

**MINISTRY OF HEALTH AND FAMILY WELFARE  
(Department of Health)**

**NOTIFICATION**

New Delhi, the 30th June, 2005

**G.S.R. 431(E).**—Whereas a draft of certain rules further to amend the Drugs and Cosmetics Rules, 1945 was published, as required by sections 12 and 33 of the Drugs and Cosmetics Act, 1940 (23 of 1040), in the Gazette of India, Extraordinary, Part II, section 3, sub section(i), dated the 8<sup>th</sup> November, 2004, in the notification of the Government of India in the Ministry of Health and Family Welfare (Department of Health), number GSR 738(E), dated the 8<sup>th</sup> November, 2004 inviting objections and suggestions from all persons likely to be affected thereby before the expiry of a period of thirty days from the date on which copies of the Official Gazette containing the said notification were made available to the public;

And whereas copies of the said Gazette were made available to the public on 11-11-04.

And whereas, objections and suggestions received from the public on the said draft rules have been considered by the Central Government;

Now, therefore, in exercise of the powers conferred by sections 12 and 33 of the said Act, the Central Government, after consultation with the Drugs Technical Advisory Board, hereby makes the following rules further to amend the Drugs and Cosmetics Rules, 1945, namely: -

1. (1) These rules may be called the Drugs and Cosmetics (5th Amendment) Rules, 2005.
- (2) They shall come into force on the date of their publication in the Official Gazette.

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2. In the Drugs and Cosmetics Rules, 1945, in Schedule M,
- (i) After the heading “ Good Manufacturing Practices and Requirements of Premises, Plant and Equipment for Pharmaceutical Products”, in the “Note”, for the words “and no other manufacturing activity shall be under taken therein”, the words “and no other manufacturing activity shall be undertaken therein except in respect of units licensed prior to 11<sup>th</sup> December, 2001” shall be substituted.
  - (ii) In PART 1, under the heading “**Good Manufacturing Practices for Premises and Materials**”, in sub-heading “1 GENERAL REQUIREMENTS”,-
    - (a) in paragraph “**3. Production area**”, in sub-paragraph 3.4, for the words “creation of recesses”, occurring in the first sentence, the words “accumulation of dust” shall be substituted.
    - (b) in paragraph “**7. Health, clothing and sanitation of workers**”, in sub-paragraph 7.7, for the words “ clean towels, hand dryers”, the words “clean towels or hand dryers” shall be substituted;
    - (c) in paragraph “**8. Manufacturing operations and controls-**”, in sub-paragraph 8.2.2., for the words “The effective segregation of these areas shall be demonstrated with adequate records of maintenance and services”., the words “ The effective segregation of these areas shall be validated with adequate records of maintenance and services” shall be substituted;
    - (d) in paragraph “**8. Manufacturing operations and controls**”, in sub-paragraph 8.2.9, for the words “ There shall be segregated enclosed areas, secured for recalled or rejected material and for such material which are to be re-processed or recovered” , the words “ There shall be segregated secured areas for recalled or rejected material and for such material which are to be re-processed or recovered” shall be substituted;
    - (e) in Paragraph “**25. Distribution records**”, in sub-paragraph 25.2, for the words “ such that finished batch of a drug can be traced to the retail level”, the words “so as” shall be substituted.

(iii) In PART 1A, under the heading " SPECIFIC REQUIREMENTS FOR MANUFACTURE OF STERILE PRODUCTS, PARENTERAL PREPARATIONS (SMALL VOLUME INJECTABLES AND LARGE VOLUME PARENTERALS) AND STERILE OPHTHALMIC PREPARATIONS",-

(1) In paragraph 3. **Air handling System (Central Air conditioning)**.-

(a) in sub-para 3.6, for the words "Grade C background and if necessary, under Laminar Air Flow work station", the words, "Grade D background and should be handled in such a way that they are not re-contaminated" shall be substituted;

(b) in sub-paragraph 3.10, for the words, figures and letters "shall not exceed 27 degree centigrade and relative humidity 55% respectively.", the words, figures and letters "shall be  $27 \pm 2$  degree centigrade and relative humidity  $55\% \pm 5$ , respectively.", shall be substituted.

(c) for TABLE 1 the following table shall be substituted, namely:-

"TABLE 1

The air borne particulate classification for manufacture of sterile products.

Grade	Maximum number of permitted particles per cubic metre equal to or above			
	At rest (b)		In Operation (a)	
	0.5 $\mu$ m	5 $\mu$ m	0.5 $\mu$ m	5 $\mu$ m
A	3500	0	3500	0
B(a)	3500	0	3,50,000	2000
C(a)	350,000	2000	35,00,000	20,000
D(a)	35,00,000	20,000	Not defined (c)	Not defined (c)

(d) in the notes, in para (a) for the words "Grade A; corresponds with class 100 or M 3.5 or class 5; Grade B with class 1000 or M 4.5 or ISO class 6", the words "Grade A and B corresponds with class 100 or M 3.5 or class 5" shall be substituted.

(e) in TABLE III, under the item Grade C, for the words "usually at risk" the words, "unusually at risk" shall be substituted;

(II) In paragraph "**6. Sanitation**", in sub-paragraph 6.4, for the words, "where alcohol or isopropyl alcohol is used for disinfectants for use as hand sprays, the preparation of the same shall be done in bulk preparation area and the diluted solution membrane filtered into suitable sterile containers held in aseptic area", the words "where alcohol or isopropyl alcohol is used for dilution of disinfectants for use as hand sprays, the preparation of the same shall be done in bulk preparation area in grade C" shall be substituted.

(III) In paragraph "**8. Water and Steam Systems**", in sub-paragraph 8.9, the words, figures and letters "The steam condensate shall meet microbiological specification of not more than 10cfu per 100 ml. The condensate shall also meet IP specification for Water for injection and shall have an endotoxin levels of not more than 0.25 EU/ml. There shall be a suitable schedule for the monitoring of steam quality." shall be omitted.

(IV) In PART IB, under the heading "SPECIFIC REQUIREMENTS FOR MANUFACTURE OF ORAL SOLID DOSAGE FORMS (TABLETS AND CAPSULES),-

(a) in paragraph 1, under the heading "**General**", in sub-paragraph 1.5, for the words, "Where the facilities are designed to provide special environmental conditions of pressure differentials between rooms, these conditions shall be regularly monitored and any specification results brought to the immediate attention of the production and quality assurance departments which shall be immediately attended to.", the words "Where the facilities are designed to provide special environmental conditions of pressure differentials between rooms, these conditions shall be regularly monitored and any deviation shall be brought to the immediate attention of the production and quality assurance departments" shall be substituted.,

(b) In paragraph **"2. Sifting, mixing and granulation-"**, in sub-paragraph 2.1 after the word, " extractors", the words, " or in a dedicated area for each operation" shall be inserted.

(V) In PART IC, under the heading "SPECIFIC REQUIREMENTS FOR MANUFACTURE OF ORAL LIQUIDS (SYRUPS, ELIXIRS, EMULSIONS AND SUSPENSION),-

(a) in paragraph **"1. Building and Equipment"**, in sub-paragraph 1.8, after the words "stainless steel", the words "or any other appropriate material which is scratch proof, washable and smooth" shall be added at the end.

(b) In paragraph **"3. Manufacturing"**, in sub-paragraph 3.1, for the words " Manufacturing personnel shall wear non-fiber shedding clothing to prevent contamination of the products", the words "Manufacturing personnel shall wear wherever required, non-fiber shedding clothing to prevent contamination of the products" shall be substituted.

(VI) In PART ID, under the heading "SPECIFIC REQUIREMENTS FOR MANUFACTURE OF TOPICAL PRODUCTS, i.e., EXTERNAL PREPARATIONS (CREAMS, OINTMENTS, PASTES, EMULSIONS, LOTIONS, SOLUTIONS, DUSTING POWDERS AND IDENTICAL PRODUCTS)",-

(a) in paragraph 2, the words "The air shall be ventilated." occurring as the last sentence shall be omitted;

(b) in paragraph 5, for the words, "No rags or dusters", the words "Suitable cleaning equipment and material" shall be substituted.

(VII) In PART II, under the heading, " Requirements of Plant and Equipment",-

(A) In paragraph **" 1. External Preparations"-**,

(a) in item (1), for the words, " Mixing and storage tanks (stainless steel);" the words, "Mixing and

- storage tanks preferably of stainless steel or any other appropriate material,” shall be substituted;
- (b) in item (2), after the words, “Jacketted Kettle”, the words, “Stainless steel container” shall be inserted;
  - (c) in item (8), the words “ (Electrically operated)” shall be omitted;
  - (d) after item 8, Under the heading “ Area.-,” after clause (2), the following shall be added, namely:-

“Note- The requirement for ancillary area in this part shall not apply to units registered before 1<sup>st</sup> January 2002.”

(B) In paragraph **2. Oral Liquids preparations,-**

- (i) in item (1), for the words “Mixing and storage tanks (Stainless steel )” the words “Mixing and storage tanks preferably of Stainless steel or any other appropriate material.” shall be substituted;
- (ii) after item (9), Under the heading “Area.-” after clause (2), the following shall be added, namely:-

“Note- The requirement for ancillary area in this part shall not apply to units registered before 1<sup>st</sup> January 2002.”;

(C) In paragraph **3. Tablets,-**

- (a) in the opening paragraph, for the words, “tablet machine”, the words, “tablet compression machine” shall be substituted;
- (b) in sub-paragraph 3.1,
  - (i) in clause (a), in item (4), for the words, “Granulator”, the words, “Granulator wherever required.” shall be substituted.
  - (ii) In clause (b), in item (5), for the words, “Dissolution test apparatus” the words “Dissolution test apparatus” wherever required.” shall be substituted;
  - (iii) In clause (d), in item (i), after the words “jacketed Kettle”, the words “stainless steel container or any other appropriate material” shall be inserted;

- (iv) In clause (d) after item (6), the following shall be added, namely:-

Note- The requirement for ancillary area in this part shall not apply to units registered before 1<sup>st</sup> January 2002.”;

- (C) in sub-paragraph 3.2, under the heading “Area.-”, the following shall be inserted at the end, namely:-

Note- The requirement for ancillary area in this part shall not apply to units registered before 1<sup>st</sup> January 2002.”;

- (d) in sub-paragraph 3.4, the words “/dispersible” shall be omitted.

- (D) in paragraph “**4. Powders.-**”,

- (i) in item (5), the words “(electrically operated)” shall be deleted;
- (ii) Under the heading “Arca.-”, the following shall be added at the end, namely:-

“Note-The requirement for additional room in this part shall not apply to units registered before 1<sup>st</sup> January 2002.

- (E) in paragraph “**5. Capsules.-**”

- (i) in item (2), the words “ ( preferably semi automatic or automatic filling machines)” shall be omitted.
- (ii) under the heading “Area.-”, the following shall be inserted at the end, namely:-

“Note- The requirement for ancillary area in this part shall not apply to units registered before 1<sup>st</sup> January 2002.

- (F) in paragraph “**7. Ophthalmic preparations.-**”,

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- (i) Under the heading "Area.-", after clause (2), the following shall be added at the end, namely:-

"Note- The requirement for ancillary area in this part shall not apply to units registered before 1<sup>st</sup> January 2002."

(G) in paragraph **"11. Parenteral Preparations.-"**

- (i) in sub-paragraph 11.1, under the heading "Area.-", after clause (3), the following shall be added, namely:-

"Note- The requirement for ancillary area in this part shall not apply to units registered before 1<sup>st</sup> January 2002."

- (ii) in sub-paragraph 11.2, under the heading "Area.-", after item (3), the following note shall be added at the end, namely :—

"Note- The requirement for ancillary area in this part shall not apply to units registered before 1<sup>st</sup> January 2002."

(VIII) In Schedule M, after part II, the following shall be added, namely:-

"Note 1:- There are certain categories of drugs such as chemicals and pharmaceutical aids, gauzes and bandages, medicinal gases, empty gelatin capsules, non chemical/mechanical contraceptives, diagnostic kits and reagents, medical devices, new dosage forms and their delivery systems, disinfectant fluids, antacids, raw-materials manufactured from sea bittern, veterinary biologicals including poultry vaccines, re-packing of drugs etc. for which this Schedule does not prescribe specific requirements of space and equipments. The Licensing Authority or Central Licence Approving Authority, as the case may be, in respect of such categories of drugs, have the discretion to modify the requirements of this Schedule, if he is of the opinion that having regard to the nature of the products and extent of manufacturing operations and for reasons to be recorded in writing, it is necessary to relax or alter them in the

circumstances of a particular case and direct the manufacturer to carry out necessary modifications in them and the modifications having been made, approve the manufacture of such categories of the drugs.

Note II:- In case of manufacturers licensed to manufacture drugs prior to the 11<sup>th</sup> December 2001, the requirements of this Schedule shall also apply to them from 1<sup>st</sup> July 2005.”

[F. No. X-11014/7/2003-DMS & PFA]  
RITA TEAOTIA, Jt. Secy.

**Foot Note :** The Principal rules were published in the Official Gazette *vide* number F-28/45H (1), dated the 21st December, 1945 and last amended *vide* G.S.R. 174(E) dated 16-3-05.