



AAVBC

AMERICAN ACADEMY OF VALUE BASED CARE

Lipid Medication Treatment Quick Reference Guide

2026

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LANDSCAPE OF LIPID TREATMENT UTILIZATION IN VALUE-BASED CARE

Dyslipidemia treatment prevents myocardial infarction (MI), ischemic stroke, peripheral artery events, and cardiovascular death by reducing exposure to atherogenic lipoproteins.^{1,2} It also lowers pancreatitis risk when triglycerides are severely elevated.^{3,4} In older adults, value is determined by absolute cardiovascular risk, time to benefit, treatment tolerance, function, medication burden, and whether the regimen can be sustained, not by a laboratory value alone.^{5,6}



AAVBC PERSPECTIVE

The **American Academy of Value-Based Care (AAVBC)** supports primary care ownership of lipid identification, risk estimation, statin initiation, adherence recovery, and stepwise intensification. The **2026 US dyslipidemia guideline** restores low-density lipoprotein cholesterol (LDL-C) and non-high-density lipoprotein cholesterol (non-HDL-C) goals while retaining **percentage reduction** as an important measure of response.^{5,7} **Primary care providers (PCPs) should use the PREVENT-ASCVD equations** for primary-prevention risk assessment in adults aged 30 to 79 years, then **personalize the estimate** with risk-enhancing factors and selectively use **coronary artery calcium (CAC)** to reclassify uncertain decisions.^{5,8}

Generic statins are the high-value foundation because they are inexpensive, effective, and supported by extensive outcomes evidence.^{2,5} When the **LDL-C goal is not reached** on a maximally tolerated statin, **generic ezetimibe** is usually the first add-on because it is oral, well tolerated, and low cost.⁵ Bempedoic acid, proprotein convertase subtilisin/kexin type 9 (**PCSK9**) monoclonal antibodies, inclisiran, and other specialty therapies should follow a **documented assessment of residual risk**, expected absolute benefit, adherence, intolerance, coverage, and patient preference.⁵

Older age alone is not a reason to withhold or discontinue treatment.⁵ For primary prevention after age 75 years, decisions should incorporate frailty, cognition, polypharmacy, life expectancy, function, and goals.^{5,6} Initiating a moderate-intensity statin may be reasonable when estimated life expectancy is at least 2.5 years after shared decision-making.⁵ Deprescribing may be reasonable when life expectancy is less than 1 year or treatment no longer supports the patient's goals.⁵ Secondary prevention generally remains high value unless burden or harm clearly outweighs benefit.⁵

Clinical Context in Older Adults

CLINICAL SCENARIO	CLINICAL SIGNIFICANCE ^{5,9,10}	KEY PCP CONSIDERATIONS ^{*5,6,10}
Established atherosclerotic cardiovascular disease (ASCVD)	Recurrent-event risk is high	Use high-intensity or maximally tolerated statin and add nonstatin therapy to the risk-based goal
LDL-C at least 190 mg/dL	Suggests severe primary hypercholesterolemia or familial hypercholesterolemia (FH)	Treat without relying on a short-term risk calculator ; assess family history and cascade screening
Diabetes, stage 3 or 4 chronic kidney disease (CKD), or human immunodeficiency virus (HIV), age 40 to 75 years	Confers elevated ASCVD risk	LDL-lowering therapy is recommended regardless of calculated risk or baseline LDL-C
Age 65 to 79 years without ASCVD	PREVENT-ASCVD remains applicable	Discuss therapy at intermediate or high 10-year risk and individualize at borderline risk
Age 75 years or older	Trial evidence is less complete and competing risks increase	Consider : life expectancy, frailty, function, polypharmacy, CAC when uncertain, and patient priorities
Frailty or life-limiting illness	Treatment burden may exceed delayed preventive benefit	Simplify or deprescribe when goal-concordant; do not use age alone
Persistent muscle symptoms	Common cause of abandonment and inaccurate allergy labeling	Evaluate : timing, interactions, thyroid disease, and rechallenge before declaring complete intolerance
Triglycerides at least 1000 mg/dL	Pancreatitis risk becomes a dominant concern	Address secondary causes urgently and institute a very-low-fat diet plus indicated triglyceride-lowering therapy

ABBREVIATIONS: ASCVD = atherosclerotic cardiovascular disease; PCP = primary care provider; LDL-C = low-density lipoprotein cholesterol; FH = familial hypercholesterolemia; CKD = chronic kidney disease; HIV = human immunodeficiency virus; PREVENT = Predicting Risk of cardiovascular disease EVENTS; CAC = coronary artery calcium
***Consult FDA labels for the most up-to-date dosage information, contraindications, and drug-drug interactions**

Direct Savings and Value Impact Matrix

VALUE LEVER	CLINICAL STRATEGY ^{5,11,12}	FINANCIAL MECHANISM AND IMPACT ^{5,13,14}
Acquisition-cost efficiency — generic statin first	Use generic atorvastatin or rosuvastatin at the intensity required, or another low-cost statin when tolerability or interactions dictate	Produces large LDL-C reductions at minimal pharmacy cost
Enhanced efficacy — goal-based combination therapy	Add generic ezetimibe before a specialty agent when clinically appropriate	Delivers incremental LDL-C lowering and may delay high-cost injectable therapy
Adherence recovery — persistence optimization	Use refill review, 90-day supplies, synchronization, and shared decisions about adverse effects	Prevents events caused by silent discontinuation and avoids unnecessary escalation
Waste avoidance — statin-intolerance pathway	Dechallenge, address reversible causes , and rechallenge with a different dose or statin	Prevents low-value permanent statin abandonment and inappropriate specialty spending
Targeted high-cost therapy — risk-based specialty treatment	Reserve PCSK9-directed therapy and inclisiran for patients with sufficient residual risk and documented need	Concentrates high-cost therapy where absolute event reduction is greatest
Diagnostic precision — Lp(a) and selective ApoB testing	Measure lipoprotein(a) once and use apolipoprotein B when standard lipids may underestimate risk	Improves targeting without converting biomarkers into indiscriminate repeated testing
Decision resolution — CAC for uncertainty	Use CAC selectively , not as routine screening or serial treatment monitoring	Resolves borderline decisions and reduces both undertreatment and low-benefit prescribing
Care-transition closure — post-event reconciliation	Reconcile lipid therapy after: MI, stroke, revascularization, or hospitalization	Prevents: failure to initiate, unintended discontinuation, and avoidable recurrent events

ABBREVIATIONS: LDL-C = low-density lipoprotein cholesterol; PCSK9 = proprotein convertase subtilisin/kexin type 9; Lp(a) = lipoprotein(a); ApoB = apolipoprotein B; CAC = coronary artery calcium; MI = myocardial infarction
***Consult FDA labels for the most up-to-date dosage information, contraindications, and drug-drug interactions**

INDICATIONS AND CONTRAINDICATIONS

Indications

Secondary Prevention

Clinical ASCVD includes prior acute coronary syndrome or MI, stable coronary disease, coronary or other arterial revascularization, ischemic stroke or transient ischemic attack attributable to atherosclerosis, and peripheral artery disease.^{5,7} **Most patients should receive a high-intensity or maximally tolerated statin.**⁵ For those at very high risk, the 2026 guideline recommends **LDL-C below 55 mg/dL and non-HDL-C below 85 mg/dL.**⁵ Patients with ASCVD who are **not at very high risk** generally have an **LDL-C goal below 70 mg/dL.**⁵ Risk status, achieved percentage reduction, tolerance, and patient preference **determine add-on therapy.**⁵

Primary Prevention

- **Severe hypercholesterolemia: LDL-C at least 190 mg/dL** generally warrants statin therapy **without PREVENT-ASCVD calculation.** Exclude secondary causes and **evaluate for FH**⁵
- **Diabetes, CKD, or HIV:** Adults aged 40 to 75 years with diabetes, stage 3 or 4 CKD, or HIV should receive LDL-lowering therapy **regardless of calculated risk or LDL-C level.** Intensity should reflect **risk enhancers** and **treatment goals**^{5,8}
- **Risk-based treatment:** For adults without ASCVD or subclinical atherosclerosis and LDL-C 70 to 189 mg/dL, use **PREVENT-ASCVD.** Therapy can be considered at **3% to less than 5% 10-year risk** and should be considered at **5% to less than 10%** after clinician-patient discussion. High-risk adults should generally receive therapy capable of **at least a 50% LDL-C reduction** and achievement of **LDL-C below 70 mg/dL**^{5,9}
- **Borderline or intermediate risk:** An LDL-C goal **below 100 mg/dL** and a **30% to 49%** reduction are generally appropriate when medication is selected, with greater lowering as risk increases^{5,7}
- **Older adults:** For adults aged **75 years or older** with life expectancy of **at least 2.5 years**, initiating **moderate-intensity statin therapy** may be reasonable after shared decision-making. Continue effective therapy when benefit remains aligned with goals^{5,6}

Hypertriglyceridemia

Statins remain the foundation for ASCVD risk reduction **when triglycerides are persistently elevated.**⁵ **Triglycerides from 150 to 499 mg/dL should prompt** correction of diet, alcohol exposure, diabetes, obesity, hypothyroidism, kidney or liver disease, and contributing medications.⁵ **Icosapent ethyl** may reduce residual cardiovascular risk in **selected statin-treated patients** with established ASCVD or diabetes plus additional risk features and **persistent triglyceride elevation.**⁵

At triglycerides at least **500 mg/dL**, and especially at least **1000 mg/dL**, **pancreatitis prevention** becomes increasingly important.⁵ Use a **very-low-fat diet** at the highest levels, **eliminate alcohol**, treat **hyperglycemia** and **secondary causes**, and consider **prescription omega-3 fatty acid** or fenofibrate therapy.⁵ Acute abdominal pain, vomiting, or systemic illness **requires urgent evaluation**.⁵

Special Considerations in Older Adults

CONSIDERATION	KEY POINTS* ^{5,15-19}
Secondary prevention	Evidence supports maximally tolerated statin ($\pm$ ezetimibe/PCSK9i) regardless of chronological age ; ASCVD benefit is robust at any age, so do not withhold on age alone
Primary prevention ≥ 75	RCT data are limited ; moderate-intensity statin may be reasonable if life expectancy ≥ 2.5 years after shared decision-making; relative benefit is smaller and no clear all-cause mortality benefit
Individualize, not by age alone	Base initiation/continuation on multimorbidity, frailty, polypharmacy, function, and patient priorities; never on chronological age or life expectancy alone
Time-to-benefit	Primary-prevention benefit takes ~ 2.5 years to accrue ; weigh against life expectancy before starting
Selective CAC use	When the decision is uncertain in adults >75 with life expectancy ≥ 2.5 years, CAC of 0 (or 1-10) can reclassify downward and support avoiding LLT
Deprescribing	Reasonable when risks outweigh benefits or priorities shift ; in life expectancy <1 year, stopping LLT is safe and may improve quality of life
Muscle symptoms	SAMS are the most common complaint ; true myopathy/rhabdomyolysis is rare but higher with high-dose statins in older/CKD patients; much reported intolerance is placebo; manage rather than reflexively stop
New-onset diabetes	Small dose-related risk in those with prediabetes risk factors ; ASCVD benefit outweighs it = not a reason to avoid or stop statins
Cognition	Evidence does not support statin-induced cognitive impairment; effects are neutral or possibly protective = not a reason to withhold
CKD and dosing	CKD favors LLT ; prefer lower-renal-clearance statins (atorvastatin), cap rosuvastatin (≤ 10 mg if eGFR <30); statin initiation is not beneficial on dialysis but may be continued
Polypharmacy/interactions	Reconcile CYP3A4 and other interacting drugs ; avoid gemfibrozil with statins; simplify regimens to reduce burden and adverse events
Hypertriglyceridemia	Statin remains foundational; treat secondary causes; icosapent ethyl may lower residual risk in selected ASCVD/diabetes patients ; at TG ≥ 500 mg/dL use fibrate or prescription omega-3 to prevent pancreatitis

CONSIDERATION	KEY POINTS* ^{5,15-19}
ABBREVIATIONS: PCSK9i = proprotein convertase subtilisin/kexin type 9 inhibitor; ASCVD = atherosclerotic cardiovascular disease; RCT = randomized controlled trial; CAC = coronary artery calcium; LLT = lipid-lowering therapy; SAMS = statin-associated muscle symptoms; CKD = chronic kidney disease; eGFR = estimated glomerular filtration rate; CYP3A4 = cytochrome P450 3A4; TG = triglycerides *Consult FDA labels for the most up-to-date dosage information, contraindications, and drug-drug interactions	



CLINICAL PEARL: CHANGES TO STATIN THERAPY RECOMMENDATIONS IN UPDATED 2026 AHA/ACC DYSLIPIDEMIA GUIDELINES

The 2026 guideline expands primary prevention assessment from ages **40-75 to 30-79** and replaces the pooled cohort equations with **PREVENT-ASCVD**.⁵ Statins remain recommended **regardless of calculated risk for LDL-C ≥ 190 mg/dL and adults ages 40-75 with diabetes**; stage ≥ 3 CKD and HIV are now added to this group.⁵ Statin therapy is also more strongly supported for **borderline 10-year risk** (3% to $<5\%$) and may be reasonable **even at low risk** when **LDL-C is 160-189 mg/dL or 30-year risk is $\geq 10\%$** .⁵ For older adults, **use PREVENT through age 79** and consider **coronary artery calcium scoring** when treatment remains uncertain.⁵ **Beyond age 79, individualize decisions** around ASCVD burden, frailty, polypharmacy, life expectancy, and patient goals.⁵

Contraindications and Major Precautions

CATEGORY	CONTRAINDICATION/ PRECAUTION* ^{5,17,17,20}	APPLIES TO*	PCP RESPONSE* ^{5,17,20}
Active serious liver injury	Unexplained acute hepatitis or decompensated hepatic disease	Statins and selected non-statins	Defer initiation during acute injury , evaluate cause, and use product labeling and specialty input
Pregnancy	FDA removed the class-wide absolute contraindication, but most pregnant patients should stop statins	Statins	Stop in most pregnancies ; consider continuation only in rare very-high-risk patients with specialist and obstetric input
Lactation	Statin exposure may occur in breast milk	Statins	Avoid breastfeeding if statin therapy must continue ; discuss alternatives
Severe hypersensitivity	Prior serious reaction to the drug or excipient	All classes	Avoid the implicated product and select an alternative

CATEGORY	CONTRAINDICATION/ PRECAUTION* ^{5,17,17,20}	APPLIES TO*	PCP RESPONSE* ^{5,17,20}
Gallbladder disease	May increase gallstone risk	Fibrates	Avoid or use with caution according to product labeling
Severe renal impairment	Raises fibrate exposure and myopathy risk	Fenofibrate and gemfibrozil	Apply renal restrictions and dose adjustments ; monitor kidney function
Hypertriglyceridemia	Bile acid sequestrants can raise triglycerides	Bile acid sequestrants	Avoid with severe hypertriglyceridemia or prior hypertriglyceridemic pancreatitis
Gout or hyperuricemia	Bempedoic acid can increase uric acid	Bempedoic acid	Review history and monitor clinically; counsel about tendon symptoms
Older age, frailty, polypharmacy, CKD, or hypothyroidism	Increases susceptibility to muscle symptoms and interactions	Statins, especially higher exposures	Select dose and product carefully; correct reversible risks and monitor symptoms

ABBREVIATIONS: PCP = primary care provider; FDA = Food and Drug Administration; CKD = chronic kidney disease

*Consult current FDA labeling for agent-specific dosing, renal thresholds, contraindications, reversal, and peri-procedural interruption



CLINICAL PEARL: ESSENTIAL PCP PRE-TREATMENT CHECKS

Before initiating or materially changing therapy:

- **Define the indication:** document primary or secondary prevention, ASCVD events and dates, FH suspicion, diabetes, CKD, HIV, triglyceride severity, and the **intended LDL-C and non-HDL-C goals**⁵
- **Establish baseline lipids:** obtain total cholesterol, LDL-C, high-density lipoprotein cholesterol (HDL-C), triglycerides, and non-HDL-C. **Repeat a surprising result** and obtain a fasting sample when triglycerides are high or a calculation is unreliable^{5,21}
- **Assess risk and enhancers:** use **PREVENT-ASCVD** when applicable; **measure Lp(a) at least once**; consider ApoB when triglycerides exceed 200 mg/dL, diabetes is present, or achieved LDL-C is below 70 mg/dL but residual risk is uncertain⁵
- **Check safety and secondary causes:** obtain alanine aminotransferase (ALT), review kidney function, diabetes status, thyroid disease, alcohol, medications, and baseline muscle symptoms. **Creatine kinase (CK)** is selective, **not routine**^{5,21}
- **Measure feasibility:** review swallowing, cognition, injection capacity, cost, pharmacy access, adherence history, and patient priorities **using teach-back**⁵

Common Drug and Clinical Interactions

COMBINATION	MAIN RISK* ^{7,17,22}	PCP STRATEGY* ^{5,7,17}
Simvastatin or lovastatin plus strong CYP3A4 inhibitor	Markedly increased statin exposure and myopathy	Hold or select a non-CYP3A4 statin ; follow agent-specific restrictions
Atorvastatin plus strong CYP3A4 inhibitor	Increased statin plasma concentration (systemic drug exposure)	Temporarily hold, reduce, or substitute according to interaction guidance
Ritonavir-boosted antiviral, including nirmatrelvir/ritonavir	Severe statin interaction for selected agents	Check the current interaction resource before prescribing ; hold contraindicated statins for the specified interval
Statin plus gemfibrozil	Increased myopathy and rhabdomyolysis risk	Avoid when possible ; fenofibrate is generally preferred if a fibrate is necessary
Statin plus cyclosporine or selected transplant drugs	Large increase in statin plasma drug levels (systemic exposure)	Use specialist-guided statin selection and dose limits
Bile acid sequestrant plus another oral medication	Reduced absorption	Separate administration according to the interacting product, commonly at least 1 hour before or 4 to 6 hours after
Icosapent ethyl plus antithrombotic therapy	Increased bleeding risk ; atrial fibrillation signal	Review bleeding and rhythm history and monitor clinically
Grapefruit with simvastatin, lovastatin, or atorvastatin	Increased exposure depending on quantity and product	Counsel using product-specific limits or choose an alternative statin

ABBREVIATIONS: CYP3A4 = cytochrome P450 3A4; PCP = primary care provider

***Consult current FDA labeling for agent-specific dosing, renal thresholds, contraindications, reversal, and peri-procedural interruption**

RISK ASSESSMENT, TREATMENT SELECTION, AND INTENSIFICATION

Calculate, Personalize, Reclassify

Use the 2026 guideline's CPR model:

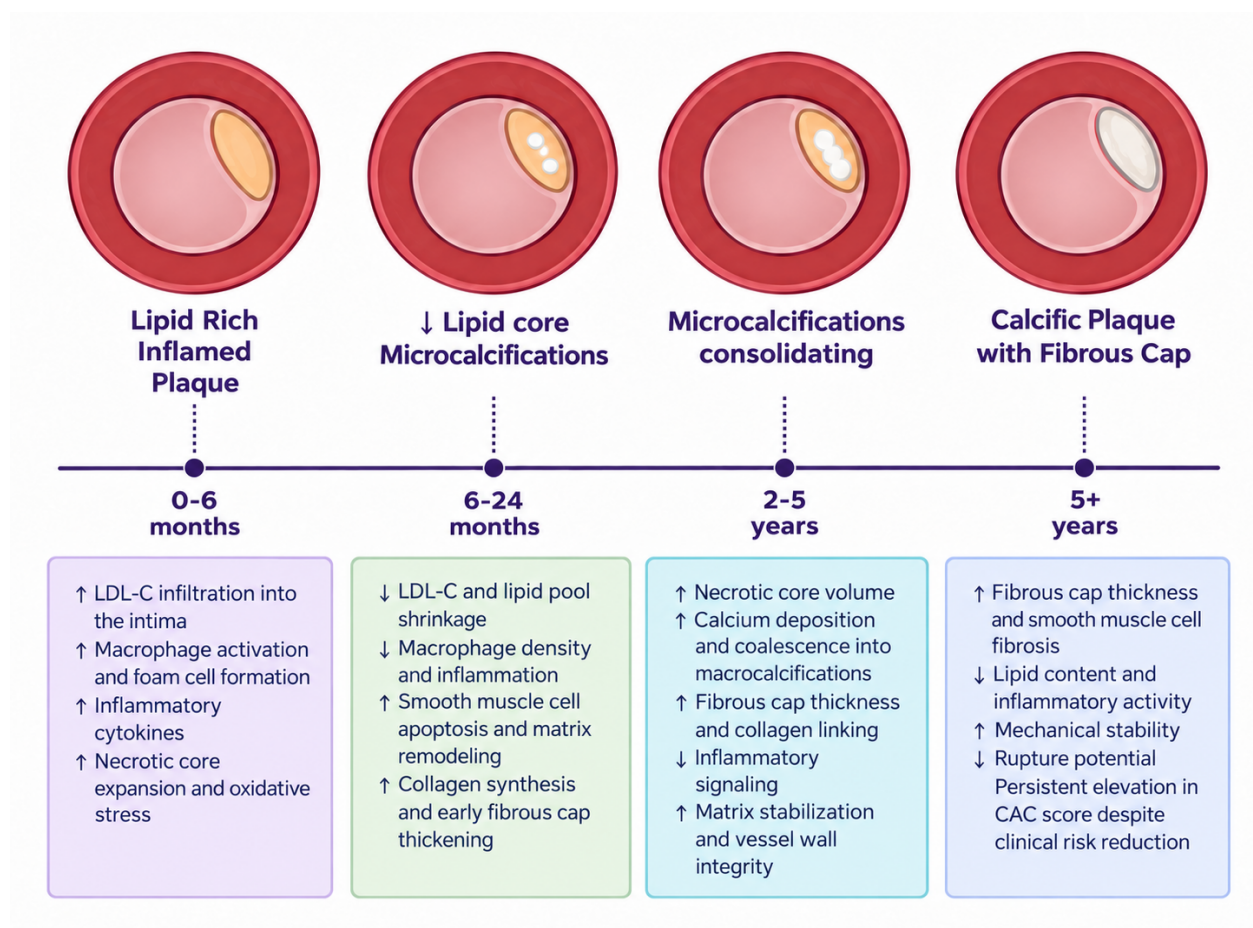
- **Calculate:** estimate **10-year PREVENT-ASCVD risk** in eligible adults aged **30 to 79 years**^{5,23}
- **Personalize:** consider **family history** of premature ASCVD, LDL-C 160 to 189 mg/dL, metabolic syndrome, inflammatory disease, elevated triglycerides, CKD, HIV, **premature menopause** or **adverse pregnancy outcomes** when relevant, and elevated Lp(a), ApoB, high-sensitivity C-reactive protein, or low ankle-brachial index^{5,24}
- **Reclassify:** use **CAC selectively** when a borderline or intermediate decision remains uncertain, then reassess the treatment recommendation with the patient^{5,25}

PREVENT-ASCVD is **not used to decide whether to treat a patient with established ASCVD**, LDL-C at least **190 mg/dL**, or **established subclinical atherosclerosis**. It should support, not replace, clinical judgment.⁵

Coronary Artery Calcium

CAC is most useful when the **decision is genuinely uncertain**.^{5,25} Any CAC supports **active LDL lowering**, with lower goals as calcium burden rises.^{5,24} A **CAC score of 0** can support **deferring medication** in selected primary-prevention adults, but it **does not create a guarantee of safety** and should not overrule diabetes, LDL-C at least 190 mg/dL, cigarette smoking, or strong premature family history.^{5,25} **In adults older than 75 years**, CAC of 0 to 10 may help **reclassify risk downward** when life expectancy is at least 2.5 years and the treatment decision remains uncertain.^{5,26}

Do not use serial CAC scanning to judge statin effectiveness. Statins may **increase plaque calcification density** while stabilizing plaque, so **a rising Agatston score does not necessarily indicate treatment failure**.^{27,28}



Practical Medication Categories

CATEGORY	TYPICAL LDL-C EFFECT* ^{5,29}	MAIN ROLE ^{5,24}	VALUE AND SAFETY CONSIDERATIONS ^{5,17,30,31}
High-intensity statin	At least 50% reduction	ASCVD, LDL-C at least 190 mg/dL, and high-risk primary prevention	Generic atorvastatin 40 to 80 mg or rosuvastatin 20 to 40 mg is usually highest value
Moderate-intensity statin	30% to 49% reduction	Borderline or intermediate primary prevention and many adults aged 75 years or older	Select by: interactions, kidney function, tolerance, and formulary
Ezetimibe	About 15% to 25% additional reduction	First oral add-on or monotherapy when statin use is limited	Generic , once daily, and generally well tolerated
Bempedoic acid	About 15% to 25% reduction	Statin intolerance or additional oral LDL lowering	Outcomes benefit demonstrated in statin-intolerant populations ; review gout and tendon risk

CATEGORY	TYPICAL LDL-C EFFECT* ^{5,29}	MAIN ROLE ^{5,24}	VALUE AND SAFETY CONSIDERATIONS ^{5,17,30,31}
PCSK9 monoclonal antibody	About 50% to 60% additional reduction	Very-high-risk ASCVD , FH, or severe residual LDL elevation	Strong LDL lowering and outcomes evidence, but specialty cost and self-injection burden
Inclisiran	About 50% reduction	Selected patients needing durable, clinician-administered LDL lowering	Twice-yearly maintenance dosing may improve implementation; cardiovascular outcomes evidence and coverage differ from PCSK9 antibodies
Bile acid sequestrant	About 15% to 25% reduction	Selected patients when triglycerides are controlled	Constipation, pill or powder burden, and binding interactions limit use in older adults

ABBREVIATIONS: LDL-C = low-density lipoprotein cholesterol; ASCVD = atherosclerotic cardiovascular disease; PCSK9 = proprotein convertase subtilisin/kexin type 9; FH = familial hypercholesterolemia

***Consult FDA labels for the most up-to-date dosage and duration information, contraindications, and drug-drug interactions**

Stepwise LDL-Lowering Pathway

This pathway applies to both **primary** and **secondary ASCVD prevention**, providing a sequential escalation strategy to reach the patient's **risk-stratified LDL-C goal** while balancing efficacy, tolerability, and treatment burden:

- Start an **evidence-based statin intensity** and document the **expected percentage reduction and goal**^{5,29}
- Repeat lipids in **4 to 12 weeks** and **assess**: actual use, tolerance, diet, secondary causes, and interactions²⁹
- Titrate to a **high-intensity** or **maximally tolerated statin when indicated**⁵
- **Add ezetimibe** when the goal remains unmet and the **expected benefit justifies treatment**^{5,24}
- For persistent high risk, select **bempedoic acid**, a **PCSK9 monoclonal antibody**, **inclisiran**, or another indicated therapy **based on**: required LDL lowering, outcomes evidence, route, adherence, contraindications, access, and patient preference^{5,24}
- **Reassess after every transition and simplify** when treatment burden exceeds likely benefit⁵

STATIN-ASSOCIATED SYMPTOMS AND DEINTENSIFICATION

Statin-associated muscle symptoms require evaluation, not automatic permanent discontinuation.⁵ **Document**: symptom location, symmetry, timing, severity, weakness, urine color, exercise change, acute illness, and relation to starting, stopping, or rechallenging therapy.⁵ Review hypothyroidism, vitamin D deficiency when clinically suspected, kidney or liver dysfunction, alcohol, interacting medications, and strenuous activity.^{5,17} **Measure CK** for severe symptoms, objective weakness, or concern for myopathy or rhabdomyolysis, not routinely for mild aches.⁵

If symptoms are clinically significant, **hold the statin until they improve, address reversible causes**, and **rechallenge** with a lower dose, a different statin, or less frequent dosing.⁵ Many patients can tolerate some statin exposure.⁵ **Combine** the maximally tolerated dose **with ezetimibe** or **another evidence-based nonstatin** rather than accepting uncontrolled risk.⁵ Persistent weakness or CK elevation after statin withdrawal warrants **urgent specialty evaluation** for rare immune-mediated necrotizing myopathy.⁵

Deintensification is appropriate when serious adverse effects recur, the medication no longer matches goals, frailty or pill burden substantially changes, or **life expectancy is too short for meaningful preventive benefit**.⁵ For patients with life expectancy less than 1 year, **discontinuation** may be reasonable **after shared decision-making**. Avoid deprescribing solely because a patient crossed an age threshold.³²



CLINICAL PEARL: COQ10 FOR STATIN-ASSOCIATED MUSCLE SYMPTOMS

The rationale is mechanistic: statins inhibit the **mevalonate pathway**, lowering endogenous ubiquinone (CoQ10) and potentially impairing mitochondrial energy production in myocytes—the leading hypothesis for **statin-associated muscle symptoms (SAMS)**.^{33–35} However, because CoQ10 is generally **well tolerated** and **inexpensive**, a time-limited trial of **200–400 mg daily** may be considered for select patients with **confirmed SAMS** who remain symptomatic and **may otherwise discontinue statin therapy**. Continue only if **symptoms clearly improve**.³⁵

TRIGLYCERIDE MANAGEMENT AND PANCREATITIS PREVENTION

Persistent triglyceride elevation should trigger **confirmation and evaluation** for uncontrolled diabetes, alcohol, high refined-carbohydrate intake, obesity, hypothyroidism, pregnancy, CKD, liver disease, nephrotic syndrome, and medications such as glucocorticoids, estrogen, retinoids, atypical antipsychotics, selected HIV therapies, or immunosuppressants.^{5,36}

For **triglycerides 150 to 499 mg/dL**, prioritize lifestyle, weight and glycemic management, and statin-based ASCVD prevention.⁵ **Do not substitute** a fibrate or nonprescription fish oil **for indicated LDL-lowering therapy**.⁵ Icosapent ethyl is not interchangeable with mixed eicosapentaenoic acid/docosahexaenoic acid supplements and should be **reserved for the evidence-based population**.^{5,37}

For **triglycerides at least 1000 mg/dL**, institute a very-low-fat eating plan, stop alcohol, treat hyperglycemia, and add indicated prescription triglyceride-lowering therapy.⁵ **Fenofibrate is generally preferred over gemfibrozil** if combination with a statin is needed.⁵ **Once pancreatitis risk is controlled, return attention to LDL-C, non-HDL-C, and global ASCVD risk.**^{5,36}

MONITORING AND CONFIRMING SAFE CONTROL

Monitoring Plan

- **Lipid panel:** repeat **4 to 12 weeks after initiation**, dose change, or adherence intervention, then **every 3 to 12 months** according to stability and risk^{5,36}
- **Percentage reduction and goals:** compare with the **untreated baseline when available** and **document LDL-C and non-HDL-C goal attainment**⁵
- **ALT:** repeat for symptoms of hepatotoxicity or a relevant clinical change, **not automatically at every lipid check**⁵
- **CK:** obtain for severe muscle symptoms, weakness, or high-risk presentations, **not routine surveillance**⁵
- **ApoB:** use selectively when triglycerides, diabetes, or very low achieved LDL-C may **leave discordant residual atherogenic particle risk**^{5,38}
- **Lp(a):** measure **at least once**; routine serial measurement is **usually unnecessary** because levels are **largely genetically determined** and no approved therapy is currently used solely to normalize Lp(a)³⁹
- **Therapy-specific monitoring:** review uric acid or gout with **bempedoic acid**, renal function with **fibrates**, injection technique and reactions with **PCSK9-directed therapy**, and atrial fibrillation or bleeding concerns with **icosapent ethyl**^{5,40}

Criteria for Safe Maintenance

Control is adequate when the patient has achieved the **risk-based LDL-C and non-HDL-C goals** or an agreed **maximally achievable reduction**, is taking the regimen consistently, has **no unacceptable toxicity**, and can **sustain the cost and administration plan**.⁵ A low LDL-C on paper does not establish value if the patient has stopped therapy, is using samples with no continuation plan, or cannot manage the injection schedule.⁵

MANAGEMENT OF APPARENT TREATMENT FAILURE

Before escalating therapy, perform a structured review:

- **Measurement:** confirm fasting status when relevant, calculation method, baseline value, laboratory variation, and whether triglycerides make LDL-C estimation unreliable⁵
- **Medication:** confirm the exact drug, dose, frequency, refill access, missed doses, intolerance, drug interactions, and unintentional discontinuation after hospitalization⁵
- **Metabolic causes:** assess hypothyroidism, nephrotic syndrome, cholestatic liver disease, uncontrolled diabetes, weight change, alcohol, and causative medications⁵
- **Lifestyle:** reassess saturated fat, ultraprocessed food, refined carbohydrate, alcohol, activity, sleep, and tobacco **without implying that lifestyle failure explains all genetic risk**⁵
- **Required magnitude:** calculate how much **additional LDL-C lowering is needed**, then select the **least burdensome therapy** capable of achieving it⁴¹

Refer to a lipid specialist or cardiology for suspected homozygous or severe heterozygous FH, LDL-C remaining markedly elevated despite verified combination therapy, recurrent ASCVD events despite goal-directed treatment, true complete statin intolerance, severe hypertriglyceridemia that persists after secondary causes are treated, pregnancy with extreme ASCVD risk, or **need for advanced agents beyond local experience**.⁵

CLOSED-LOOP PCP-LED CARE PATHWAY

- **Identify the indication:** primary prevention, secondary prevention, severe LDL elevation, FH, or pancreatitis prevention⁵
- **Calculate and personalize risk:** use **PREVENT-ASCVD** when applicable, risk enhancers, Lp(a), and selective ApoB⁵
- **Set the treatment goal:** document LDL-C, non-HDL-C, and expected percentage reduction⁵
- **Choose the highest-value feasible regimen: generic statin first**, then generic ezetimibe when appropriate¹³
- **Close access:** confirm formulary coverage, pharmacy pickup, 90-day supply when appropriate, and a **sustainable plan** after samples or prior authorization⁴²
- **Reassess in 4 to 12 weeks:** evaluate response, adherence, adverse effects, and secondary causes⁵

- **Intensify or simplify:** add therapy according to residual risk and required reduction, or deprescribe when burden exceeds benefit^{5,41}
- **Close transitions:** reconcile lipid therapy after every cardiovascular hospitalization, specialist visit, and formulary switch⁵

PCP and Team Responsibilities

- **PCP:** identify risk, order baseline and follow-up testing, initiate statins, manage common intolerance, add ezetimibe, address lifestyle and tobacco, and document goals^{42,43}
- **Clinical pharmacist:** complete interaction review, formulary comparison, adherence outreach, prior authorization support, and injection education^{42,43}
- **Dietitian:** provide a culturally and financially realistic **cardioprotective eating plan** and a **very-low-fat plan** when severe hypertriglyceridemia requires it⁴²
- **Cardiologist:** optimize very-high-risk secondary prevention and coordinate therapy after acute coronary, cerebrovascular, or revascularization events¹³
- **Lipid specialist:** manage FH, complex intolerance, refractory triglycerides, advanced therapies, and cascade screening^{5,42}
- **Patient and caregiver:** use teach-back to confirm medication identity, schedule, adverse-effect plan, and refill strategy⁴³

COST-SMART PRESCRIBING IN THE UNITED STATES

Start with the generic therapy that can meet the clinical goal. Atorvastatin and rosuvastatin provide high-intensity options; pravastatin, lower-dose rosuvastatin, and other statins may solve interaction or tolerability problems.⁵ **Generic ezetimibe** usually provides the **lowest-cost incremental LDL reduction**.^{5,41} Avoid expensive fixed-dose combinations when **separate generics provide the same clinically appropriate components at lower net cost** and acceptable pill burden.⁵

Do not assume a discount-card price is always better. For Medicare beneficiaries, prescriptions purchased outside Part D generally do not count toward plan spending or the 2026 annual Part D **out-of-pocket threshold of \$2,100**.^{44,45} Compare the plan copay, preferred pharmacy, mail order, cash price, and impact on **total annual spending**.⁴⁶ Commercial manufacturer copay cards generally cannot be used by Medicare beneficiaries, but Extra Help, charitable assistance, and plan-specific programs may apply.⁴⁶

For PCSK9 monoclonal antibodies, inclisiran, and other specialty therapies, document the indication, baseline and treated LDL-C, prior statin doses and dates, adverse effects and rechallenges, ezetimibe use, FH criteria, and ASCVD history **before submission**.^{5,41} Confirm whether coverage is under the pharmacy or medical benefit and who owns renewal. **Avoid nonmedical gaps** during prior authorization by **maintaining the best tolerated oral regimen**.^{5,41}

The lowest acquisition price is not automatically the lowest total cost. A specialty agent can be **high value** in a patient with **very-high-risk ASCVD** and major **residual LDL elevation**, while it can be **low value** in a **low-risk patient** who has not tried a statin or ezetimibe.^{41,47} Judge value by absolute event reduction, safety, persistence, and total cost of care.⁴⁸

BRAND OR HIGHER-COST OPTION ^{37,41,49,50}	GENERIC/ALTERNATIVE (EST. COST)* ^{49,51-53}	EST. SAVINGS + COST-SMART TIP* ^{41,54,55}
Brand atorvastatin (Lipitor) ~\$30/mo cash for 20 mg; historically up to 10× generic)	Generic atorvastatin 10–80 mg (~\$4–\$15/ mo; median OOP ~\$5/ mo)	~\$25/mo cash High-intensity generic (40–80 mg) is the highest-value foundation ; no evidence brand is clinically superior = prescribe generic to protect adherence
Brand rosuvastatin (Crestor) ~\$260–\$310/mo cash)	Generic rosuvastatin 5–40 mg (~\$8–\$30/ mo)	~\$230–\$300/mo High-intensity option (20–40 mg); full generic competition made rosuvastatin high-value → no advantage remains for the brand
Brand ezetimibe (Zetia) ~\$400–\$500/mo cash)	Generic ezetimibe 10 mg (~\$8–\$30/mo; net <\$10/mo)	~\$370–\$490/mo First-line oral add-on before any injectable ; high value (<\$50,000/QALY) added to maximally tolerated statin
Brand ezetimibe/statin fixed-dose combo	Separate generic statin + generic ezetimibe (~\$12–\$45/ mo combined)	Variable; usually lower with generics Two low-cost generics achieve ≥50% LDL-C reduction ; avoid brand fixed-dose combos when separate generics give the same components
Bempedoic acid (Nexletol) or bempedoic acid/ezetimibe (Nexlizet), no generic (~\$400–\$500/mo)	Generic ezetimibe ± maximally tolerated/alternative statin first (~\$8–\$30/mo)	~\$370–\$490/mo Reserve for statin intolerance or residual LDL after statin + ezetimibe; intermediate value (\$50,000–\$150,000/QALY); review gout/tendon risk
PCSK9 mAb — evolocumab (Repatha) or alirocumab (Praluent) (~\$500–\$600/mo; self-injected)	Maximally tolerated statin + generic ezetimibe first (~\$12–\$45/mo)	Reserve; adds cost Highest value in very-high-risk ASCVD/FH with major residual LDL after statin + ezetimibe; proven MACE benefit but 61% payer rejection/15% abandonment when access not secured
Inclisiran (Leqvio), no generic (~\$3,250/dose, twice yearly; clinician-administered)	PCSK9 mAb or statin + ezetimibe per residual risk	Reserve; adds cost Second-line pending CVOTs (ORION-4/VICTORION-2P); consider for documented poor adherence to PCSK9 mAb; confirm medical-vs pharmacy-benefit billing and buy-and-bill site

BRAND OR HIGHER-COST OPTION ^{37,41,49,50}	GENERIC/ ALTERNATIVE (EST. COST)* ^{49,51-53}	EST. SAVINGS + COST-SMART TIP* ^{41,54,55}
Brand icosapent ethyl (Vascepa ~\$300-\$360/mo)	Generic icosapent ethyl (~\$70-\$150/mo); do NOT substitute OTC fish oil	~\$180-\$290/mo Reserve for evidence-based REDUCE-IT population (statin-treated ASCVD/diabetes, TG 150-499); OTC mixed EPA/DHA supplements are not equivalent
Brand fenofibrate/fenofibric acid (branded formulations ~\$150-\$300/mo)	Generic fenofibrate 40-200 mg or gemfibrozil 600 mg BID (~\$10-\$40/mo)	~\$120-\$280/mo For TG ≥500 mg/dL to prevent pancreatitis ; fenofibrate preferred over gemfibrozil with a statin; reduce dose in CKD
Preventable PCSK9/statin nonadherence, unfilled specialty scripts, or cost-related discontinuation	Out-of-pocket cost review + mail-order/ 90-day fills + Extra Help/patient assistance; confirm Part D vs cash	Avoids event-driven spending 1 in 8 CVD patients report cost-related nonadherence ; review OOP at initiation, use assistance programs; IRA \$2,000 (2025)/ \$2,100 (2026) cap lowers specialty burden

ABBREVIATIONS: OOP = out-of-pocket; QALY = quality-adjusted life-year; LDL-C = low-density lipoprotein cholesterol; PCSK9 = proprotein convertase subtilisin/kexin type 9; mAb = monoclonal antibody; ASCVD = atherosclerotic cardiovascular disease; FH = familial hypercholesterolemia; MACE = major adverse cardiovascular events; CVOT = cardiovascular outcomes trial; OTC = over-the-counter; TG = triglycerides; EPA = eicosapentaenoic acid; DHA = docosahexaenoic acid; BID = twice daily; CKD = chronic kidney disease; CVD = cardiovascular disease; IRA = Inflation Reduction Act

***Consult FDA labels for the most up-to-date dosage and duration information, contraindications, and drug-drug interactions**

NON-PHARMACOLOGICAL AND PATIENT-CENTERED CARE

- **Emphasize:** vegetables, fruits, whole grains, beans, nuts, seeds, unsaturated fats, and lean proteins while **limiting ultraprocessed food, trans fat, and excess saturated fat**^{5,56}
- **Replace saturated fats with unsaturated fats** rather than simply lowering total fat^{5,56}
- Use **soluble fiber** and **plant sterol-containing foods** as adjuncts, not replacements for indicated medication^{57,58}
- Support physical activity, smoking cessation, healthy sleep, blood-pressure control, diabetes care, and weight management **according to ability and goals**^{5,13}
- **Screen for:** food insecurity, swallowing problems, cognitive impairment, depression, medication beliefs, and financial toxicity⁵⁶
- Preserve protein intake, muscle, and function in frail older adults; **avoid restrictive diets that worsen malnutrition**

- Consider a **time-limited CoQ¹⁰** trial for select patients with **confirmed statin-associated muscle symptoms**, and **vitamin K-2** in those with **vascular calcification concerns**, while counseling that outcomes evidence for both remains limited³³
- Explain that plaque stabilization may increase calcification density and that **repeat CAC is not a medication-response test**^{28,59}
- Ask about **red yeast rice** and other supplements because potency, contamination, hepatotoxicity, myopathy, and interactions **may be clinically important**^{60,61}

Patient-Centered Conversation Framework

- **Explain the target:** "The goal is to **lower cholesterol enough to reduce heart attack and stroke risk** and reach your risk-based LDL-C and non-HDL-C goal—**not to chase a single 'normal' number**." Frame the discussion around **estimated ASCVD risk** and **anticipated benefit**⁵
- **Use teach-back:** Ask the patient to restate the drug and dose, timing (including bile acid sequestrant separation and PCSK9 dosing intervals), lab schedule, supplements/OTC products to avoid, warning symptoms, and who to call^{5,41}
- **Separate lifestyle foundation from pharmacotherapy:** Explain that diet, weight loss, and activity are **foundational but usually do not replace an indicated statin**, which provides the outcome-proven LDL-C reduction^{5,17}
- **Discuss treatment burden:** Compare oral daily agents (statins, ezetimibe, bempedoic acid), self-injected PCSK9 antibodies, twice-yearly inclisiran, and prescription omega-3/fibrates by route, cost, prior authorization, and monitoring—not percentage LDL-C reduction alone⁴¹
- **Make the intolerance record durable:** Document **confirmed statin-associated muscle symptoms**, hypersensitivity, or true intolerance in the adverse-reaction field and discharge materials; **avoid labeling a nocebo or reversible reaction as permanent**, since most patients tolerate some exposure after rechallenge^{5,17}
- **Define emergencies and drug-specific warning signs:** Urgent evaluation for **severe muscle pain/weakness with dark urine** (rhabdomyolysis), **jaundice or RUQ pain** (hepatotoxicity), **angioedema** with injectables, **acute tendon pain** (bempedoic acid), new palpitations or bleeding (icosapent ethyl), and **severe abdominal pain with vomiting** (pancreatitis)^{5,17}

QUALITY ALIGNMENT

Key HEDIS/MIPs Measures

Several lipid medication and cardiovascular disease-specific MIPS and HEDIS measures are active in MY2026:

- **MIPS #441:** Ischemic Vascular Disease (IVD) All or None Outcome Measure (Optimal Control)
- **MIPS #438:** Statin Therapy for the Prevention and Treatment of Cardiovascular Disease
- **HEDIS SPC-E:** Statin Therapy For Patients With Cardiovascular Disease
- **HEDIS SMC:** Cardiovascular Monitoring for People With Cardiovascular Disease and Schizophrenia
- **HEDIS MAC:** Medication Adherence For Cholesterol (Statins)
- **HEDIS SPD-E:** Statin Therapy For Patients With Diabetes
- **HEDIS SUPD:** Statin Use in Persons With Diabetes

2026 MEASURE	STANDARD	LIPID MEDICATION APPLICABILITY* ^{1,7,57-59}
MIPS #441: Ischemic Vascular Disease (IVD) All-or-None Outcome Measure (Optimal Control)	Denominator: patients ≥ 18 with IVD (prior ASCVD event or procedure, or active CAD/ischemic diagnoses); Numerator: patients meeting ALL control targets—most recent BP $< 140/90$ mmHg AND on a statin (or documented statin use) AND on daily aspirin/antiplatelet unless contraindicated (higher rate is better); Exclusions: documented contraindications/intolerance, patient factors	This is a secondary-prevention population where statin benefit is robust regardless of age; in older adults, avoid failing the statin/antiplatelet component reflexively → document true intolerance or bleeding contraindication rather than omitting therapy, and individualize the BP component to avoid orthostasis and falls

2026 MEASURE	STANDARD	LIPID MEDICATION APPLICABILITY* ^{1,7,57-59}
MIPS #438: Statin Therapy for the Prevention and Treatment of Cardiovascular Disease	Denominator: three groups (1) ≥21 with clinical ASCVD; (2) ≥21 with LDL-C ≥190 mg/dL; (3) 40–75 with diabetes; Numerator: prescribed/on a statin of appropriate intensity during the year (higher rate is better); Exclusions: pregnancy/lactation, ESRD, cirrhosis, rhabdomyolysis/myopathy, hospice, active-liver-disease/statin-intolerance documentation	Most older adults with: ASCVD, severe hypercholesterolemia, or diabetes fall in the denominator; age alone is not a reason to withhold; in secondary prevention continue maximally tolerated statin, and clearly record intolerance or a valid exclusion rather than leaving a numerator gap
HEDIS SPC-E: Statin Therapy for Patients With Cardiovascular Disease	Denominator: males 21–75 and females 40–75 with clinical ASCVD (event or diagnosis); Numerator: two rates (1) dispensed a moderate/high-intensity statin during the year; (2) statin adherence with PDC ≥80% (higher rate is better); Exclusions: pregnancy/lactation, IVF, ESRD, cirrhosis, myopathy/rhabdomyolysis, hospice/palliative, frailty+advanced illness (age criteria)	Secondary-prevention statin benefit persists in older adults, so treat and support adherence; the "-E" version allows exclusion for members ≥66 with frailty plus advanced illness or long-term institutional residence → use it when time-to-benefit is exceeded rather than deprescribing without documentation
HEDIS SMC: Cardiovascular Monitoring for People With Cardiovascular Disease and Schizophrenia	Denominator: patients 18–64 with schizophrenia/schizoaffective disorder AND cardiovascular disease; Numerator: an LDL-C test during the measurement year (higher rate is better); Exclusions: hospice, palliative care	Although the base denominator caps at 64, the principle extends to older adults with serious mental illness who often have monitoring gaps; pair routine lipid testing with caregiver-supported adherence review, as cognitive/psychiatric barriers raise nonadherence and dosing-error risk
HEDIS MAC: Medication Adherence for Cholesterol (Statins)	Denominator: patients ≥18 with ≥2 fills of a statin on different dates; Numerator: proportion of days covered (PDC) ≥80% during the measurement period (higher rate is better); Exclusions: ESRD, hospice, and (per Part D/Star specifications) certain end-of-life and institutional statuses	Adherence, not just prescribing, drives event prevention; in older adults, use 90-day fills, refill synchronization, and cost/OOP review; simplify polypharmacy and manage (rather than reflexively stop) muscle symptoms to protect the PDC and prevent silent discontinuation

2026 MEASURE	STANDARD	LIPID MEDICATION APPLICABILITY* ^{1,7,57-59}
HEDIS SPD-E: Statin Therapy for Patients With Diabetes	Denominator: patients 40–75 with diabetes and NO clinical ASCVD; Numerator: two rates (1) dispensed a moderate/high-intensity statin; (2) statin adherence PDC $\geq 80\%$ (higher rate is better); Exclusions: pregnancy/lactation, ESRD, cirrhosis, myopathy/rhabdomyolysis, hospice/palliative, and frailty+advanced illness for the "E" set	Nearly all diabetic patients 40–75 qualify; those >75 with diabetes remain high-risk and typically benefit from moderate-intensity therapy weighed against life expectancy; document statin prescription and adherence while accounting for polypharmacy and drug-interaction burden common in this group
HEDIS SUPD: Statin Use in Persons With Diabetes	Denominator: patients 40–75 with diabetes dispensed a diabetes medication (including insulin); Numerator: dispensed at least one statin of ANY intensity during the year (higher rate is better); Exclusions: pregnancy/lactation, ESRD, cirrhosis, myalgia/myopathy/rhabdomyolysis, hospice/palliative care	Any diabetes-medication dispensing places older adults in this denominator; ensure a statin is prescribed unless a documented exclusion exists; the small statin-associated diabetes risk is outweighed by ASCVD benefit and is not a reason to withhold or stop therapy

ABBREVIATIONS: MIPS = Merit-based Incentive Payment System; IVD = ischemic vascular disease; ASCVD = atherosclerotic cardiovascular disease; CAD = coronary artery disease; BP = blood pressure; LDL-C = low-density lipoprotein cholesterol; ESRD = end-stage renal disease; HEDIS = Healthcare Effectiveness Data and Information Set; SPC-E = Statin Therapy for Patients With Cardiovascular Disease (Electronic); PDC = proportion of days covered; IVF = in vitro fertilization; SMC = Cardiovascular Monitoring for People With Cardiovascular Disease and Schizophrenia; MAC = Medication Adherence for Cholesterol; OOP = out-of-pocket; SPD-E = Statin Therapy for Patients With Diabetes (Electronic); SUPD = Statin Use in Persons With Diabetes

***Consult FDA labels for the most up-to-date dosage information, contraindications, and drug-drug interactions**

TEN KEY TAKEAWAYS FOR LIPID MEDICATION

1 The 2026 guideline reshapes primary prevention eligibility

The central change is a broader, earlier, and more granular risk framework: primary-prevention assessment now spans ages 30–79 (previously 40–75), PREVENT-ASCVD replaces the Pooled Cohort Equations, and stage ≥ 3 CKD and HIV join diabetes, LDL-C ≥ 190 mg/dL, and established ASCVD as conditions warranting statins independent of calculated risk.

2 Use PREVENT-ASCVD, not the retired Pooled Cohort Equations

Apply PREVENT for eligible primary-prevention adults aged 30 to 79 years, recognizing it generally yields lower estimated risk than the prior equations.

3 Treat statin-indicated conditions without waiting for a calculator

Established ASCVD, LDL-C at least 190 mg/dL, diabetes (ages 40–75), and now stage ≥ 3 CKD and HIV justify statins independent of a computed score.

4 Set both a percentage-reduction expectation and a risk-based goal

Document an anticipated LDL-C lowering and a risk-stratified LDL-C and non-HDL-C target at initiation.

5 Generic high-intensity statins plus generic ezetimibe are the core cost-smart sequence

Escalate along this backbone before advancing to higher-cost agents.

6 Measure Lp(a) at least once and use ApoB selectively

Apply ApoB when standard lipids may underestimate residual risk, particularly in hypertriglyceridemia or metabolic syndrome

7 Use CAC only to resolve genuinely uncertain decisions

CAC can refine borderline/intermediate risk, but CAC 0 does not override major high-risk conditions such as diabetes, LDL-C ≥ 190 mg/dL, or established ASCVD.

8 Evaluate and rechallenge statin-associated symptoms before labeling intolerance

Assess timing, secondary causes, and interactions, and attempt rechallenge or an alternative statin before declaring complete intolerance.

9 Treat severe hypertriglyceridemia and pancreatitis risk urgently

Address secondary causes and initiate triglyceride-lowering therapy promptly, especially at triglycerides at least 1000 mg/dL.

10 Individualize therapy in adults older than 75 and close the loop after transitions

Weigh life expectancy, frailty, function, polypharmacy, and goals rather than chronological age, and reassess after every initiation, cardiovascular hospitalization, prior authorization, and formulary change.

AAVBC SUMMARY AND CONCLUSION

High-value lipid treatment pairs accurate risk assessment with reliable long-term implementation.⁵ **PCPs should calculate PREVENT-ASCVD risk when appropriate**, personalize the result with clinical and biomarker risk enhancers, use CAC selectively, and establish explicit LDL-C and non-HDL-C goals.^{5,8} **Generic statins remain the first-line clinical and financial anchor**, with generic ezetimibe as the usual first add-on.^{5,13} Advanced therapy is appropriate when residual risk and required LDL lowering justify its cost and burden.^{5,41} **In older adults**, treatment should be continued, intensified, simplified, or deprescribed **according to expected benefit, frailty, function, life expectancy, and patient priorities**.⁵ The value endpoint is fewer cardiovascular and pancreatitis events **with sustainable treatment**, not prescription volume or an isolated lipid result.⁵

REFERENCES

1. Ference BA, Ginsberg HN, Graham I, et al. Low-density lipoproteins cause atherosclerotic cardiovascular disease. 1. Evidence from genetic, epidemiologic, and clinical studies. A consensus statement from the European Atherosclerosis Society Consensus Panel. *Eur Heart J*. 2017;38(32):2459-2472. doi:10.1093/eurheartj/ehx144
2. Collins R, Reith C, Emberson J, et al. Interpretation of the evidence for the efficacy and safety of statin therapy. *Lancet Lond Engl*. 2016;388(10059):2532-2561. doi:10.1016/S0140-6736(16)31357-5
3. Kleindorfer DO, Towfighi A, Chaturvedi S. 2021 Guideline for the Prevention of Stroke in Patients With Stroke and Transient Ischemic Attack. *Stroke*. 2021;52(7):e364-e467. doi:10.1161/STR.0000000000000375
4. Marston Nicholas A., Bergmark Brian A., Alexander Veronica J., et al. Olezarsen for Managing Severe Hypertriglyceridemia and Pancreatitis Risk. *N Engl J Med*. 2026;394(5):429-441. doi:10.1056/NEJMoa2512761
5. Blumenthal RS, Morris PB, Gaudino M, et al. 2026 ACC/AHA/AACVPR/ABC/ACPM/ADA/AGS/APhA/ASPC/NLA/PCNA Guideline on the Management of Dyslipidemia: A Report of the American College of Cardiology/American Heart Association Joint Committee on Clinical Practice Guidelines. *Circulation*. 2026;153(17):e1154-e1276. doi:10.1161/CIR.0000000000001423

6. Bittner V, Linnebur SA, Dixon DL, et al. Managing Hypercholesterolemia in Adults Older Than 75 years Without a History of Atherosclerotic Cardiovascular Disease: An Expert Clinical Consensus From the National Lipid Association and the American Geriatrics Society. *J Am Geriatr Soc*. 2025;73(6):1674-1696. doi:10.1111/jgs.19398
7. Wiggins BS, Barac A, Benziger CP, et al. 2026 Dyslipidemia Guideline-at-a-Glance. *J Am Coll Cardiol*. 2026;87(19):2617-2623. doi:10.1016/j.jacc.2026.02.4872
8. Abbasi J. What to Know About the New Lipid Guidelines. *JAMA*. 2026;335(15):1285-1287. doi:10.1001/jama.2026.3968
9. Anderson TS, Wilson LM, Sussman JB. Implications of the 2026 Dyslipidemia Guideline for Primary Prevention Statin Therapy. *JAMA*. Published online July 20, 2026. doi:10.1001/jama.2026.11246
10. Stein JH, Tattersall MC. Familial Hypercholesterolemia. *JAMA*. Published online June 29, 2026. doi:10.1001/jama.2026.8822
11. Reston JT, Buelt A, Donahue MP, Neubauer B, Vagichev E, McShea K. Interventions to Improve Statin Tolerance and Adherence in Patients at Risk for Cardiovascular Disease : A Systematic Review for the 2020 U.S. Department of Veterans Affairs and U.S. Department of Defense Guidelines for Management of Dyslipidemia. *Ann Intern Med*. 2020;173(10):806-812. doi:10.7326/M20-4680
12. Rosenson RS, Baker S, Banach M, et al. Optimizing Cholesterol Treatment in Patients With Muscle Complaints. *J Am Coll Cardiol*. 2017;70(10):1290-1301. doi:10.1016/j.jacc.2017.07.752
13. Virani Salim S., Newby L. Kristin, Arnold Suzanne V., et al. 2023 AHA/ACC/ACCP/ASPC/NLA/PCNA Guideline for the Management of Patients With Chronic Coronary Disease. *JACC*. 2023;82(9):833-955. doi:10.1016/j.jacc.2023.04.003
14. Kohli-Lynch CN, Bellows BK, Zhang Y, et al. Cost-Effectiveness of Lipid-Lowering Treatments in Young Adults. *J Am Coll Cardiol*. 2021;78(20):1954-1964. doi:10.1016/j.jacc.2021.08.065
15. Ponce OJ, Larrea-Mantilla L, Hemmingsen B, et al. Lipid-Lowering Agents in Older Individuals: A Systematic Review and Meta-Analysis of Randomized Clinical Trials. *J Clin Endocrinol Metab*. 2019;104(5):1585-1594. doi:10.1210/jc.2019-00195
16. Effect of statin therapy on muscle symptoms: an individual participant data meta-analysis of large-scale, randomised, double-blind trials. *Lancet Lond Engl*. 2022;400(10355):832-845. doi:10.1016/S0140-6736(22)01545-8
17. Newman CB, Preiss D, Tobert JA, et al. Statin Safety and Associated Adverse Events: A Scientific Statement From the American Heart Association. *Arterioscler Thromb Vasc Biol*. 2019;39(2):e38-e81. doi:10.1161/ATV.0000000000000073
18. Goldstein LB, Toth PP, Dearborn-Tomazos JL, et al. Aggressive LDL-C Lowering and the Brain: Impact on Risk for Dementia and Hemorrhagic Stroke: A Scientific Statement From the American Heart Association. *Arterioscler Thromb Vasc Biol*. 2023;43(10):e404-e442. doi:10.1161/ATV.0000000000000164
19. Kidney Disease: Improving Global Outcomes (KDIGO) CKD Work Group, Stevens PE, Ahmed SB, et al. KDIGO 2024 Clinical Practice Guideline for the Evaluation and Management of Chronic Kidney Disease. *Kidney Int*. 2024;105(4):S117-S314. doi:10.1016/j.kint.2023.10.018
20. Lawrence E, Johns T. Bempedoic Acid (Nexletol) for the Treatment of Hyperlipidemia and Familial Hypercholesterolemia. *Am Fam Physician*. 2021;103(6):377-378.

21. Samson SL, Vellanki P, Blonde L, et al. American Association of Clinical Endocrinology Consensus Statement: Algorithm for Management of Adults With Type 2 Diabetes – 2026 Update. *Endocr Pract.* 2026;32(4):473-518. doi:10.1016/j.eprac.2026.01.006
22. Olshansky B, Bhatt DL, Miller M, et al. Cardiovascular Benefits of Icosapent Ethyl in Patients With and Without Atrial Fibrillation in REDUCE-IT. *J Am Heart Assoc.* 2023;12(5):e026756. doi:10.1161/JAHA.121.026756
23. Ndumele Chiadi E., Rodriguez Fatima, Dixon Dave L., et al. 2026 AHA/ACC/ADA/ASN Guideline for the Prevention, Detection, Evaluation, and Management of Cardiovascular-Kidney-Metabolic Syndrome. *JACC.* 2026;87(22_Supplement):e1889-e2007. doi:10.1016/j.jacc.2026.03.056
24. Lloyd-Jones DM, Braun LT, Ndumele CE, et al. Use of Risk Assessment Tools to Guide Decision-Making in the Primary Prevention of Atherosclerotic Cardiovascular Disease: A Special Report From the American Heart Association and American College of Cardiology. *J Am Coll Cardiol.* 2019;73(24):3153-3167. doi:10.1016/j.jacc.2018.11.005
25. Rikhi R, Chen H, Mirzai S, et al. The Role of Coronary Artery Calcium in Statin Eligibility Based on the American Heart Association's PREVENT Calculator: Insights From MESA. *J Am Coll Cardiol.* Published online July 7, 2026:S0735-1097(26)06707-0. doi:10.1016/j.jacc.2026.05.039
26. Arnett DK, Blumenthal RS, Albert MA, et al. 2019 ACC/AHA Guideline on the Primary Prevention of Cardiovascular Disease: Executive Summary: A Report of the American College of Cardiology/American Heart Association Task Force on Clinical Practice Guidelines. *Circulation.* 2019;140(11):e563-e595. doi:10.1161/CIR.0000000000000677
27. Zheutlin AR, Chokshi AK, Wilkins JT, Stone NJ. Coronary Artery Calcium Testing—Too Early, Too Late, Too Often. *JAMA Cardiol.* 2025;10(5):503-509. doi:10.1001/jamacardio.2024.5644
28. Orringer CE, Blaha MJ, Blankstein R, et al. The National Lipid Association scientific statement on coronary artery calcium scoring to guide preventive strategies for ASCVD risk reduction. *J Clin Lipidol.* 2021;15(1):33-60. doi:10.1016/j.jacl.2020.12.005
29. Grundy SM, Stone NJ, Bailey AL, et al. 2018 AHA/ACC/AACVPR/AAPA/ABC/ACPM/ADA/AGS/APhA/ASPC/NLA/PCNA Guideline on the Management of Blood Cholesterol: A Report of the American College of Cardiology/American Heart Association Task Force on Clinical Practice Guidelines. *J Am Coll Cardiol.* 2019;73(24):e285-e350. doi:10.1016/j.jacc.2018.11.003
30. Nissen Steven E., Lincoff A. Michael, Brennan Danielle, et al. Bempedoic Acid and Cardiovascular Outcomes in Statin-Intolerant Patients. *N Engl J Med.* 2023;388(15):1353-1364. doi:10.1056/NEJMoa2215024
31. Ray KK, Raal FJ, Kallend DG, et al. Inclisiran and cardiovascular events: a patient-level analysis of phase III trials. *Eur Heart J.* 2023;44(2):129-138. doi:10.1093/eurheartj/ehac594
32. Kutner JS, Blatchford PJ, Taylor DHJ, et al. Safety and benefit of discontinuing statin therapy in the setting of advanced, life-limiting illness: a randomized clinical trial. *JAMA Intern Med.* 2015;175(5):691-700. doi:10.1001/jamainternmed.2015.0289
33. Raizner AE, Quiñones MA. Coenzyme Q(10) for Patients With Cardiovascular Disease: JACC Focus Seminar. *J Am Coll Cardiol.* 2021;77(5):609-619. doi:10.1016/j.jacc.2020.12.009
34. Qu H, Guo M, Chai H, Wang WT, Gao ZY, Shi DZ. Effects of Coenzyme Q10 on Statin-Induced Myopathy: An Updated Meta-Analysis of Randomized Controlled Trials. *J Am Heart Assoc.* 2018;7(19):e009835. doi:10.1161/JAHA.118.009835

35. Kennedy C, Köller Y, Surkova E. Effect of Coenzyme Q10 on statin-associated myalgia and adherence to statin therapy: A systematic review and meta-analysis. *Atherosclerosis*. 2020;299:1-8. doi:10.1016/j.atherosclerosis.2020.03.006
36. Virani Salim S., Morris Pamela B., Agarwala Anandita, et al. 2021 ACC Expert Consensus Decision Pathway on the Management of ASCVD Risk Reduction in Patients With Persistent Hypertriglyceridemia. *JACC*. 2021;78(9):960-993. doi:10.1016/j.jacc.2021.06.011
37. Wilkins JT, Lloyd-Jones DM. Novel Lipid-Lowering Therapies to Reduce Cardiovascular Risk. *JAMA*. 2021;326(3):266-267. doi:10.1001/jama.2021.2244
38. Johannesen CDL, Mortensen MB, Langsted A, Nordestgaard BG. Apolipoprotein B and Non-HDL Cholesterol Better Reflect Residual Risk Than LDL Cholesterol in Statin-Treated Patients. *J Am Coll Cardiol*. 2021;77(11):1439-1450. doi:10.1016/j.jacc.2021.01.027
39. Reyes-Soffer G, Ginsberg HN, Berglund L, et al. Lipoprotein(a): A Genetically Determined, Causal, and Prevalent Risk Factor for Atherosclerotic Cardiovascular Disease: A Scientific Statement From the American Heart Association. *Arterioscler Thromb Vasc Biol*. 2022;42(1):e48-e60. doi:10.1161/ATV.000000000000147
40. Spitz J, Patel J, Agarwala A, et al. A Critical Appraisal of Lipid Management in the Post-Statins Era: Comparison on Guidelines, Therapeutic Targets, and Screening in a Case-Based Framework of Lipid Management. *JACC Adv*. 2025;4(6 Pt 2):101823. doi:10.1016/j.jacadv.2025.101823
41. Lloyd-Jones Donald M., Morris Pamela B., Ballantyne Christie M., et al. 2022 ACC Expert Consensus Decision Pathway on the Role of Nonstatin Therapies for LDL-Cholesterol Lowering in the Management of Atherosclerotic Cardiovascular Disease Risk. *JACC*. 2022;80(14):1366-1418. doi:10.1016/j.jacc.2022.07.006
42. Cheeley MK, Kirkpatrick CF, Brown EE, et al. Multidisciplinary teams in clinical lipidology and cardiometabolic care: A National Lipid Association Expert Clinical Review. *J Clin Lipidol*. 2025;19(4):737-747. doi:10.1016/j.jacl.2025.05.002
43. NLA Expert Panel, Jacobson TA, Maki KC, et al. National Lipid Association Recommendations for Patient-Centered Management of Dyslipidemia: Part 2. *J Clin Lipidol*. 2015;9(6):S1-S122.e1. doi:10.1016/j.jacl.2015.09.002
44. Cai CL, Bhaskar A, Kesselheim AS, Rome BN. Changes in Medicare Part D Plan Designs After the Inflation Reduction Act. *JAMA Intern Med*. 2025;185(10):1266-1273. doi:10.1001/jamainternmed.2025.4003
45. Narasimmaraj PR, Oseran A, Tale A, et al. Out-of-Pocket Drug Costs for Medicare Beneficiaries With Cardiovascular Risk Factors Under the Inflation Reduction Act. *J Am Coll Cardiol*. 2023;81(15):1491-1501. doi:10.1016/j.jacc.2023.02.002
46. Lowe EF, Gerasta D, Balser M, et al. Contributors and Solutions to High Out-of-Pocket Costs for Heart Failure Medications: A State-of-the-Art Review. *J Am Coll Cardiol*. 2025;85(4):365-377. doi:10.1016/j.jacc.2024.11.011
47. Sabatine MS, Giugliano RP. Low-Density Lipoprotein Cholesterol Treatment in the Proprotein Convertase Subtilisin/Kexin Type 9 Inhibitor Era: Getting Back on Target. *JAMA Cardiol*. 2017;2(9):935-936. doi:10.1001/jamacardio.2017.2293

48. Ray KK, Corral P, Morales E, Nicholls SJ. Pharmacological lipid-modification therapies for prevention of ischaemic heart disease: current and future options. *Lancet Lond Engl*. 2019;394(10199):697-708. doi:10.1016/S0140-6736(19)31950-6
49. Lin S yu, Baumann K, Zhou C, Zhou W, Cuellar AE, Xue H. Trends in Use and Expenditures for Brand-name Statins After Introduction of Generic Statins in the US, 2002-2018. *JAMA Netw Open*. 2021;4(11):e2135371-e2135371. doi:10.1001/jamanetworkopen.2021.35371
50. Green JB, Ross JS, Jackevicius CA, Shah ND, Krumholz HM. When Choosing Statin Therapy: The Case for Generics. *JAMA Intern Med*. 2013;173(3):229-232. doi:10.1001/jamainternmed.2013.1529
51. Luo J, Seeger JD, Donneyong M, Gagne JJ, Avorn J, Kesselheim AS. Effect of Generic Competition on Atorvastatin Prescribing and Patients' Out-of-Pocket Spending. *JAMA Intern Med*. 2016;176(9):1317-1323. doi:10.1001/jamainternmed.2016.3384
52. U.S. Food and Drug Administration. *Approved Drug Products with Therapeutic Equivalence Evaluations (Orange Book)*. 46th ed. U.S. Food and Drug Administration; 2026. <https://www.fda.gov/drugs/drug-approvals-and-databases/approved-drug-products-therapeutic-equivalence-evaluations-orange-book>
53. Sasidharan A, Bagepally BS, Kumar SS, Jagadeesh KV, Natarajan M. Cost-effectiveness of Ezetimibe plus statin lipid-lowering therapy: A systematic review and meta-analysis of cost-utility studies. *PloS One*. 2022;17(6):e0264563. doi:10.1371/journal.pone.0264563
54. Desai NR, Farbaniec M, Karalis DG. Nonadherence to lipid-lowering therapy and strategies to improve adherence in patients with atherosclerotic cardiovascular disease. *Clin Cardiol*. 2023;46(1):13-21. doi:10.1002/clc.23935
55. Kazi DS, DeJong C, Chen R, Wadhera RK, Tseng CW. The Inflation Reduction Act and Out-of-Pocket Drug Costs for Medicare Beneficiaries With Cardiovascular Disease. *J Am Coll Cardiol*. 2023;81(21):2103-2111. doi:10.1016/j.jacc.2023.03.414
56. Lichtenstein AH, Appel LJ, Vadiveloo M, et al. 2021 Dietary Guidance to Improve Cardiovascular Health: A Scientific Statement From the American Heart Association. *Circulation*. 2021;144(23):e472-e487. doi:10.1161/CIR.0000000000001031
57. Brum J, Ramsey D, McRorie J, Bauer B, Kopecky SL. Meta-Analysis of Usefulness of Psyllium Fiber as Adjuvant Antilipid Therapy to Enhance Cholesterol Lowering Efficacy of Statins. *Am J Cardiol*. 2018;122(7):1169-1174. doi:10.1016/j.amjcard.2018.06.040
58. Jellinger PS, Handelsman Y, Rosenblit PD, et al. American Association of Clinical Endocrinologists and American College of Endocrinology Guidelines for Management of Dyslipidemia and Prevention of Cardiovascular Disease. *Endocr Pract*. 2017;23:1-87. doi:10.4158/EP171764.APPGL
59. Razavi AC, Agatston AS, Shaw LJ, et al. Evolving Role of Calcium Density in Coronary Artery Calcium Scoring and Atherosclerotic Cardiovascular Disease Risk. *JACC Cardiovasc Imaging*. 2022;15(9):1648-1662. doi:10.1016/j.jcmg.2022.02.026
60. Cicero AFG, Fogacci F, Zambon A. Red Yeast Rice for Hypercholesterolemia: JACC Focus Seminar. *J Am Coll Cardiol*. 2021;77(5):620-628. doi:10.1016/j.jacc.2020.11.056
61. Department of Veterans Affairs, Department of Defense. *VA/DoD Clinical Practice Guideline on Lipid Management for Cardiovascular Disease Risk Reduction*. Department of Veterans Affairs and Department of Defense; 2025. Accessed August 18, 2026. https://www.healthquality.va.gov/HEALTHQUALITY/guidelines/CD/lipids/Lipids-CPG_2025-Guideline_final_20260106.pdf

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