

## Policies Repository



**Policy Title** Atorvastatin (Lipitor®) / Amlodipine/atorvastatin (Caduet®)

**Policy Number** FS.CLIN.42

*Application of Pharmacy Policy is determined by benefits and contracts. Benefits may vary based on product line, group or contract. Some medications may be subject to precertification, age, gender or quantity edits. Individual member benefits must be verified.*

*This Pharmacy Policy document describes the status of pharmaceutical information and/or technology at the time the document was developed. Since that time, new information relating to drug efficacy, interactions, contraindications, dosage, administration routes, safety or FDA approval may have changed. If the Medical/Pharmacy Reviewer is aware of any new information on the subject of this document, please provide it promptly to the Medical/Pharmacy Policy Department. This information may include new FDA approved indications, withdrawals or other FDA alerts. This type of information is relevant not only when considering whether this Policy should be updated, but also when applying it to current requests for coverage.*

*Members are advised to use participating pharmacies in order to receive the highest level of benefits.*

**Policy** **Atorvastatin (Lipitor®)** is indicated for the treatment of hypercholesterolemia and the prevention of cardiovascular disease in individuals with multiple risk factors for coronary heart disease.

**Amlodipine/atorvastatin (Caduet®)** is indicated for the treatment of hypercholesterolemia and hypertension in individuals for whom treatment with both amlodipine and atorvastatin is appropriate.

The use of atorvastatin (Lipitor®) and amlodipine/atorvastatin (Caduet®) require prior authorization (ie, clinical pharmacy and/or Medical Director review).

**Policy Description**

**Atorvastatin (Lipitor®)** is an HMG-CoA reductase inhibitor. HMG-CoA reductase is the rate-limiting enzyme in de novo cholesterol synthesis. HMG-CoA reductase inhibitors are frequently used to reduce the levels of plasma total cholesterol and low-density lipoprotein (LDL) cholesterol in individuals with hypercholesterolemia. It is believed that HMG-CoA reductase inhibitors lower cholesterol levels by reducing the production of mevalonic acid, which results in the following:

- A reduction in hepatic cholesterol synthesis
- A compensatory increase in the expression of high-affinity LDL receptors on hepatocyte membranes
- Stimulation of LDL catabolism

**Amlodipine/atorvastatin (Caduet®)** is a combination of two drugs, a dihydropyridine calcium channel blocker (amlodipine) and an HMG-CoA reductase inhibitor (atorvastatin). The amlodipine component of amlodipine/atorvastatin (Caduet®) inhibits the transmembrane influx of calcium ions into vascular smooth muscles and cardiac muscle to cause a reduction in peripheral vascular resistance and a reduction in blood pressure.

**Policy Guideline Inclusion**

**Atorvastatin (Lipitor®)** or **amlodipine/atorvastatin (Caduet®)** is approved when **all** of the following inclusion criteria are met:

- Documentation of a minimum 30-day trial and failure or contraindication/intolerance/allergy to one of the following agents:
  - Lovastatin-containing product
  - Pravastatin-containing product
  - Simvastatin-containing product
- Documentation of a minimum 30-day trial and failure or contraindication/intolerance/allergy to rosuvastatin calcium (Crestor®)

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#### Policy Guideline Exclusion

**Atorvastatin (Lipitor®) or amlodipine/atorvastatin (Caduet®)** is denied when **any** of the following exclusion criteria are present:

- No documentation of a minimum 30-day trial and failure or contraindication/intolerance/allergy to one of the following agents:
  - Lovastatin-containing product
  - Pravastatin-containing product
  - Simvastatin-containing product
- No documentation of a minimum 30-day trial and failure or contraindication/intolerance/allergy to rosuvastatin calcium (Crestor®)

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#### Policy List of Applicable Drugs

Brand Name	Generic Name
Caduet	amlodipine/atorvastatin
Lipitor	atorvastatin

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#### Dosing and Administration

Refer to the specific manufacturer's prescribing information for administration and dosage details for each specific agent.

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#### Policy References

Ballantyne CM, Blazing MA, King TR, Brady WE, Palmisano J. Efficacy and safety of ezetimibe co-administered with simvastatin compared with atorvastatin in adults with hypercholesterolemia. *Am J Cardiol.* 2004;93(12):1487-1494.

Branchi A, Fiorenza AM, Torri A, et al. Effect of atorvastatin 10 mg and simvastatin 20 mg on serum triglycerides levels in patients with hypercholesterolemia. *Curr Ther Res Clin Exp.* 2001;62(5):408-415.

Caduet® [package insert]. New York, NY: Pfizer, Inc.; 2007. Also available at: <http://www.caduet.com>. Accessed January 6, 2010.

Cannon CP, Braunwald MD, McCabe CH, et al. Intensive versus moderate lipid lowering with statins after acute coronary syndrome [published correction appears in *N Engl J Med.* 2006;354(7):778]. *N Engl J Med.* 2004;350(15):1495-1504.

Chan WB, Ko GT, Yeung VT, et al. A comparative study of atorvastatin and simvastatin as monotherapy for mixed hyperlipidaemia in Type 2 diabetic patients. *Diabetes Res Clin Pract.* 2004;66(1):97-99.

Davidson M, Ma P, Stein EA, et al. Comparison of effects on low-density lipoprotein cholesterol and high-density lipoprotein cholesterol with rosuvastatin versus atorvastatin in patients with type IIa or IIb hypercholesterolemia. *Am J Cardiol.* 2002;89(3):268-275.

Facts and Comparisons. Lipitor. [Facts and Comparisons Web site]. Available at: <http://www.factsandcomparisons.com> [via subscription only]. Accessed January 6, 2010.

Facts and Comparisons. Vytorin. [Facts and Comparisons Web site]. Available at: <http://www.factsandcomparisons.com> [via subscription only]. Accessed January 6, 2010.

Gartlehner G, Hansen R, Kahwati L, et al. Drug class review on HMG-CoA reductase inhibitors (statins). Oregon Health and Science University (OHSU). Final Report Update 1, July 2005.

Grace KA, Swiecki J, Hyatt R, et al. Implementation of a therapeutic-interchange clinic for HMG-Co A reductase inhibitors. *Am J Health Syst Pharm*. 2002;59(11):1077-1082.

Grundy SM, Cleeman JI, Merz CN, et al. Implication of recent clinical trials for the National Cholesterol Education Program Adult Treatment Panel III Guidelines [published correction appears in *Circulation*. 2004;110(6):763]. *Circulation*. 2004;110(2):227-239.

Hippisley-Cox J, Cater R, Pringle M, Coupland C. Cross sectional survey of effectiveness of lipid lowering drugs in reducing serum cholesterol concentration in patients in 17 general practices. *BMJ*. 2003;326(7391):689.

Jones PH, Davidson MH, Stein EA, et al. STELLAR Study Group. Comparison of the efficacy and safety of rosuvastatin versus atorvastatin, simvastatin, and pravastatin across doses (STELLAR Trial). *Am J Cardiol*. 2003;92(2):152-160.

Jones P, Kafonek S, Laurora I, Hunninghake D. Comparative dose efficacy study of atorvastatin versus simvastatin, pravastatin, lovastatin, and fluvastatin in patients with hypercholesterolemia (the CURVES Study) [published correction appears in *Am J Cardiol*. 1998;82(1):128]. *Am J Cardiol*. 1998;81(5):582-587.

Kadikoylu G, Yukselen V, Yavasoglu I, Bolaman Z. Hemostatic effects of atorvastatin versus simvastatin. *Ann Pharmacother*. 2003;37(4):478-484.

LaRosa JC, Grundy SM, Waters DD, et al. Intensive lipid lowering with atorvastatin in patients with stable coronary disease. *N Engl J Med*. 2005;352(14):1425-1435.

Lipitor® [package insert]. New York, NY: Pfizer, Inc.; 2005. Also available online at: <http://www.lipitor.com>. Accessed January 6, 2010.

Marz W, Wollschlager H, Klein G, Neiss A, Wehling M. Safety of low-density lipoprotein cholesterol reduction with atorvastatin versus simvastatin in a coronary heart disease population (the TARGET TANGIBLE trial). *Am J Cardiol*. 1999;84(1):7-13.

McManus B. FDC Reports. Lipitor adds claims for high-dose use from TNT, IDEAL- but label is mixed. 2007;69:11. [The Pink Sheet Web site]. 03/12/07. Available at: [http://www.thepinksheet.com/fdcreports/story/search/submitSearch.do?doClear=true&subsNum=&publications=pink&query=lipitor&find\\_button.x=13&find\\_button.y=6](http://www.thepinksheet.com/fdcreports/story/search/submitSearch.do?doClear=true&subsNum=&publications=pink&query=lipitor&find_button.x=13&find_button.y=6) [via subscription only]. Accessed November 11, 2008.

Pearson TA, Denke MA, McBride PE, et al. A community-based, randomized trial of ezetimibe added to statin therapy to attain NCEP ATP III goals for LDL cholesterol in hypercholesterolemic patients: The Ezetimibe Add-on to Statin for Effectiveness (EASE) Trial. *Mayo Clin Proc*. 2005;80(5):587-595.

Pedersen TR, Faergeman O, Kastelein JJ, et al. High-dose atorvastatin vs usual-dose simvastatin for secondary prevention after myocardial infarction: the IDEAL study: a randomized controlled trial [published correction appears in *JAMA*. 2005;294(24):3092]. *JAMA*. 2005;294(19):2437-2445.

Taylor AJ, Grace K, Swiecki J, et al. Lipid-lowering efficacy, safety, and costs of a large-scale therapeutic statin formulary conversion program. *Pharmacotherapy*. 2001;21(9):1130-1139.

Vytorin® [package insert]. North Wales, PA: Merck/Schering-Plough Pharmaceuticals; 2008. Also available online at: [http://www.vytorin.com/vytorin/shared/documents/vytorin\\_pi.pdf](http://www.vytorin.com/vytorin/shared/documents/vytorin_pi.pdf). Accessed January 6, 2010.

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