

30 or above. "high sun protection product."

(2) For products containing any ingredient identified in §352.10, the following labeling statement may be used under the heading "Other information" or anywhere outside of the "Drug Facts" box or enclosure. "Sun alert: Limiting sun exposure, wearing protective clothing, and using sunscreens may reduce the risks of skin aging, skin cancer, and other harmful effects of the sun." Any variation of this statement will cause the product to be misbranded under section 502 of the act.

(f) Products labeled for use only on specific small areas of the face (e.g., lips, nose, ears, and/or around eyes) and that meet the criteria established in §201.66(d)(10) of this chapter. The title, headings, subheadings, and information described in §201.66(c) of this chapter shall be printed in accordance with the following specifications:

(1) The labeling shall meet the requirements of §201.66(c) of this chapter except that the title, headings, and information described in §201.66(c)(1), (c)(3), and (c)(7) may be omitted, and the headings, subheadings, and information described in §201.66(c)(2), (c)(4), (c)(5), and (c)(6) may be presented as follows:-

(i) The active ingredients (§201.66(c)(2) of this chapter) shall be listed in alphabetical order.

(ii) The heading and the indication required by §201.66(c)(4) may be limited to: "Use [in bold type] helps prevent sunburn."

(iii) The "external use only" warning in §201.66(c)(5)(i) of this chapter may be omitted.

(iv) The subheadings in §201.66(c)(5)(iii) through (c)(5)(vii) of this chapter may be omitted, provided the information after the heading "Warnings" states: "Keep out of eyes." and "Stop use if skin rash occurs."

(v) The warning in §201.66(c)(5)(x) of this chapter may be limited to the following: "Keep out of reach of children."

(vi) For a lipstick, the warnings "Keep out of eyes" in §352.52(f)(1)(iv) and "Keep out of reach of children" in §352.52(f)(1)(v) and the directions in §352.52(d) may be omitted.

(2) The labeling shall be printed in accordance with the requirements of §201.66(d) of this chapter except that any requirements related to §201.66(c)(1), (c)(3), and (c)(7), and the horizontal barlines and hairlines described in §201.66(d)(8), may be omitted.

EFFECTIVE DATE NOTE: At 68 FR 33380, June 4, 2003, §352.52 was amended by revising the heading in paragraphs (c)(2) and (d)(4) and by revising paragraphs (f)(1)(ii) and (f)(1)(vi), effective June 4, 2004. For the convenience of the user, the revised text is set forth as follows:

§352.52 Labeling of sunscreen drug products.

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(c) * * *

(2) For products containing any ingredient identified in §352.10 marketed as a lip protectant or lipstick. * * *

(d) * * *

(4) For products marketed as a lip protectant or lipstick. * * *

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(f) * * *

(1) * * *

(ii) The heading and the indication required by §201.66(c)(4) of this chapter may be limited to: "Use [in bold type] helps protect against sunburn." For a lip protectant product, the heading and the indication required by §201.66(c)(4) may be limited to: "Use [in bold type] helps protect against sunburn and chapped lips."

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(vi) For a lip protectant product or lipstick, the warnings "Keep out of eyes" in §352.52(f)(1)(iv) and "Keep out of reach of children" in §352.52(f)(1)(v) and the directions in §352.52(d) may be omitted.

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§352.60 Labeling of permitted combinations of active ingredients.

Statements of identity, indications, warnings, and directions for use, respectively, applicable to each ingredient in the product may be combined to eliminate duplicative words or phrases so that the resulting information is clear and understandable.

(a) *Statement of identity.* For a combination drug product that has an established name, the labeling of the

product states the established name of the combination drug product, followed by the statement of identity for each ingredient in the combination, as established in the statement of identity sections of the applicable OTC drug monographs. For a combination drug product that does not have an established name, the labeling of the product states the statement of identity for each ingredient in the combination, as established in the statement of identity sections of the applicable OTC drug monographs.

(b) *Indications.* The labeling of the product states, under the heading "Uses," the indication(s) for each ingredient in the combination as established in the indications sections of the applicable OTC drug monographs, unless otherwise stated in this paragraph. Other truthful and nonmisleading statements, describing only the indications for use that have been established in the applicable OTC drug monographs or listed in this paragraph (b), may also be used, as provided by § 330.1(c)(2) of this chapter, subject to the provisions of section 502 of the Federal Food, Drug, and Cosmetic Act (the act) relating to misbranding and the prohibition in section 301(d) of the act against the introduction or delivery for introduction into interstate commerce of unapproved new drugs in violation of section 505(a) of the act.

(1) In addition, the labeling of the product may contain any of the "other allowable statements" that are identified in the applicable monographs.

(2) For permitted combinations containing a sunscreen and a skin protectant identified in § 352.20(b).

(c) *Warnings.* The labeling of the product states, under the heading "Warnings," the warning(s) for each ingredient in the combination, as established in the warnings section of the applicable OTC drug monographs. For permitted combinations containing a sunscreen and a skin protectant identified in § 352.20(b).

(d) *Directions.* The labeling of the product states, under the heading "Directions," directions that conform to the directions established for each ingredient in the directions sections of the applicable OTC drug monographs, unless otherwise stated in this para-

graph. When the time intervals or age limitations for administration of the individual ingredients differ, the directions for the combination product may not contain any dosage that exceeds those established for any individual ingredient in the applicable OTC drug monograph(s), and may not provide for use by any age group lower than the highest minimum age limit established for any individual ingredient. For permitted combinations containing a sunscreen and a skin protectant identified in § 352.20(b).

EFFECTIVE DATE NOTE: At 68 FR 33380, June 4, 2003, § 352.60 was amended by revising paragraphs (b)(2), (c), and (d), effective June 4, 2004. For the convenience of the user, the revised text is set forth as follows:

§ 352.60 Labeling of permitted combinations of active ingredients.

* * * * *

(b) * * *

(2) For permitted combinations containing a sunscreen and a skin protectant identified in § 352.20(b), any or all of the applicable indications for sunscreens in § 352.52(b) and the indication for skin protectants in § 347.50(b)(2)(i) of this chapter should be used. For products marketed as a lip protectant, the indication in § 352.52(f)(1)(ii) should be used.

(c) *Warnings.* The labeling of the product states, under the heading "Warnings," the warning(s) for each ingredient in the combination, as established in the warnings section of the applicable OTC drug monographs, except that the warning for skin protectants in § 347.50(c)(3) of this chapter is not required for permitted combinations containing a sunscreen and a skin protectant identified in § 352.20(b). For products marketed as a lip protectant or lipstick, § 352.52(f)(1)(iii), (f)(1)(iv) (except "Keep out of eyes," which may be omitted), and (f)(1)(vi) apply.

(d) *Directions.* The labeling of the product states, under the heading "Directions," directions that conform to the directions established for each ingredient in the directions sections of the applicable OTC drug monographs, unless otherwise stated in this paragraph. When the time intervals or age limitations for administration of the individual ingredients differ, the directions for the combination product may not contain any dosage that exceeds those established for any individual ingredient in the applicable OTC drug monograph(s), and may not provide for use by any age group lower than the highest minimum age limit established for any individual ingredient. For permitted combinations containing a sunscreen and a

skin protectant identified in §352.20(b), the directions for sunscreens in §352.52(d) should be used. For products marketed as a lip protectant or lipstick, §352.52(d)(4) applies.

Subpart D—Testing Procedures

§ 352.70 Standard sunscreen.

(a) *Laboratory validation.* A standard sunscreen shall be used concomitantly in the testing procedures for determining the SPF value of a sunscreen drug product to ensure the uniform evaluation of sunscreen drug products. The standard sunscreen shall be an 8-percent homosalate preparation with a mean SPF value of 4.47 (standard deviation =1.279). In order for the SPF determination of a test product to be considered valid, the SPF of the standard sunscreen must fall within the standard deviation range of the expected SPF (i.e., 4.47 ± 1.279) and the 95-percent confidence interval for the mean SPF must contain the value 4.

(b) *Preparation of the standard homosalate sunscreen.* (1) The standard homosalate sunscreen is prepared from two different preparations (preparation A and preparation B) with the following compositions:

COMPOSITION OF PREPARATION A AND PREPARATION B OF THE STANDARD SUNSCREEN

Ingredients—	Percent by weight
Preparation A	
Lanolin	5.00
Homosalate	8.00
White petrolatum	2.50
Stearic acid	4.00
Propylparaben	0.05
Preparation B	
Methylparaben	0.10
Edetate disodium	0.05
Propylene glycol	5.00
Triethanolamine	1.00
Purified water U.S.P	74.30

(2) Preparation A and preparation B are heated separately to 77 to 82 °C, with constant stirring, until the contents of each part are solubilized. Add preparation A slowly to preparation B while stirring. Continue stirring until the emulsion formed is cooled to room temperature (15 to 30 °C). Add sufficient purified water to obtain 100 grams of standard sunscreen preparation.

(c) *Assay of the standard homosalate sunscreen.* Assay the standard

homosalate sunscreen preparation by the following method to ensure proper concentration:

(1) *Preparation of the assay solvent.* The solvent consists of 1 percent glacial acetic acid (V/V) in denatured ethanol. The denatured ethanol should not contain a UV radiation absorbing denaturant.

(2) *Preparation of a 1-percent solution of the standard homosalate sunscreen preparation.* Accurately weigh 1 gram of the standard homosalate sunscreen preparation into a 100-milliliter volumetric flask. Add 50 milliliters of the assay solvent. Heat on a steam bath and mix well. Cool the solution to room temperature (15 to 30 °C). Then dilute the solution to volume with the assay solvent and mix well to make a 1-percent solution.

(3) *Preparation of the test solution (1:50 dilution of the 1-percent solution).* Filter a portion of the 1-percent solution through number 1 filter paper. Discard the first 10 to 15 milliliters of the filtrate. Collect the next 20 milliliters of the filtrate (second collection). Add 1 milliliter of the second collection of the filtrate to a 50-milliliter volumetric flask. Dilute this solution to volume with assay solvent and mix well. This is the test solution (1:50 dilution of the 1-percent solution).

(4) *Spectrophotometric determination.* The absorbance of the test solution is measured in a suitable double beam spectrophotometer with the assay solvent and reference beam at a wavelength near 306 nanometers.

(5) *Calculation of the concentration of homosalate.* The concentration of homosalate is determined by the following formula which takes into consideration the absorbance of the sample of the test solution, the dilution of the 1-percent solution (1:50), the weight of the sample of the standard homosalate sunscreen preparation (1 gram), and the standard absorbance value (172) of homosalate as determined by averaging the absorbance of a large number of batches of raw homosalate:

$$\text{Concentration of homosalate} = \text{absorbance} \times 50 \times 100 \times 172 = \text{percent concentration by weight.}$$