

**CENTER FOR DRUG EVALUATION AND
RESEARCH**

APPLICATION NUMBER: 20-550/S-012

MICROBIOLOGY REVIEW

**MICROBIOLOGY REVIEW
DIVISION OF ANTIVIRAL DRUG PRODUCTS (HFD-530)**

**NDA 20-550 Serial #s SLR-010
SE-2 (012)**

REVIEWER : N. Biswal

Date Submitted : 2/26/2001

Date Assigned : 3/14/2001

Date Received : 3/05/2001

Date Reviewed : 3/21/2001

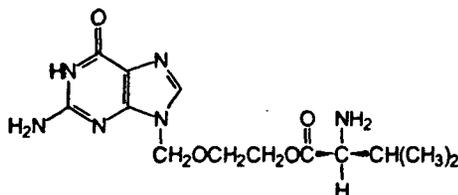
**Sponsor: GlaxoSmithKline, Inc.
PO.Box 13398
Five Moore Drive
Research Triangle Park, NC 27709**

Product Names:

a. Code Names: Valaciclovir, Valtrex, 256U87

b. Chemical: L-Valine 2-[(2-Amino-1,6-dihydro-6-oxo-9H-purin-9-yl)methoxy]ethyl ester hydrochloride

Structural Formula:



Empirical Formula: C₁₃H₂₀N₆O₄

Molecular Weight.: 324.34

Drug category: Antiviral

Indication: Treatment of Initial and Recurrent Episodes of Genital Herpes Simplex Virus Infections, Suppression of Recurrent Genital Herpes, Treatment of Herpes Zoster

Dosage Form/Route of Administration: 500 mg, 1000 mg Caplets/Oral

Supporting Documents: NA

BACKGROUND AND SUMMARY: In this amendment the sponsor has accepted the revisions to the Microbiology section of the Valtrex® label as recommended earlier. However, there are a number of minor revisions still needed to standardize the contents of the label as follows.

RECOMMENDATIONS: Please revise the MICRIOLGY section as suggested below.

MICROBIOLOGY:

- Under **Mechanism of Antiviral Action**

Line 26. _____

Line 28. Remove _____ The sentence in lines 27 and 28 should read as follows.

"The inhibitory activity of acyclovir is highly selective due to its affinity for the enzyme thymidine kinase (TK) encoded by HSV and VZV."

Line 30. _____

- Under **Antiviral Activities**

Line 35. _____

Line 38. Rewrite the sentence in lines 37, 38 and 39 to read as follows.

"Sensitivity testing results, expressed as the concentration of the drug required to inhibit the growth of virus in cell culture by 50% (IC₅₀), vary greatly depending upon a number of factors."

Line 40, 42 and 43. _____

Nilambar Biswal, Ph.D.

CONCURRENCES:

**HFD-530/Assoc Dir
HFD-530/TLMicro**

CC:

**HFD-530/Orig. IND
HFD-530/Division File
HFD-530/TLMicro
HFD-530/Review Micro
HFD-530/Young, RPM**

Nilambar Biswal
3/26/01 01:19:41 PM
MICROBIOLOGIST

Julian O Rear
3/26/01 02:44:50 PM
MICROBIOLOGIST

James Farrelly
4/4/01 08:14:29 AM
PHARMACOLOGIST