

**CENTER FOR DRUG EVALUATION AND
RESEARCH**

APPLICATION NUMBER:

20-944

CHEMISTRY REVIEW(S)

Anti-inflammatory, Analgesic, and Ophthalmic Drug Products, HFD-550
Review of Chemistry, Manufacturing, and Controls

NDA #: 20-944 REVIEW #1

DATE REVIEWED: November 12, 1998

<u>SUBMISSION TYPE</u>	<u>SUBMISSION DATE</u>	<u>USER FEE DATE</u>
Original	12/19/97	12/22/98
Amendment 1	2/12/98	(Certify ready for inspection on or after 4/30/98)
Amendment 2	2/23/98	(Claim Categorical Exclusion for EA submission)
Amendment 3	3/20/98	(A new stability testing site, corrections in dissolution data and stability results)
Amendment 4	7/16/98	(To provide method validation)

Reviewed: All

NAME & ADDRESS OF APPLICANT:

Whitehall-Robins, 5 Giralda Farms, Madison, NJ 07940-0871

DRUG PRODUCT NAME

Proprietary: Advil^R Chewable Tablet

Established: Ibuprofen

Code Name/#:

Chem.Type/Ther.Class:

Temporary relief of headache, backache, muscular aches, toothache, the minor pain of arthritis, the pain of menstrual cramps, the minor aches and pains associated with the common cold, sore throat, and reduction of fever

PHARMACOL. CATEGORY: NSAID

DOSAGE FORM: Chewable Tablet

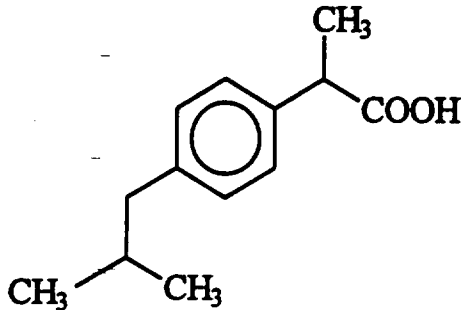
STRENGTHS: 50 mg and 100 mg

ROUTE OF ADMINISTRATION: Oral

DISPENSED: Rx OTC

RELATED DOCUMENTS (if applicable): None

CHEMICAL NAME, STRUCTURAL FORMULA, MOLECULAR FORMULA AND WEIGHT:



IBUPROFEN

C₁₃H₁₈O₂

206.28

CONSULTS:

The trade name "Advil Chewable Tablet" is found acceptable by Nomenclature Committee. A copy of Nomenclature Committee's recommendation is attached to this review.

Dissolution specification: Acceptable to Biopharm. Copies of the request are attached to this review.

REMARKS:

EER status: Acceptable. A copy of the EER is attached.

Method validation: A copy of method validation was sent [redacted] 2/13/98.

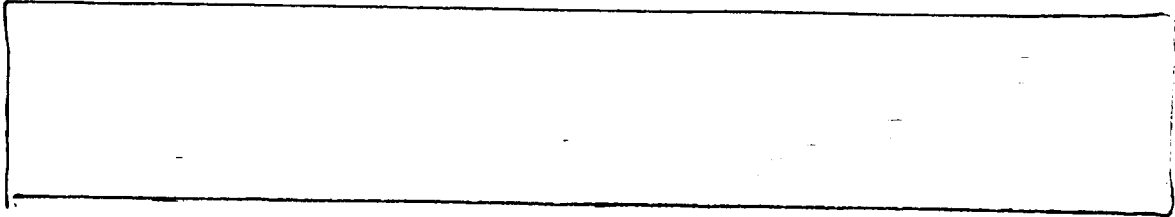
Expiration dating period (Proposed): 24 months for bottles, pouches and [redacted] blisters and 18 months for [redacted] blisters.

Container/closure (C/C) system:

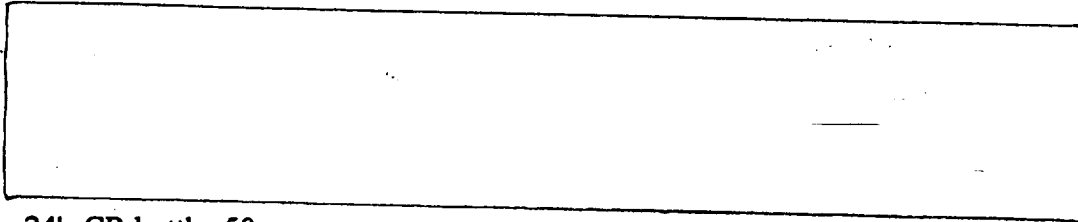
2's, CR pouch: rectangular, heat sealed, foil pouch

[redacted]

1's CR blister: opaque, white blister with aluminum foil lidding



1's CR blister: opaque, white blister with aluminum foil lidding



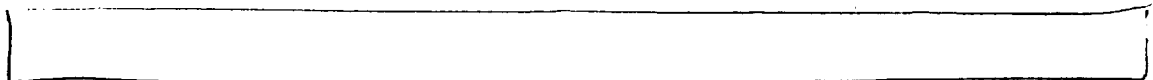
24's CR bottle, 50 mg:



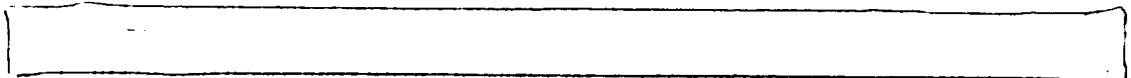
24's CR bottle, 100 mg:



100's CR bottle, 50 mg:



100's CR bottle, 100 mg:

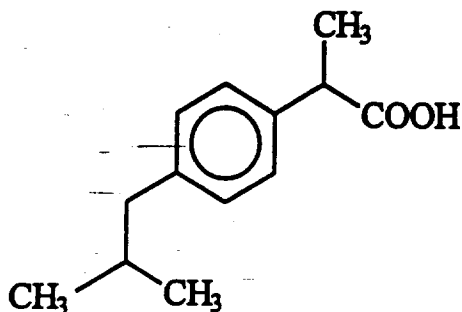


ENVIRONMENTAL ASSESSMENT:

Claims Categorical Exclusion for the submission of EA citing 21 CFR 25.31(a) in the amendment dated 2/23/98.

SUPPORTING DOCUMENTS:

***This page of the document
contains confidential
information that will not
be included in the
redacted portion of the
document for the public to
obtain.***



IBUPROFEN

$C_{13}H_{18}O_2$

206-28

CONSULTS:

The trade name "Advil Chewable Tablet" is found acceptable by Nomenclature Committee. A copy of Nomenclature Committee's recommendation is attached to this review.

Dissolution specification: Acceptable to [] Copies of the request are attached to this review.

REMARKS:

EER status: Acceptable. A copy of the EER is attached.

Method validation: A copy of method validation was sent []
2/13/98.

Expiration dating period (Proposed): 24 months for bottles, pouches and []
blisters and 18 months for [] blisters.

APPL. AS THIS WAY
ON ORIGINAL

Conclusions and Recommendations:

In this submission, the applicant has satisfactorily responded to all the deficiencies and comments noted in chemist's review #1. The applicant has provided adequate information on the chemistry, manufacture and controls of the drug substance and drug product. From the chemistry standpoint, the NDA is recommended for approval.

/S/

12/7/98.

Vispi Bhavnagri, Review/Chemist

/S/

12/7/98

Team Leader, Hasmukh Patel

cc:

Orig. NDA 20-944

HFD-550/Division File

HFD-550/BHo/Bhavnagri

HFD-550/Cook

HFD-550/Patel

HFD-830/Chi-wan Chen

R/D Init by: Patel