

Contains Nonbinding Recommendations
Draft – Not for Implementation
Draft Guidance on Atorvastatin Calcium
October 2024

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Active Ingredient:	Atorvastatin calcium
Dosage Form:	Suspension
Route:	Oral
Strength:	20 mg/5 mL
Recommended Study:	One in vivo bioequivalence study with pharmacokinetic endpoints

1. Type of study: Fasting
Design: Single-dose, two-treatment, two-period crossover in vivo
Strength: 20 mg/5 mL at a dose of 80 mg (20 mL)
Subjects: Healthy males and non-pregnant, non-lactating females
Additional comments: Applicants may consider using a reference-scaled average bioequivalence approach. If using this approach, provide evidence in the studies of high variability in the bioequivalence parameters of area under the plasma concentration time curve and/or peak concentration (i.e., within-subject variability $\geq 30\%$). For detailed information on this approach, refer to the most recent version of the FDA guidance for industry on *Bioequivalence Studies with Pharmacokinetic Endpoints for Drugs Submitted Under an ANDA*.^a

Analytes to measure: Atorvastatin and its active metabolites, ortho and para-hydroxylated atorvastatin in plasma

Submit the metabolite data as supportive evidence of comparable therapeutic outcome. For the metabolites, the following data should be submitted: individual and mean concentrations, individual and mean pharmacokinetic parameters, and geometric means and ratios of means for AUC and C_{max}.

Bioequivalence based on (90% CI): Atorvastatin

Waiver request of in vivo testing: Not applicable

Dissolution test method and sampling times: The dissolution information for this drug product can be found in the FDA's Dissolution Methods database, <http://www.accessdata.fda.gov/scripts/cder/dissolution/>. Conduct comparative dissolution testing on 12 dosage units for each of the test product and reference listed drug (RLD).¹ Specifications will be determined upon review of the abbreviated new drug application.

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^a For the most recent version of a guidance, check the FDA guidance website at <https://www.fda.gov/regulatory-information/search-fda-guidance-documents>.

¹ If the RLD is not available, refer to the most recent version of the FDA guidance for industry on *Referencing Approved Drug Products in ANDA Submissions*.