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APPLICATION NUMBER for: 020702, S018

CHEMISTRY REVIEW(S)

CHEMIST'S REVIEW

1. Organization CDER/HFD-510 Division of Metabolism and Endocrine Drug Products		2. NDA # 20-702 Approved: 17-DEC-1996	
3. Name And Address Of Applicant Parke-Davis Pharmaceutical Research Division of Warner-Lambert Company 2800 Plymouth Road P.O. Box 1047 Ann Arbor, MI 48106-1047 (313) 966-5000		4. Supplement SEI-018 Doc.03-MAR-1999 Rec.04-MAR-1999	
		5. Name of the Drug Lipitor Tablets	
		6. Nonproprietary Name Atorvastatin Calcium	
7. Supplement provides support for the use of Lipitor (Atorvastatin Calcium) tablets to decrease the non HDL-C/HDL-C ratio and to increase HDL-C in patients with primary hypercholesterolemia (heterozygous familial) and mixed dislipidemia (Fredrickson Type IIa and Type IIb)		8. Amendment -	

9. Pharmacological Category Lipid Modifier. HMG-CoA reductase inhibitor/ Antihyperlipoproteinemic agent.	10. How Dispensed	11. Related -N. A.-
12. Dosage Form Tablet	13. Potency 10-, 20- And 40-mg	

14. Chemical Name And Structure

Atorvastatin Calcium

$(C_{33}H_{34}FN_2O_5)_2Ca$

CAS 134523-03-8
CAS 134523-00-5 (atorvastatin)
FW $2 \times 557.7 + 40.0 = 1155.38$
FW calcium salt trihydrate $(C_{33}H_{34}FN_2O_5)_2Ca \cdot 3H_2O = 1209.42$
FW free acid $C_{33}H_{35}FN_2O_5 = 558.66$

1H-Pyrrole-1-heptanoic acid, 2-(4-fluorophenyl)-β,δ-dihydroxy-5-(1-methylethyl)-3-phenyl-4-[(phenylamino)carbonyl]-, calcium salt (2:1), [R-(R*,R*)]-

15. Comments: This Efficacy Supplement provides the information to support for the use of Lipitor (Atorvastatin Calcium) 10-, 20- and 40-mg tablets to decrease the non HDL-C/HDL-C ratio and to increase HDL-C in patients with primary hypercholesterolemia (heterozygous familial) and mixed dislipidemia (Fredrickson Type IIa and Type IIb): Approved drug substance, Atorvastatin Calcium, and drug product, Lipitor 10-, 20- and 40-mg. Tablets will be used for this indications. No Environmental Assessment is required.

16. Conclusions And Recommendations Since there are no changes in chemistry, manufacture and controls of both drug substance and drug product, this supplement is approvable from the chemistry view point.

17. Reviewer Name (and signature) <u>Xavier Ysern, PhD</u>	Date Completed 18-OCT-1999
R/D Init.	filename:

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AE Xavier Ysern
10/18/99