and ED in any model. Finally, among those age 55+ without CVD or diabetes, there were elevated ORs for treated hyperlipidemia and ED but confidence intervals included 1.00 (Model 1: OR=1.84, 95% CI: 0.76, 4.43), while ORs for untreated hyperlipidemia showed no association (Model 1: OR=0.89, 95% CI: 0.23, 3.40).

To consider whether the observed association between treated hyperlipidemia and ED among younger men with chronic illness would change if the analysis was restricted to particular anti-lipemic drug classes, we reran our Model 1s excluding those 46 men who were not exclusively using statins. In this statins-only analysis, the OR for treated hyperlipidemia was still substantial (OR=8.86, 95% CI: 2.69, 29.20), while the other estimates for treated hyperlipidemia remained statistically insignificant. The inverse association between untreated hyperlipidemia and ED among men aged <55 with no diabetes or CVD was unchanged in the statins-only analysis (OR=0.24, 95% CI=0.10, 0.56).

DISCUSSION

We examined the characteristics of men with treated and untreated hyperlipidemia, and estimated the association between use of lipid-lowering medications and ED, as well as the association between untreated hyperlipidemia and ED (compared to men without ED) in a population-based sample of community-dwelling men. We observed that men taking lipid-lowering drugs had lower mean erectile function scores compared to men with untreated hyperlipidemia and no hyperlipidemia. However, men on hyperlipidemia drug treatment were more likely to have comorbidities and take relevant medications, suggesting confounding by other risk factors for ED. In multivariate analyses that included other risk factors, we observed a large association between use of lipid-lowering medications and erectile dysfunction but only among men who were <55 and had diabetes and/or CVD. We were unable to examine all of our classes of lipid-lowering medications due to sample size, but repeating this analysis among those taking only statins did not change our conclusions. It is important to note that this result was based on relatively small numbers in the hyperlipidemia treatment group (n=60), as reflected in the imprecise confidence interval.

Finally, our multivariate results suggest lipid-lowering medications were not independently associated with ED among older men (55+), with or without comorbidities.

Our results are novel in that we find an association of lipid-lowering medications and ED only among younger men with comorbidities. In this group, we further considered whether testosterone levels were lower among men with ED on anti-lipemics compared to treated men without ED, but this was not explanatory (data not shown). Despite our multivariate adjustment including adjustment for additional medications for CVD as a proxy for CVD severity, we cannot rule out that the association we observed in this group could be due to interactions with other medications, or confounding by severity of pre-existing hyperlipidemia, ED, or other co-morbid disease as the severity of these at drug initiation was unknown. Because younger men with comorbidities who are prescribed anti-lipemics may be more mindful of their disease at a time when their peers may be healthier, they may have more of the psychological covariants that negatively affect sexual function. However, the possibility of a lipid-lowering drug effect among younger, sicker men should not be dismissed, and could be addressed in future clinical trials. Clinical trials of statins have not reported ED as an adverse event, except one that found no statistically significant difference in reporting between treatment and placebo.²⁹ Other authors have pointed out that ED is generally underreported and patients in trials were not specifically queried about ED.¹¹ A prior longitudinal study of men with existing cardiovascular risk factors found that statin initiation caused median IIEF-5 scores to fall from 21.0 to 6.5 at evaluation six months later. Although these men were generally older than BACH participants (mean age 61), the results support our findings by suggesting that statins may have a greater adverse impact on ED among men with comorbidities.¹² A clinical study of younger men that excluded those with CVD and diabetes but matched on other risk factors found an increased association of statins with ED (OR=1.46, 95% CI: 1.27, 1.68).³⁰ In contrast, a clinical trial of atorvastatin to improve lower urinary tract symptoms and benign prostatic enlargement among men aged 50+ included two questions from the IIEF but concluded there was no impact of atorvastatin on erectile function compared to placebo after 26 weeks.³¹ A small study of 74 hyperlipidemic men receiving atorvastatin reported a trend towards improvement in IIEF score over one year; however the difference in score from baseline was small (<2) and there was no comparison group.³² Prior epidemiologic studies have not found statins to be associated with ED in longitudinal³³ or cross-sectional analyses,²⁵ although the proportion of subjects taking statins in these studies was small.

We did not observe an independent effect of untreated hyperlipidemia on ED compared to men without hyperlipidemia, rather; we observed that younger men (<55) with untreated hyperlipidemia reported significantly less ED than men with no hyperlipidemia. While this finding remains unexplained, we note small numbers in this subgroup, and do not suggest that untreated hyperlipidemia could potentially reduce the risk of erectile dysfunction. Men in this group were not severely hyperlipidemic; it is possible that hyperlipidemia may take many years to induce the endothelial dysfunction thought to be related to ED. Supporting this is the observation that the "protective effect" of untreated hyperlipidemia was only observed among those <55.

Strengths of our study include a racially/ethnically and socioeconomically diverse sample of a broad age range, for whom comorbidities and medication information were systematically collected. We captured ED using a validated scale³⁴ and had the ability to consider use of other medications. Study limitations are also present. We had missing data for ED, although were able to use other information to impute ED status. In addition, we had small numbers in some subgroups, which led to imprecise confidence intervals. We did not collect a fasting blood sample, which may have caused misclassification of our untreated hyperlipidemia group, where we were relying on serum measures alone. In our study, there were differences

in the characteristics of those giving a blood sample versus not, and the observed proportion of men who had high cholesterol as measured in serum and/or who were using lipidlowering medications was 27.8%. However, this proportion is similar to that observed in the National Health Examination and Nutrition Survey (1999-2000): 24.9% for men aged 20-75.³⁵ In this cross-sectional analysis, we were unable to determine the severity of hyperlipidemia or the presence of ED before treatment, but when we excluded men on multiple types of hyperlipidemia treatment (a potential proxy for severity) our conclusions remained unchanged. Similarly, we were not able to control the severity of CVD, although we included current use of CVD medications as a severity proxy and we were able to consider variables other than lipids included in the validated Framingham coronary heart disease risk score (age, smoking, diabetes (self-reported) and blood pressure (self-reported). ^{36, 37} We do not capture recommendations for lifestyle changes as treatment for hyperlipidemia; our analysis only includes drug treatments. Because lipid-lowering medications are specific for hyperlipidemia, we do not feel we have overestimated the proportion treated with drugs, however. Finally, we have previously studied the effects of statins on testosterone in this population, and concluded there was no effect that was independent of comorbidity and body size, suggesting that statins are not inducing ED via reduction of testosterone.³⁸

CONCLUSION

Our results add to the available evidence that suggests that lipid-lowering agents, including statins, may be associated with erectile dysfunction among certain men (in our analysis, these were men <55 with CVD and/or diabetes), but our overall results may be reassuring for older men both with and without comorbidities, in whom no association was found in multivariate analyses. Given available treatments for erectile dysfunction and the body of clinical evidence for statins' reduction of major coronary events, all-cause mortality and other outcomes,³⁹ adverse effects of statins should always be weighed against the well-established benefits. The body of available evidence suggests that hyperlipidemia itself is an important part of the increased global cardiometabolic risk profile⁹ and when present in patients with ED should prompt management with diet and exercise as well as appropriate pharmacotherapy. Research is on-going to determine whether timely identification and aggressive treatment of ED and hyperlipidemia will lower cardiovascular morbidity and mortality.

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Figure 1. Severity of erectile dysfunction, by age and hyperlipidemia treatment status.

Table 1

Characteristics of those with treated, untreated and no hyperlipidemia* among men contributing blood samples in Boston Area Community Health Survey, 2002–2005, N=1899.

Continuous covariates †	Freated hyp (Me	oerlipidemia (n=290) an, Standard Error)	(n=2	Untreated hyperlipidemia 51) (Mean, Standard Error)	No hyperlipidemia (n=1358) (Mean, Standard Error)
Age		58.3 (1.0)		44.9 (1.0)	45.8 (0.5)
Median		59.8		42.4	43.2
Years education		14.2 (0.4)		14.5 (0.4)	15.0 (0.2)
Total cholesterol (mg/dl)		187.5 (4.1)		268.0 (2.5)	188.8 (1.2)
Categorical covariates †		Treated hyperlipidem	ia %	Untreated hyperlipidemia %	No hyperlipidemia %
Socioeconomic status					<u> </u>
	Low		32.8	17.5	23.2
	Middle		36.9	58.4	49.7
	High		30.2	24.0	27.1
Race/ethnicity					
	Black		23.1	20.0	26.4
	Hispanic		7.5	16.0	13.6
	White		69.4	64.0	60.0
Health insurance					
	Private		67.5	69.3	65.9
	Public		28.0	12.9	17.4
	None		4.5	17.8	16.8
Current smoker					
	Never		31.8	37.5	49.8
	Former		44.5	28.7	25.7
	Current		23.7	33.7	24.5
Alcohol use					
	None		39.1	19.0	24.8
	<1/day		41.2	34.4	41.0
	1-2/day		15.9	31.3	24.9
	3+/day		3.9	15.3	9.2
Physical activity score (PASE)	1				
Le	ow (<100)		40.8	21.3	23.1
Medium ((100–249)		41.0	47.9	49.7
Hi	gh (250+)		18.2	30.8	27.2
BMI categories					
	<25.0		16.6	22.2	29.6
:	25.0–29.9		37.8	36.4	40.4
	30.0+		45.6	41.4	29.9
Cardiovascular disease \ddagger			46.0	12.7	15.6
Hypertension			58.2	16.4	21.7
Diabetes (Type I and/or Type	II)		31.7	18	65

Categorical covariates †	Treated hyperlipidemia %	Untreated hyperlipidemia %	No hyperlipidemia %
Depression	15.0	10.0	14.5
Erectile dysfunction (IIEF-5 score ≤16)	44.4	8.9	17.9
Self-reported history of hyperlipidemia	85.3	37.7	15.8

**Treated hyperlipidemia was defined as taking prescription anti-lipid medication, untreated hyperlipidemia was measured serum total cholesterol of \geq 240 mg/dL without taking prescription anti-lipid medication, and no hyperlipidemia was defined as no use of anti-lipid medications and total serum cholesterol <240.

 † Variable names marked with bold were significantly different (P<0.05) across groups by the chisquare test of heterogeneity or Wald F test (for continuous variables). All estimates were weighted by the inverse of the probability of being sampled. Percents shown are column percents.

 ‡ Any history of coronary artery bypass surgery or angioplasty, heart attack, angina, having a pacemaker, congestive heart failure, transient ischemic attack, stroke, carotid artery surgery, intermittent claudication, surgery or angioplasty for arterial disease of the leg, pulmonary embolism, aortic aneurysm, heart-rhythm disturbance, deep vein thrombosis, Reynaud's disease or peripheral vascular disease.

Table 2

Characteristics of those with treated, untreated and no erectile dysfunction (ED)^{*} among men contributing blood samples in Boston Area Community Health Survey, 2002–2005, N=1899.

Continuous covariates † Treat	ted ED (n=40) (Me Standard Err	an, Untreated	l ED (n=472) (Mean, Standard Error)	No ED (n=1387) (Mean, Standard Error)
Age	52.6 (2	2.3)	55.3 (1.1)	45.5 (0.5)
Median	5	2.3	56.2	43.2
Years education	17.3 (0).6)	12.9 (0.4)	15.2 (0.2)
Total cholesterol (mg/dl)	195.5 (7	7.0)	188.8 (2.8)	201.5 (1.8)
Categorical covariates [†]	Treated ED %	Untreated ED %	No ED %	
Socioeconomic status				
Low	6.4	49.4	18.4	
Middle	40.2	36.8	52.1	
High	53.4	13.8	29.5	
Race/ethnicity				
Black	17.6	31.1	23.9	
Hispanic	10.7	16.8	12.2	
White	71.7	52.1	64.0	
Health insurance				
Private	82.9	47.1	70.6	
Public	12.6	38.1	13.8	
None	4.5	14.8	15.5	
Current smoker				
Never	53.3	33.7	48.1	
Former	29.1	37.7	26.7	
Current	17.6	28.6	25.1	
Alcohol use				
None	9.9	43.2	22.7	
<1/day	49.1	31.5	42.0	
1–2/day	37.4	16.7	25.8	
3+/day	3.5	8.6	9.5	
Physical activity score (PASE)				
Low (<100)	16.1	46.1	20.9	
Medium (100–249)	65.5	40.5	49.4	
High (250+)	18.4	13.4	29.7	
BMI categories				
<25.0	35.4	23.1	27.3	
25.0–29.9	41.5	38.8	39.6	
30.0+	23.2	38.1	33.1	
Cardiovascular disease \ddagger	43.7	34.7	15.2	
Hypertension	53.7	44.5	21.1	
Diabetes (Type I and/or Type II)	32.0	19.4	6.5	

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Categorical covariates †	Treated ED %	Untreated ED %	No ED %
Depression	17.3	24.6	11.4
Self-reported history of hyperlipidemia	53.8	42.8	24.7
Treated hyperlipidemia ${}^{\&}$	48.4	29.2	10.2
Untreated hyperlipidemia ${}^{\&}$	5.5	5.8	15.0
No hyperlipidemia §	46.0	65.1	74.8

Treated ED was defined as any use of phosphodiesterase inhibitors, papaverine or prostaglandin, untreated ED was defined as an IIEF-5 score of \leq 16 and no evidence of ED drug treatment, while no ED was defined as an IIEF-5 score \geq 17 and no evidence of ED drug treatment.

 † Variable names marked with bold were significantly different (P<0.05) across groups by the chisquare test of heterogeneity or Wald F test (for continuous variables); all estimates were weighted by the inverse of the probability of being sampled. Percents shown are column percents.

[‡]Any history of coronary artery bypass surgery or angioplasty, heart attack, angina, having a pacemaker, congestive heart failure, transient ischemic attack, stroke, carotid artery surgery, intermittent claudication, surgery or angioplasty for arterial disease of the leg, pulmonary embolism, aortic aneurysm, heart-rhythm disturbance, deep vein thrombosis, Reynaud's disease or peripheral vascular disease.

 $^{\$}$ Treated hyperlipidemia was defined as taking prescription anti-lipid medication, untreated hyperlipidemia was measured serum total cholesterol of \geq 240 mg/dL without taking prescription anti-lipid medication, and no hyperlipidemia was defined as no use of anti-lipid medications and total serum cholesterol <240.

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Table 3

Prevalence of use of medications by hyperlipidemia* and erectile dysfunction (ED)[†] status among men contributing blood samples in Boston Area Community Health Survey, 2002–2005, N=1899.

Statins $d e$ Non-etotin ontilinomize e	(defina nyperupua (n=25) %	l) % (n=1358)	% (n=40) %	Untreated ED (n=472) %	No ED (n=1387) %
Non-statin antilinamias l	89.8	0	0 48.4	26.4	8.9
	17.4	0	0 5.2	5.9	1.7
Beta blockers d e	30.8	1.7	5.2 13.4	25.1	5.3
Calcium channel blockers d e	19.7	4.1	3.6 13.9	14.1	3.9
ACE inhibitors $d e$	37.9	3.9	5.5 42.4	22.6	6.9
Angiotensin II receptor agonists d	6.2	0.8	0.9 4.1	3.2	1.2
Loop diuretics d e	5.6	0.2	1.3 2.3	6.6	0.6
Thiazide diuretics $d e$	12.7	2.9	1.0	9.6	4.4
Miscellaneous diuretics d e	6.0	0.7	0.6 (3.0	1.1
Any anti-hypertensive drug d e	50.0	6.1 1	2.6 48.6	33.5	12.3
SSR1 anti-depressants ^e	10.6	8.0	.0 0.4	13.0	9.1
Tricyclic anti-depressants e	9.8	1.9	1.6 14.6	7.8	1.3
Anti-psychotics ^e	4.3	2.1	3.5 0.4	6.7	2.7
Anti-ED drugs ^{d e}	9.1	1.2	1.8 100	0	0
Alpha blockers or 5-alpha reductase d^e inhibitors	7.9	0.4	2.7 7.9	6.4	2.2

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n D á medication, and no hyperlipidemia was defined as no use of anti-lipid medications and total serum cholesterol <240. † Treated ED was defined as any use of phosphodiesterase inhibitors, papaverine or prostaglandin, untreated ED was defined as an IIEF-5 score of \leq 16 and no evidence of ED drug treatment, while no ED was defined as an IIEF-5 score ≥ 17 and no evidence of ED drug treatment.

 \sharp Estimates were weighted by the inverse of the probability of being sampled. Percents shown are column percents.

 $^{\$}$ Variable names marked with bold were significantly different (P<0.05) across groups by the chisquare test of heterogeneity

 $d_{\rm represents}$ statistical significance for hyperlipidemia comparison

 e represents statistical significance for ED comparison.

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Mean (standard error) IIEF-5 score for erectile dysfunction by age group and hyperlipidemia status^{*} among men contributing blood samples in the Boston Area Community Health Survey, 2002–2005, N=1899.

Age group	Total	Treated hyperlipidemia (any treatment) st	Treated with statins ${}^{\sharp}$ only	Untreated hyperlipidemia	No hyperlipidemia
	Mean [†] (SE)	$\operatorname{Mean} \mathring{r} (\operatorname{SE})$	Mean [†] (SE)	Mean [†] (SE)	Mean [†] (SE)
Total	20.2 (0.2)	17.0 (0.6)	17.0 (0.6)	21.6 (0.4)	20.6 (0.2)
Z	1899	290	244	251	1358
30–39	21.9 (0.3)	13.7 (2.4)	15.3 (2.9)	22.5 (0.6)	22.1 (0.2)
Z	512	11	2	72	429
40-49	21.0 (0.2)	19.3 (0.9)	19.1 (0.9)	22.3 (0.5)	21.0 (0.3)
Z	554	62	54	86	406
50-59	19.4 (0.5)	18.1 (1.0)	17.7 (1.2)	20.2 (1.0)	19.6 (0.6)
Z	436	81	65	55	300
69-09	17.6 (0.6)	16.7 (0.9)	16.6 (1.0)	18.9 (1.5)	18.1 (0.8)
N	260	91	77	27	142
<i>70–79</i>	14.9 (0.8)	15.2 (1.3)	15.5 (1.4)	16.4 (2.8)	14.5 (1.0)
Z	137	45	41	11	81

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taking prescription anti-lipid adist

 $\mathring{\tau}_{\rm All}$ means were weighted by the inverse of the probability of being sampled.

 t^{j} Statins were atorvastatin, fluvastatin, lovastatin, pravastatin, rosuvastatin, or simvastatin.

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Table 5

Prevalence of erectile dysfunction $(ED)^*$ by hyperlipidemia^t, age and comorbidity status among men contributing blood samples in Boston Area Community Health Survey, 2002–2005, N=1899.

Subgroup	Total N	Overall	Treated hyperlipidemia	Untreated hyperlipidemia	No hyperlipidemia
Age <55	1291	14.1%	39.5%	3.9%	13.5%
N with ED		234	41	21	172
Age 55+	608	38.1%	47.4%	29.7%	34.0%
N with ED		257	91	25	141
Age <55 and CVD [§] and/or diabetes	239	23.1%	60.7%	3.8%	13.0%
N with ED		99	29	4	34
Age <55, no CVD or diabetes	1052	12.1%	13.7%	3.9%	13.6%
N with ED		168	12	17	139
Age 55+ and CVD and/or diabetes	267	51.4%	52.8%	50.4%	49.9%
N with ED		143	65	10	68
Age 55+, no CVD or diabetes	341	27.5%	36.5%	22.5%	25.7%
N with ED		114	25	15	73

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7. medication, and no hyperlipidemia was defined as no use of anti-lipid medications and total serum cholesterol <240.

 \sharp Prevalence percentages were weighted by the inverse of the probability of being sampled.

claudication, surgery or angioplasty for arterial disease of the leg, pulmonary embolism, aortic aneurysm, heart-rhythm disturbance, deep vein thrombosis, Reynaud's disease or peripheral vascular disease. ⁸Any history of coronary artery bypass surgery or angioplasty, heart attack, angina, having a pacemaker, congestive heart failure, transient ischemic attack, stroke, carotid artery surgery, intermittent

Table 6

Odds ratios and 95% confidence intervals from multivariate logistic regression models for ED^{*} and hyperlipidemia status^t by age and comorbidity subgroups among men contributing blood samples in Boston Area Community Health Survey, 2002-2005, N=1899.

Subgroup	Ν	Model	Treated hyperlipidemia	Untreated hyperlipidemia	No hyperlipidemia (referent)
	730	1^{\sharp}	10.39 (3.25, 33.20)	0.59 (0.12, 2.97)	1.00
Age <55 and UVD and/or diabetes	607	28	10.65 (3.11, 36.45)	0.40 (0.07, 2.22)	1.00
	1050	11	1.03 (0.36, 2.92)	0.24 (0.10, 0.56)	1.00
Age <25, no UVD of diabetes	7001	2§	1.02 (0.35, 3.02)	0.24 (0.10, 0.57)	1.00
		1//	1.10 (0.48, 2.52)	1.18 (0.30, 4.60)	1.00
Age 55+ and CVD ana/or madetes	107	2§	1.04 (0.43, 2.55)	$1.10\ (0.29, 4.14)$	1.00
A con 55 - and CVD on dishered	341	1**	1.84 (0.76, 4.43)	0.89 (0.23, 3.40)	1.00
Age 33+, IIO CAD OI UIADEICS	140	2§	1.84 (0.68, 5.02)	0.85 (0.23, 3.15)	1.00

, having a pacemaker, congestive aortic aneurysm, heart-rhythm disturbance, deep vein thrombosis, Reynaud's disease or peripheral vascular disease.

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7. medication, and no hyperlipidemia was defined as no use of anti-lipid medications and total serum cholesterol <240.

Z parsimonious model solution controlled for age, socioeconomic status, insurance status, physical activity, depression, alcohol use, and use of other CVD drugs and other drugs that contribute to ED

 $\frac{\delta}{8}$ Full model solution controlled for age, socioeconomic status, race/ethnicity, insurance, physical activity, body mass index category, depression, health status, pack-years smoking & use of CVD drugs & drugs that may cause ED.

Parsimonious model solution controlled for age, socioeconomic status, and use of other CVD drugs.

n Parsimonious model solution controlled for age, socioeconomic status, body mass index category, depression, and use of other CVD drugs.

** Parsimonious model solution controlled for age, socioeconomic status, depression, and use of other drugs that contribute to ED.