Statin Use in the U.S. for Secondary Prevention of Cardiovascular Disease Remains Suboptimal

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Background: Atherosclerotic cardiovascular disease (ASCVD) remains the leading cause of mortality in the United States. The purpose of this study is to examine the rates of statin use for secondary prevention of ASCVD events in the United States over the last decade and determine whether disparities in the treatment of ASCVD still persist among women and racial/ethnic minorities.

Methods: We conducted a trend analysis using data from 2008 through 2016 to describe age-adjusted trends in the use of statins for secondary prevention using the Medical Expenditure Panel Survey. We also conducted a multivariable logistic regression analysis to determine whether sociodemographic characteristics are associated with statin use during the 3 years that followed the publication of the 2013 American College of Cardiology/American Heart Association (ACC/AHA) guideline (2014 through 2016).

Results: The prevalence of statin use among those with a history of ASCVD remained unchanged from 2008 through 2016. In 2014 to 2016, more than 40% of those aged 40 years and older with a history of ASCVD did not use statins, corresponding to approximately 9.5 million Americans. Increasing age and having been diagnosed with high cholesterol (odds ratio [OR], 6.22; P < .001) were associated with higher odds of statin use while being female (OR, 0.65; P < .001) or Hispanic (OR, 0.69; P = .011) were associated with lower odds of statin use.

Conclusions: Our study found there was no increase in the national rates of statin use following the ACC/AHA 2013 secondary prevention guideline and the availability of generic statins. Significant gender and ethnic disparities in ASCVD treatment remained in the United States. (J Am Board Fam Med 2019; 32:807-817.)

Keywords: Atherosclerosis, Cardiovascular Diseases, Chronic Disease, Hypercholesterolemia, Hyperlipidemia, Logistic Models, Preventive Medicine, Secondary Prevention, Statins, Surveys and Questionnaires

Atherosclerotic cardiovascular disease (ASCVD) remains the leading cause of mortality in the United States despite a strong emphasis on preven-

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tion, accounting for more than 800,000 deaths in 2017. The 2013 American College of Cardiology/ American Heart Association (ACC/AHA) Guideline on the Treatment of Blood Cholesterol to Reduce Atherosclerotic Cardiovascular Risk in Adults recommends statin use for secondary prevention for adults ≤75 years of age with a prior history of ASCVD, specifically recommending that "high-intensity statin therapy should be initiated for those with clinical ASCVD who are not receiving statin therapy" or intensity raised for those already on a lower dose unless contraindicated (eg, history of statin intolerance).² The ACC/AHA guideline also recommends moderate-intensity statin use be considered for secondary prevention in those >75 years of age with a history of ASCVD.

These recommendations were based on numerous studies showing that statins are effective in reducing low-density lipoprotein cholesterol levels, and the risk of death and recurrent coronary and cardiovascular events in those with a history of ASCVD, even in older adults.^{3–7}

However, despite this proven effectiveness and the resulting ACC/AHA guideline, several studies have shown persistent underuse of statins in this high-risk population.^{8–14} Of these, we are aware of only a few studies that extend beyond the year that the guideline was published and even then only include narrowly defined populations (eg, only privately insured and/or Medicare patients, those covered by a single insurer, those within 30 days postmyocardial infarction (MI).^{10,11,13,14}

We examined nationally representative data for the years 2008 through 2016 to determine whether there has been an increase in use of statins in persons with a history of ASCVD following the publication of the ACC/AHA guideline in 2013. Prior studies have tended to focus on narrower samples whereas ours contains a nationally representative sample of adults with any history of ASCVD, the wider population targeted by the ACC/AHA guideline. We also assessed whether individual sociodemographic characteristics are associated with statin use for individuals with a history of ASCVD. Given known disparities in ASCVD treatment among women and racial/ethnic minorities, 15-18 we wanted to understand whether these disparities persisted using the most current data available. We also capture those who are on Medicaid (only or dually eligible for Medicare) or uninsured, populations that are often missed in claims-based analyses that typically include only those with Medicare and/or private coverage. This study assesses whether rates of statin use for secondary prevention are improving and examines disparities to identify opportunities for improvement.

Methods

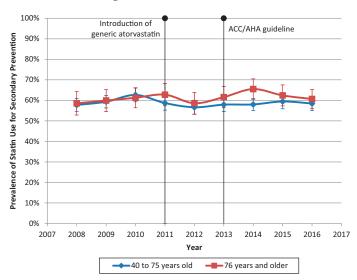
We used the Medical Expenditure Panel Survey (MEPS), a nationally representative survey of the civilian noninstitutionalized population conducted annually by the Agency for Health Care Research and Quality since 1996. ¹⁹ Our analytic sample contains 6084 observations representing 23.4 million people. Our analysis had 2 main components. First, we conducted a trend analysis using data from 2008

through 2016 to describe age-adjusted trends in the use of statins among adults with a history of diagnosed ASCVD. We calculated prevalence of statin use among those with a history of ASCVD in the 40 to 75 years and 76 years and older age groups for each year. We omitted adults aged 21 to 39 years, who are also potentially covered by the ACC/AHA guidelines, because prevalence of ASCVD was low. Then, we conducted a multivariable logistic regression analysis of whether sociodemographic characteristics were associated with statin use among those with a history of ASCVD using pooled data from 2014 through 2016, the 3 years that follow the publication of the 2013 ACC/AHA guidelines. ^{15–18}

Statin use for secondary prevention was defined as 1 or more prescriptions filled within the calendar year among our population of interest, those aged 40 and older with a history of diagnosed ASCVD.²⁰ We identified generic and brand-name statin fills from all prescription drug fills reported using Cerner Multum drug codes and/or medication names. We ascertained history of ASCVD events, to identify the population for which secondary prevention is recommended, from household respondents' answers to the question of whether a doctor or other professional had ever told them they had coronary heart disease, angina, a heart attack, or stroke. Regression models included sociodemographic characteristics (age, sex, race/ethnicity, education, marital status), health status (SF-12 physical component score), health insurance status (Medicare; any private; any Medicaid; uninsured), and geographic area (Census region, living in a Metropolitan Statistical Area). We also examined the association between additional ASCVD risk factors and statin use, including diagnosed hypertension, high cholesterol, and diabetes, as well as smoking status.21

We conducted all analyses with Stata/MP 15 (StataCorp, LLC, College Station, TX) using appropriate sample weights and survey estimation techniques to account for the complex sampling design. Multiple imputation was used to fill in a small number of missing values (at most 2.5% of sample) for demographic (ie, marital status, education) and health measures (ie, 12-Item Short Form Health Survey (SF-12) Physical Component Summary (PCS), ASCVD risk factors, household respondent-reported diagnosis), matching the approach used in a recently published study using MEPS to estimate the impact of US Preventive

Figure 1. Statin use among adults with history of diagnosed Ascvd, 2008 to 2016. Source: Agency for Health Care Research and Quality, Medical Expenditure Panel Survey, Full-Year Consolidated Files, 2008 to 2016. Note: All years were age-adjusted to match the 2016 age distribution.



Services Task Force (USPSTF) recommendations on statin use and costs for primary prevention.²³

Results

Trends in Statin Use for Secondary Prevention

Figure 1 shows the age-adjusted US trends for statin use among adults with a history of diagnosed ASCVD aged 40 to 75 years old and aged 76 years and older, respectively, over the 9-year period from 2008 to 2016. The prevalence rate of diagnosed ASCVD did not change significantly over this period (see Appendix). It also shows the years in which the generic statin, atorvastatin, was brought to market in 2011, which may increase utilization through lower costs. Rates of statin use for secondary prevention of ASCVD remained essentially unchanged over this time frame. The observed prevalence of statin use among those with a history of ASCVD went from 57.8% in 2008 to 58.4% in 2016 for 40-to-75-year-olds and from 58.6% in 2008 to 60.7% in 2016 for those 76 years and older, but these changes were not statistically significant. We also see no significant changes following the introduction of the ACC/AHA guideline in 2013. In Table 1, we focus on the postguideline period of 2014 to 2016 to describe the population size and characteristics of those using statins for secondary prevention.

Statin Use and Patient Characteristics

Table 1 shows the sociodemographic characteristics and risk factors for the study population aged

40 years and older with a history of ASCVD. Our MEPS analytic sample included approximately 6000 person-years, representing an estimated average of 23.4 million individuals with ASCVD per year of our study period. An estimated 16.6 million (71%) were aged 40 to 75 years with the remaining 6.7 million (29%) aged 76 years and older. A narrow majority of our weighed sample was male (53.7%). Non-Hispanic whites represented nearly three-quarters of the sample (73.5%) followed by non-Hispanic African Americans (11.0%) and Hispanics (9.5%). Nearly all had some form of health insurance (eg, Medicare, Medicaid, or private insurance) with over two-thirds having Medicare (69.7%) and another one-fifth having private insurance (20.4%). Only 3.9% reported being uninsured. More than three-quarters (76.2%) reported history of coronary heart disease, nearly 40% reported a prior stroke (39.5%), and almost a third (31.7%) reported having diabetes. These are not mutually exclusive, respondents could be represented in one or more categories. Nearly all respondents in this population reported having a nonemergency department usual source of care (92.4%).

Current Statin Use

Table 2 shows the proportion of the total population with ASCVD who were taking statins. Among adults aged 40 years and older with ASCVD, nearly 14 million (59.4%) were using statins on average from 2014 through 2016 after the ACC/AHA rec-

Table 1. Demographic and Health Characteristics of Adults Aged 40 Years and Older with History of Diagnosed ASCVD, 2014 to 2016

Characteristic	n	Weighted %	95% CI
Age.in years			
40 to 49	595	8.7	(7.6 to 9.8)
50 to 59	1,247	17.3	(15.9 to 18.7
60 to 64	804	12.6	(11.4 to 13.7
65 to 69	871	15.2	(13.7 to 16.7
70 to 75	946	17.4	(15.8 to 19.0
76 to 79	513	9.2	(8.1 to 10.2)
80 to 84	562	9.9	(8.9 to 10.9)
85 and above	546	9.8	(8.6 to 11.0)
Sex			, , ,
Women	3,039	46.3	(44.5 to 48.1
Men	3,045	53.7	(51.9 to 55.5
Race/ethnicity			
Hispanic	1,124	9.5	(8.0 to 11.0)
Non-Hispanic, white	3,198	73.5	(71.3 to 75.7
Non-Hispanic, African American	1,320	11.1	(9.7 to 12.6)
Non-Hispanic, Asian	254	2.7	(2.0 to 3.5)
Non-Hispanic, Other	188	3.1	(2.2 to 4.0)
Marital status			
Unmarried	3,159	44.5	(42.3 to 46.7
Married	2,925	55.5	(53.3 to 57.7
Education			
Did not complete high school	2,322	31.1	(29.3 to 32.8
High school graduate or GED	1,304	22.4	(20.7 to 24.1
Some college	1,407	24.9	(23.1 to 26.7
College degree or higher	991	21.0	(19.2 to 22.9
Insurance coverage			
Medicare	4,064	69.7	(67.8 to 71.6
Private	1,124	20.4	(18.7 to 22.1
Medicaid	568	6.0	(5.1 to 6.9)
Uninsured	328	3.9	(3.2 to 4.6)
Census region			
Northeast	1,003	17.8	(15.8 to 19.8
Midwest	1,231	22.5	(20.2 to 24.9
South	2,628	40.8	(38.1 to 43.5
West	1,222	18.8	(17.0 to 20.7
MSA			
MSA	5,024	81.3	(78.1 to 84.6
Non-MSA	1,060	18.7	(15.4 to 21.9
History of cardiovascular disease			
CHD to MI to or angina	4,555	76.2	(74.6 to 77.9
Stroke	2,538	39.5	(37.8 to 41.2

Continued

Table 1. Continued

Characteristic	n	Weighted %	95% CI
ASCVD risk factors			
Doctor ever told high cholesterol	4,504	75.1	(73.4 to 76.8)
Doctor ever told had hypertension	4,982	80.9	(79.4 to 82.4)
Doctor ever told had diabetes	2,122	31.7	(29.8 to 35.6)
Current smoker	1,020	16.8	(15.1 to 18.4)
Usual source of care			
No	531	7.9	(6.8 to 8.9)
Yes	5,553	92.1	(91.1 to 93.2)

Source: Agency for Healthcare Research and Quality, Medical Expenditure Panel Survey 2014 to 2016.

ASCVD, atherosclerotic cardiovascular disease; CHD, coronary heart disease; CI, confidence interval; ED, emergency department; GED, General Equivalency Diploma; MI, myocardial infarction; MSA, Metropolitan Statistical Area.

ommendation for statin use for secondary prevention was published. Among the 16.6 million persons with ASCVD who are aged 40 to 75 years, 9.7 million (58.3%) were using statins while among the 6.7 million persons with ASCVD aged 76 years or older, 4.2 million (62.3%) were using statins, indicating that older adults in this at risk population are using statins at a higher rate.

Table 3 shows the adjusted odds of statin use associated with various sociodemographic and health status characteristics. Statin use for secondary prevention was more likely with increasing age, plateauing in the 65-to-84-years age range before beginning to drop off. All persons age 50 years and higher had significantly higher odds of statin use relative to those aged 40 to 49 years with a history of ASCVD. The age groups with the highest odds of statin use were among 76-to-79-year-olds (odds ratio [OR], 3.43; 95% CI, 2.14 to 5.49) and 65-to-69-year-olds (OR, 3.36; 95% CI, 2.15 to 5.22). Female sex (OR, 0.65; 95% CI, 0.55 to 0.77) was associated with decreased odds of using statins for secondary prevention. Hispanic ethnicity was associated significantly lower odds of reporting statin use (OR, 0.69; 95% CI, 0.52 to 0.92), nearly a third lower than that of non-Hispanic whites. Married persons were more likely to be taking statins for secondary prevention (OR, 1.26; 95% CI, 1.02 to 1.57). All education levels (high school diploma or GED, some college, college degree or higher) had higher estimated odds of statin use than those who did not complete high school; however, only the

Table 2. Statin Use in Adults with History of Diagnosed ASCVD by Age Group, 2014 to 2016

		Total Population	Statin Users	
	N	Weighted Population, Millions (95% CI)	Weighted Users, Millions (95% CI)	Weighted % (95% CI)
Age 40 years and older	6,084	23.4 (22.1 to 24.6)	13.9 (13.0 to 14.8)	59.4 (57.4 to 61.5)
Age 40 to 75 years	4,463	16.6 (15.7 to 17.6)	9.7 (9.0 to 10.4)	58.3 (55.9 to 60.6)
Age 76 years and older	1,621	6.7 (6.2 to 7.3)	4.2 (3.8 to 4.7)	62.3 (59.0 to 65.7)

ASCVD, atherosclerotic cardiovascular disease; CI, confidence interval.

Source: Agency for Healthcare Research and Quality, Medical Expenditure Panel Survey, 2014 to 2016.

result for some college was statistically significant (OR, 1.39; 95% CI, 1.08 to 1.80). Among risk factors for ASCVD, those who reported ever being told by a doctor that they had high cholesterol had the highest odds of being on a statin (OR, 6.22; 95% CI, 5.02 to 7.72). Those who were ever told they had diabetes (OR, 1.51; 95% CI, 1.24 to 1.85) or hypertension (OR, 1.33; 95% CI, 1.08 to 1.65) also had significantly higher odds of being on a statin; however, current smoking was not associated with increased odds of statin use after controlling for other risk factors and socioeconomic characteristics. A 1-point increase in SF-12 Physical Component Score (a scale from 0 to 100), indicating better health, was associated with slightly higher odds of statin use (OR, 1.01; 95% CI, 1.00 to 1.02). Having a usual source of care, regardless of setting, was associated with a near doubling of odds of statin use for secondary prevention (OR, 1.87; 95% CI, 1.37 to 2.56). An alternate model (not shown) that split usual sources of care by setting found no significant differences in odds of statin use between those with a nonemergency department usual source of care (hospital OR, 1.92; nonhospital OR, 1.79; yes but setting unknown OR, 1.74) so we did not include that in the model presented. There were no significant differences in the odds of statin use by health insurance status, Census region, or living in an Metropolitan statistical area (MSA).

Discussion

Our study is the first to show national trends in statin use following the AHA/ACC 2013 guideline. We find nearly flat trends in statin use for secondary prevention, despite the introduction of a generic statin over the last decade (eg, generic atorvastatin in 2011). Unfortunately, gaps in recommended ASCVD services are not unique to statins for secondary prevention or other recommended clinical services.^{24,25} For example, less than 35% of patients who reported having an acute myocardial infarction received cardiac rehabilitation. 26,27

We find a gender disparity in statin use for secondary prevention, consistent with prior research, 28,29 finding that women have approximately 35% lower odds of statin use after accounting for other demographic and health factors. Though the prevalence of ASCVD is higher in men, ASCVD mortality is actually higher among women and there is no evidence that statins are less effective for secondary prevention or not tolerated as well by women as men.^{29–32} People of Hispanic ethnicity had approximately 30% lower odds of receiving statins for secondary prevention.33-36 We also found that adults who had a usual source of care, were married, and had higher education had greater odds of statin use, which is consistent with prior research.^{37–40}

Understanding barriers to statin use—including patient, clinician, and delivery system factors as well as costs—is important when identifying strategies to improve quality of care. Statins are now available to many Americans at little-to-no out-ofpocket cost regardless of whether they are being used for primary or secondary prevention. 41,42 Many insurers had already lowered or eliminated cost sharing for statins for those with a history of ASCVD. The Patient Protection and Affordable Care Act of 2010 (ACA) required that private insurers provide first-dollar coverage for services and medications with an "A" or "B" grade from the USPSTF without cost-sharing for primary prevention. 43-45 Some insurers responded by reducing the out-of-pocket costs for certain generic statins to zero regardless of indication.46 Lower out-ofpocket costs have been associated with increased prescription fills for and adherence to statin ther-

Table 3. Odds of Current Statin Use and Sociodemographic Characteristics, Adults Aged 40 Years and Older with History of Diagnosed ASCVD, 2014 to 2016 (n = 6.039)

Characteristic	Odds Ratio	(95% CI)	P-Value
Age, in years			
40 to 49 (reference)	1.00	_	_
50 to 59	1.60	(1.14 to 2.26)	.007
60 to 64	2.43	(1.66 to 3.56)	<.001
65 to 69	3.36	(2.15 to 5.22)	<.001
70 to 75	3.09	(2.02 to 4.74)	<.001
76 to 79	3.43	(2.14 to 5.49)	<.001
80 to 84	3.03	(1.93 to 4.74)	<.001
85 and above	2.03	(1.27 to 3.24)	.003
Female	0.65	(0.55 to 0.77)	<.001
Race/ethnicity			
Non-Hispanic, white (reference)	1.00	_	_
Hispanic	0.69	(0.52 to 0.92)	.011
Non-Hispanic, African American	0.81	(0.63 to 1.04)	.102
Non-Hispanic, Asian	1.25	(0.82 to 1.90)	.297
Non-Hispanic, other	0.70	(0.40 to 1.21)	.201
Has a usual source of care	1.87	(1.37 to 2.56)	<.001
Doctor ever told high cholesterol	6.22	(5.02 to 7.72)	<.001
Doctor ever told had high blood pressure	1.33	(1.08 to 1.65)	.009
Doctor ever told had diabetes	1.51	(1.24 to 1.85)	<.001
Current smoker	0.90	(0.70 to 1.14)	.375
SF-12 Physical Component Score	1.01	(1.00 to 1.02)	.022
Married	1.26	(1.02 to 1.57)	.033
Education			
Did not complete high school (reference)	1.00	_	_
High school graduate or GED	1.23	(1.00 to 1.52)	.055
Some college	1.39	(1.08 to 1.80)	.011
College degree or higher	1.15	(0.87 to 1.51)	.339
Insurance coverage			
Private (reference)	1.00	_	_
Medicare	1.05	(0.76 to 1.44)	.781
Medicaid	1.13	(0.78 to 1.63)	.527
Uninsured	0.90	(0.60 to 1.35)	.607
Census region			
Northeast (reference)	1.00	_	_
Midwest	1.17	(0.88 to 1.55)	.271
South	0.82	(0.62 to 1.08)	.165
West	0.90	(0.67 to 1.22)	.504
MSA	1.11	(0.85 to 1.44)	.451
Constant	0.04	(0.02 to 0.08)	<.001

ASCVD, atherosclerotic cardiovascular disease; CI, confidence interval; GED, General Equivalency Diploma; MSA, metropolitan statistical area; SF-12, 12-Item Short Form Health Survey (SF-12).

Source: Agency for Healthcare Research and Quality, Medical Expenditure Panel Survey, 2014 to 2016.

apy, ^{43,47} which have been subsequently associated with a reduction in rates of major cardiovascular events among those with a prior myocardial infarction. ⁴⁴ Despite removal of several potential system level barriers to statin use—the availability of generic statins, and potentially lower costs—we found

no significant change in overall rates of statin use for secondary prevention.

Prior research has documented that primary care physicians perceive side effects to be one of the most common reasons for statin discontinuation. ⁴⁸ Patients also perceive side effects to be a significant

barrier, as well as potential risks to health, and concern about prolonged dependence on medications. 49 Any interaction with a provider provides an opportunity for physicians and patients to discuss both side effects, which are associated with discontinuation of statin therapy, 48 and patient preferences, 50-53 which contribute to shared decision making between providers and patients. Despite this, there has been little assessment of how practice characteristics (eg, size, specialty, payer mix) interact with uptake of guideline-recommended statin use as of yet. 48,50-53,48 The association of provider and/or practice characteristics associated with diffusion of clinical guidelines is an area ripe for future research to increase delivery of guideline-concordant care.

Limitations

Our analysis has several limitations. First, MEPS is nationally representative of the civilian noninstitutionalized population and therefore excludes active-duty military personnel, people residing in nursing homes or other long-term health care facilities, and people incarcerated in jail or prison. Second, in MEPS, ASCVD history and risk factors are based on self reports of current smoking status and reports of whether a doctor has ever told them that they have various health conditions. Though underreporting of visits is common in MEPS, self reports of health status have been found to be generally consistent with clinical measures.^{54,55} Our estimates of the proportion of the population with ASCVD, also based on household reporting are quantitatively similar to estimates derived from National Health and Nutrition Examination Survey using clinical measurements.35,56 Third, in MEPS, statin use and prescription fills are based on household-reported data combined with records provided by household-reported pharmacies, which may be subject to error.²⁰ This could lead to our estimates being an understatement of the prevalence of statin use for secondary prevention; however, total prescription fills for lipid-lowering agents reported in MEPS do correspond reasonably well with industry estimates, and these data have been used in other studies of statin use. 9,12,20,57-60,61 Fourth, some people may experience statin-associated side effects. While national datasets do not capture the clinical reasons for why someone may stop a medication to distinguish true intolerance from other side effects, the 2018 ACC/

AHA guidelines note that most patients are able to tolerate statins by changing to an alternative statin, or changing the regimen or dose.⁶² Fifth, while peripheral arterial disease was not included in our study's definition of ASCVD, prior research has found that more than half of patients with peripheral arterial disease also have cardiovascular disease (CVD),⁶³ and thus, would have been included in our study population. Lastly, we were not able to distinguish whether people were receiving high, moderate, or low intensity statins, which is another important component of examining appropriate statin use.

Conclusions

Significant differences in statin use by age and gender and ethnic disparities in statin use for secondary prevention persist, despite no clinical basis for lower effectiveness among women and Hispanics. For populations that are at high risk of a recurrent ASCVD event, this gap of nearly 40% of those with a history of ASCVD not taking statins is an area where improvement is critically needed. Health information technology and electronic health records may potentially be leveraged for practice improvement.⁶⁴ Behavioral change will require health systems, providers, and patients to all play a role in increasing cardiovascular health and lowering the large burden of AS-CVD on morbidity and mortality.

The findings and conclusions in this document are those of the author(s), who are responsible for its content, and do not necessarily represent the views of AHRQ. No statement in this report should be construed as an official position of AHRO or of the US Department of Health and Human Services.

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Appendix

Table 4. Prevalence Rate of Diagnosed ASCVD, 2008 to 2016

Year	Prevalence Per 1000	95% CI
2008	161.0	151.3 to 170.6
2009	157.5	148.8 to 166.3
2010	153.7	145.5 to 162.0
2011	149.5	140.7 to 158.3
2012	147.6	138.7 to 156.6
2013	155.3	146.9 to 163.7
2014	156.6	147.9 to 165.3
2015	153.4	144.6 to 162.2
2016	152.6	144.6 to 160.6

ASCVD, atherosclerotic cardiovascular disease. Adjusted Wald test to Prob > F = 0.1931.