MANUFACTURING & QUALITY AUDIT DIVISION
CENTRE FOR DRUG ADMINISTRATION

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GUIDANCE NOTES ON PREPARATION OF A SITE MASTER FILE

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GUIDANCE NOTES

Subject: PREPARATION OF A SITE MASTER FILE

WHAT IS A SITE MASTER FILE ?

A Site Master File (SMF) is a document prepared by the manufacturer containing specific and factual Good Manufacturing Practice (GMP) information about the production and/or control of pharmaceutical manufacturing operations carried out at the named site. If only part of a manufacturing operation is carried out on the site, a SMF need only describe those activities, eg analysis, packaging etc.

HOW SHOULD A SITE MASTER FILE BE SUBMITTED ?

A Site Master File should be concisely written in English and, as far as possible, not exceed 25-30 A4 sheets.

The Site Master File should have an edition number and an effective date, and preferably be submitted on loose individually numbered A4 sheets. The sheets should be ring-bound to ensure the integrity of the document. Wherever possible, simple plans, outline drawings or schematic layouts should be used instead of narrative. These plans etc., should fit on A4 sheets of paper. The format and heading of the site master file should be set out as follows:-

Chapter 1

C.1 GENERAL INFORMATION

REQUIREMENT

C.1.1 Brief information on the site (including name and address), relation to other sites and, particularly, any information relevant to understand the manufacturing operations.

GUIDANCE
(Not more than 250 words or one A4 sheet)

C.1.1 Outline the company's activities, other sites (if any), in addition to the site which is the subject of this report.

REQUIREMENT

C.1.2 Pharmaceutical manufacturing activities as licensed by the Competent Authorities.

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C.1.2.1 *Indicate whether the site has been approved by your own national authority, or any foreign Competent Authority (if the latter, name the authority and state if approval granted is for the manufacture of medicinal products of the same or different description from that in the application).*

C.1.2.2 *Quote the relevant document (licence) as issued by the Competent Authority. State the period of validity of licence document (if the validity of the document is given in the country concerned). Any conditions and/or restrictions should be stated.*

REQUIREMENT

C.1.3 Any other manufacturing activities carried out on the site.

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C.1.3 *This covers both pharmaceutical and non-pharmaceutical activities.*

NB: See Para. C.1.6

REQUIREMENT

C.1.4 Name and exact address of the site, including telephone, fax and 24-hour telephone numbers.

GUIDANCE

C.1.4.1 *Name of Company, Site Address and Mailing Address (if different from site address).*

C.1.4.2 *Telephone and Fax Nos. of contact person.*

C.1.4.3 *24-hour contact Telephone No.*

REQUIREMENT

C.1.5 Type of actual products manufactured on the site (see list at Appendix) and information about specifically toxic or hazardous substances handled,

mentioning the way they are manufactured (in dedicated facilities or on a campaign basis).

GUIDANCE

C.1.5.1 *Quote the type of products manufactured as described at Appendix, specifying if the product is manufactured under a contractual agreement with a contract giver.*

C.1.5.2 *Note any toxic, hazardous, highly sensitising substances handled e.g. antibiotics, hormones, cytostatics. Note whether the products are manufactured in a dedicated facility or on a campaign basis.*

C.1.5.3 *Mention if human and veterinary products are both prepared on the site.*

REQUIREMENT

C.1.6 Short description of the site (size, location and immediate environment and other manufacturing activities on the site).

GUIDANCE

(Not more than 250 words or one A4 sheet).

C.1.6.1 *Provide a map indicating the location of the manufacturing site and the surrounding area. Mark the site. Describe the surrounding area and use of properties near by.*

C.1.6.2 *The size of the site, types of buildings and their ages.*

C.1.6.3 *Other manufacturing activities on the site.*

REQUIREMENT

C.1.7 Number of employees engaged in quality assurance, production, quality control, storage and distribution.

GUIDANCE

(Note: Include employees working only part-time on full-time equivalent basis. Give the rate of the academic and non-academic persons)

C.1.7.1 *Quality Assurance*

C.1.7.2 Production

C.1.7.3 Quality Control

C.1.7.4 Storage & Distribution

C.1.7.5 Technical & Engineering Support Services

C.1.7.6 Total of the above

REQUIREMENT

C.1.8 Use of outside scientific, analytical or other technical assistance in relation to manufacture and analysis (if so, see Chapter 7 for details).

GUIDANCE

For each outside contractor (including contract testing laboratories), give:

C.1.8.1 Name, address, telephone no. and fax. no. of contractor.

C.1.8.2 Brief outline of the activity being undertaken in not more than 100 words or half an A4 sheet.

REQUIREMENT

C.1.9 Short description of the quality management system of the company responsible for manufacture.

GUIDANCE
(Not more than 750 words or three A4 sheets).

C.1.9.1 State the company's Quality Policy.

C.1.9.2 Define the responsibility of the Quality Assurance function.

C.1.9.3 Describe the elements of the QA system e.g. organisational structure, responsibilities, procedures, processes.

C.1.9.4 Describe the audit programmes (self-inspection or audits by external organisations undertaken).

C.1.9.5 *Describe how results are reviewed to demonstrate the adequacy of the quality system in relation to the objective i.e. quality, efficacy and safety of the product. (see also C.6.1.2)*

C.1.9.6 *Record if standards such as ISO 9001 etc are used by the company to assess its suppliers.*

C.1.9.7 *When suppliers of critical starting materials and packaging materials - actives, excipients, containers, closures and printed packaging materials are assessed, give details of how this is done.*

C.1.9.8 *Describe the release for sale procedure for finished products.*

Chapter 2

C.2 PERSONNEL

REQUIREMENT

C.2.1 Organisation chart showing the arrangements for quality assurance, including production and quality control.

C.2.2 Qualifications, experience and responsibilities of key personnel.

C.2.3 Outline of arrangements for basic and in-service training and how records are maintained.

C.2.4 Health requirements for personnel engaged in production.

C.2.5 Personnel hygiene requirements, including clothing.

GUIDANCE
(Not more than 500 words or two A4 sheets)

C.2.1 *Organogram for quality assurance including production and quality control. Record senior managers and supervisors only.*

C.2.2 *Brief details of academic qualifications, work-related qualifications and years of relevant experience since qualifying.*

C.2.3 Give brief details of the training programme and include induction and continuous training, as follows:

C.2.3.1 Describe how training needs are identified and by whom.

C.2.3.2 Give details of training relative to GMP requirements.

C.2.3.3 State the form of training e.g. in-house, external, and how practical experience is gained and which staff are involved.

C.2.3.4 Explain how the efficacy of the training is assessed e.g. by questionaire.

C.2.3.5 Explain how retraining needs are identified.

C.2.3.6 Give brief details of training records kept.

C.2.4 Health requirements for personnel engaged in production

C.2.4.1 Who is responsible for checking health of employees?

C.2.4.2 Is there a pre-employment medical examination?

C.2.4.3 Are employees routinely checked from time to time depending on the nature of their work?

C.2.4.4 Is there a system for reporting sickness or contact with sick people before working in a critical area?

C.2.4.5 Is there a system of reporting back after illness?

C.2.4.6 Are those who work in clean areas (Grade A-D) subject to additional monitoring?

C.2.5 *Personnel hygiene requirements including clothing*

C.2.5.1 Are there suitable washing, changing and rest areas?

C.2.5.2 Is the clothing suitable for the activity undertaken? Briefly describe the clothing.

C.2.5.3 *Are there clear instructions on how protective clothing should be used and when it should be changed? Detailed procedures are not needed. Is in-house or external laundry used?*

Chapter 3

C.3 PREMISES AND EQUIPMENT

REQUIREMENT

Premises

C.3.1 Simple layout plan and description of manufacturing areas with indication of scale (architectural or engineering drawings not required).

C.3.2 Nature of construction and finishes.

C.3.3 Brief description of ventilation systems. More details should be given for critical areas with potential risks of airborne contamination (including schematic drawings of the systems). Classification of the rooms used for the manufacture of sterile products should be mentioned.

C.3.4 Special areas for the handling of highly toxic, hazardous and sensitising materials.

C.3.5 Brief description of water systems (schematic drawings of the systems are desirable) including sanitation.

C.3.6 Maintenance (description of planned preventive maintenance programmes and recording system).

Equipment

C.3.7 Brief description of major production and quality control laboratories equipment (a list of the equipment is NOT required).

C.3.8 Maintenance (description of planned preventive maintenance programmes and recording system).

C.3.9 Qualification and calibration, including the recording system. Arrangements for computerized systems validation.

Sanitation

C.3.10 Availability of written specifications and procedures for cleaning manufacturing areas and equipment.

GUIDANCE

C.3.1 Layout of premises

C.3.1.1 Provide a site layout plan highlighting all production areas, warehouses, laboratories and other functional areas.

C.3.1.2 Provide a simple plan of each production area with indication of scale. Label areas and annotate plan with names.

C.3.1.3 Plans should be legible and on A4 sheets. Plans could be on A3 sheets if considered necessary.

C.3.1.4 For all controlled areas, particularly sterile product areas, indicate room and area classification and pressure differentials between adjoining areas of different classifications.

C.3.2 Nature of construction and finishes
(500 words or two A4 sheets).

C.3.2.1 To reduce narrative for a large complex plant, the details should be limited to critical areas.

C.3.2.2 These areas must include all processing areas, packaging areas and critical storage areas.

C.3.2.3 A narrative format is preferred.

C.3.3 Brief description of ventilation systems etc.
(500 words or two A4 sheets).

Note 1: More details should be given for critical areas with potential risks of airborne contamination. This will include sterile product areas as well as

areas for processing powders, granulation and tabletting. For sterile product areas a summary of the results of the most recent qualification/re-qualification should be given.

Note 2: *To reduce the narrative, schematic drawings should be used. The following data should be given:*

C.3.3.1 *Design criteria e.g.*
- *Specification of the air supply*
- *Temperature*
- *Humidity*
- *Pressure differentials and air change rate*
- *Single pass or recirculation (%)*

C.3.3.2 *Filter design and efficiency e.g.*
- *Bag 99% efficiency*
- *HEPA 99.997% efficiency*
- *Details of any alarms on the ventilation system should be given.*

C.3.3.3 *The limits for changing the filters should be given.*

C.3.3.4 *If DOP (dioctylphthalate) is introduced, the point must be shown.*

C.3.3.5 *Give the frequency of revalidation of the system.*

C.3.4 *Follow the same layout as C.3.1 above for description of special areas for the handling of highly toxic, hazardous and sensitizing materials.*

C.3.5 *Brief description of water system, including sanitation (500 words or two A4 sheets).*

Note : *Schematic drawings of the systems are preferred. The following information must appear:*

C.3.5.1 *The schematic drawing must go back to the city supply system.*

C.3.5.2 *The capacity of the system (maximum quantity produced per hour).*

C.3.5.3 *Construction materials of the vessels and pipework.*

C.3.5.4 *Specification of any filters in the system must be given.*

C.3.5.5 If water is stored and circulated, the temperature at the point of return.

C.3.5.6 The specification of the water produced
a) chemical
b) conductivity
c) microbiological

C.3.5.7 The sampling points and frequency of testing.

C.3.5.8 The procedure and frequency of sanitation.

C.3.6 Maintenance (250 words or one A4 sheet).

Note : For the purpose of this guide "maintenance" is carried out by the manufacturer and "servicing" is by an outside contractor.

C.3.6.1 Describe the planned preventive maintenance programme.

C.3.6.2 Are there written procedures and suitable reporting forms for maintenance and servicing? Do the documents record type/frequency of service/checks, details of service, repairs and modifications?

C.3.6.3 Are the maintenance routines that could affect product quality clearly identified?

C.3.6.4 Are the reports made known to the users?

C.3.7 Brief description of major production and control laboratory equipment (Not more than 250 words or one A4 sheet)

Note : Makes and model numbers of the equipment are not required. However the following points should be addressed.

C.3.7.1 Is the machinery constructed of appropriate material (e.g. AISI grade 316 stainless steel for product contact equipment)?

C.3.7.2 Have other materials been suitably validated e.g. polypropylene, chrome-plated brass, PVC, non-reactive plastic materials?

C.3.7.3 Is the equipment designed with ease of cleaning in mind?

C.3.7.4 Only a general description is required e.g. a rotary tablet press etc. If the equipment has additional devices, these should be recorded e.g. automatic weighing machines with printer, a labeller incorporating a bar code reader for the label; a lot number and expiry date over-printer; a freeze drier equipped with a steam sterilisation facility.

C.3.7.5 In the quality control laboratory only general descriptions such as pH meters, chromatographic equipment GLC, HPLC with computer systems, particle size analysers.

C.3.7.6 In microbiology use general descriptions such as incubators (temperature ranges), facilities for LAL testing, membrane filtration, sterility testing etc.

C.3.7.7 In particular give brief information on the use of computers, microprocessors etc. in the premise.

C.3.8 *Maintenance (250 words or one A4 sheet).*

C.3.8.1 Who is responsible for maintenance and servicing?

C.3.8.2 Are there written procedures and contractual details for outside work?

C.3.8.3 Are maintenance routines which could affect product quality clearly identified?

C.3.8.4 Are records kept of:
a) type and frequency of service/check
b) details of service repairs and modifications

C.3.8.5 Are reports made known to the users?

C.3.9 Qualification, Validation and Calibration (750 words or three A4 sheets).

C.3.9.1 *Briefly describe the company's general policy and protocols for qualification and validation (prospective and retrospective).*

C.3.9.2 *Is there regular revalidation of critical equipment?*

C.3.9.3 *An outline of process validation may be given here or cross-referenced to Production (Para. C.5.5).*

C.3.9.4 *Describe the system for the release for sale or supply of development and validation batches.*

C.3.9.5 *What are the arrangements for computer validation, including software validation?*

C.3.9.6 *Describe equipment calibration policy and records kept. (See also Para. C.4.2.8)*

C.3.10 Cleaning procedures for manufacturing areas and equipment
(250 words or one A4 sheet).

C.3.10.1 *Are there written specifications and procedures for cleaning, cleaning agents and their concentration for the method of cleaning and the frequency?*

C.3.10.2 *Are cleaning agents changed from time to time?*

C.3.10.3 *Have the cleaning procedures been validated and what was the method of evaluating the effectiveness of cleaning?*

C.3.10.4 *Are cleaning methods monitored routinely by chemical and/or microbiological methods?*

C.3.10.5 *What are the cleaning methods (and their frequency) for the water system, air handling system and dust extraction system?*

Chapter 4

C.4 DOCUMENTATION

REQUIREMENT

C.4.1 Arrangements for the preparation, revision and distribution of necessary documentation for manufacture, including storage of master documents.

C.4.2 Any other documents related to product quality which is not mentioned elsewhere (e.g. microbiological controls on air and water).

GUIDANCE
(500 words or two A4 sheets).

Note : This section refers to all documentation used in manufacture. Manufacture involves all activities relating to the production and control of pharmaceutical products.

C.4.1 Arrangement for the preparation, revision and distribution of documentation

C.4.1.1 Is there a description of the documentation system?

C.4.1.2 Who is responsible for the preparation, revision and distribution of documents?

C.4.1.3 Where are the master documents stored?

C.4.1.4 Is there a standard format and instruction of how documents are to be prepared?
a) Product/Process Specifications
b) Raw Material Specifications
c) Packaging Component Specifications
d) Standard Process Instructions including packaging
e) Batch Records (including Packaging)
f) Analytical Methods
g) *QA Release Procedures*

C.4.1.5 How is the documentation controlled?

C.4.1.6 For how long are the documents kept after the release of the batch?

C.4.1.7 Detail any arrangement for electronic or microfilmed records.

C.4.2 Other documentation related to product quality

Are the following documents available and in use?

C.4.2.1 Equipment Specifications
C.4.2.2 Specifications for disposables i.e. cleaning materials
C.4.2.3 Standard Operating Procedures
C.4.2.4 Quality Control Procedures
C.4.2.5 Training Procedures
C.4.2.6 Computer Programme Specifications
C.4.2.7 Documentation Control of Process Deviations
C.4.2.8 Calibration and Test Documents (see Para. C.3.9.6)
C.4.2.9 Validation Documents (see Para. C.3.9 and Para. C.5.5)
C.4.2.10 Reconciliation of batches of raw materials and major packaging components i.e. product-contact and printed materials.
C.4.2.11 List and briefly explain the use of any additional standard documentation used routinely.

Chapter 5

C.5 PRODUCTION

REQUIREMENT

C.5.1 Brief description of production operations using, wherever possible, flow sheets and charts specifying important parameters (see at Appendix the list of products manufactured)

C.5.2 Arrangements for the handling of starting materials, packaging materials, bulk and finished products, including sampling, quarantine, release and storage.

C.5.3 Arrangements for reprocessing or rework.

C.5.4 Arrangements for the handling of rejected materials and products.

C.5.5 Brief description of general policy for process validation.

GUIDANCE

C.5 Production

Note : This narrative should be kept to a minimum and generalized schematic layouts used where possible. The following points should be addressed.

C.5.1 Describe the operations capable of being carried out at the site with the existing facilities and specify the types of pharmaceutical products. (See Para 1.5.1 and the Appendix for types of products manufactured).

When only packaging is undertaken, give a brief description only, e.g. labelling, filling etc. and the nature of containers used e.g. sachets, tamper proof glass containers.

If cytotoxic or radio-active substances are handled, give details of the products.
Describe the production operations using flow charts if possible. Technical details are not required.

Describe how products are identified during production and how in-process storage is organised.

C.5.2 Arrangements for handling sterile materials, packing materials, bulk and finished products including sampling, quarantine, release and storage

a) Identification of suppliers' lot number with the company's lot number.
b) Sampling plans.
c) Status labelling e.g. by using labels or by computer.
d) Issue of materials to manufacture and package.
e) The control of weighing.
f) Checking methods.
g) How are materials being used for manufacture identified and released.

C.5.2.1 Control of manufacturing:

a) Checks on key parameters during manufacture e.g. blend times, filter integrity tests.
b) Records of key parameters.
c) In-process checks.
d) *Records of in-process checks.*
e) *Compliance with the Marketing Authorisation*

C.5.2.2 Packing

a) Release of bulk, semi-finished products, packing materials.

b) Confirmation of identity and line clearance checks.
c) In-process checks.

C.5.2.3 Quarantine and release of finished products; compliance with Marketing Authorisation. (see also Para. C.1.9.8)

C.5.2.4 Explain the role of the Authorised Person(s).

C.5.3 Arrangements for Reprocessing or Rework

C.5.3.1 What arrangements are in place for reprocessing or reworking batches of products?

C.5.4 Arrangements for Handling Rejected Materials and Products

C.5.4.1 Are rejected materials and products clearly labelled? Are they stored separately in restricted area?

C.5.4.2 Describe arrangements for sentencing the materials and their disposal. Is destruction recorded?

C.5.5 An outline of process validation protocol only is required (see Para 3.9.3).

Chapter 6

C.6 QUALITY CONTROL

REQUIREMENT

C.6.1 Description of the Quality Control system and of the activities of the Quality Control Department. Procedures for the release of finished products.

GUIDANCE

C.6.1 Activities of the Quality Control Department

C.6.1.1 (a) Describe the elements of the QC system e.g. specifications, test methods, and other quality related data collection.

(b) Briefly describe the activities of analytical testing, packaging, component testing, biological and microbiological testing.

C.6.1.2 *If the review of batch documentation and release of final documentation takes place in this department, give details (see Para. C.1.9.5).*

C.6.1.3 *Outline the involvement in the arrangements for the preparation, revision and distribution of documents in particular those for specification test methods and release criteria, if not mentioned elsewhere (see Chapter 4, Documentation).*

Chapter 7

C.7 CONTRACT MANUFACTURE AND ANALYSIS

REQUIREMENT

C.7.1 Description of the way in which the GMP compliance of the contract acceptor is assessed.

GUIDANCE

C.7.1 *Describe briefly the details of the technical contract between the contract giver and acceptor and the way in which the GMP compliance, or compliance with other appropriate standards, is assessed to ensure product compliance with the Marketing Authorisation/Product Licence. The selected standards should be assessed for the suitability of its application. The type of products manufactured by the contract acceptor should be specified.*

Chapter 8

C.8 DISTRIBUTION, COMPLAINTS AND PRODUCT RECALLS

REQUIREMENT

C.8.1 Arrangements and recording system for distribution.

C.8.2 Arrangements for the handling of complaints and product recalls.

GUIDANCE

C.8.1 A Description of Storage and Distribution Practices

a) Is the warehouse secure?
b) Is it environmentally controlled?
c) Is there refrigerated storage?
d) How are the materials stored e.g. pallet racking?
e) How is the status of the product controlled e.g. by computer, label?
f) What are the methods of distribution to customers?
g) Does the despatch order ensure first in first out (FIFO) and identify the lot number?

C.8.2 Records of Distribution

Do the retained records permit full batch traceability from the factory to the customer in terms of the date of sale, customer details and quantity despatched?

C.8.2.1 Complaints

C.8.2.1.1 Is there a written procedure for product complaints?

C.8.2.1.2 Who is responsible for:
a) Logging;
b) Classifying;
c) Investigating complaints

C.8.2.1.3 Are written reports prepared?

C.8.2.1.4 Who reviews these reports?

C.8.2.1.5 For how long are complaint records kept?

C.8.2.2 Product Recalls.

C.8.2.2.1 Is there a written procedure which describes the sequence of actions to follow including:
(a) Retrieval of distribution data;

(b) Notification of customers;
(c) Receipt/segregation/inspection of returned products;
(d) Investigation/reporting of cause;
(e) Reporting corrective action.

C.8.2.2.2 Who is responsible for coordinating product recalls?

C.8.2.2.3 Who notifies the Competent Authority of complaints and recalls?

C.8.2.2.4 Is the Competent Authority involved in complaints and the decision to recall?

C.8.2.2.5 Can recalls be effected below wholesale level?

Chapter 9

C.9 SELF INSPECTION

REQUIREMENT

C.9.1 Short description of the self-inspection system (see Para. C.1.9.4).

GUIDANCE

C.9.1.1 Describe how the self-inspection system verifies that those activities that have a bearing on product quality comply with the planned arrangement.

C.9.1.2 Are the quality systems effective?

C.9.1.3 Are there documented procedures for the self-inspection system and for the follow-up actions?

C.9.1.4 Are the results of the self-inspection system documented, brought to the attention of the personnel having responsibility for the area and activities inspected?

C.9.1.5 *Does the system ensure that those responsible for the area or activity take timely corrective action on the deficiencies found?*

REFERENCE

1. PIC/S PE 008-1 (effective 01 November 2002) : Explanatory Notes for Industry on the Preparation of a Site Master File

END OF DOCUMENT

APPENDIX

TYPE OF PRODUCTS MANUFACTURED
(referred to in paragraph C.1.5)

A. Sterile products

A.1 Liquid dosage forms (large volume solutions, including LVP and rinsing solutions)

A.1.1 Aseptically prepared

A.1.2 Terminally sterilized

A.2 Liquid dosage forms (small volume solutions, including SVP and eye drops)

A.2.1 Aseptically prepared

A.2.2 Terminally sterilized

A.3 Semi-solid dosage forms

A.4 Solid dosage forms

A.4.1 Solid fill

A.4.2 Freeze-dried

B. Non-sterile products

B.1 Liquid dosage forms

B.2 Semi-solid dosage forms

B.3 Solid dosage forms

B.3.1 Unit dose form (eg tablets, capsules, suppositories, pessaries)

B.3.2 Multi dose form (eg powders, granules)

C. Biological products

C.1 Vaccines

C.2 Sera

C.3 Blood products

C.4 Others (describe)

D. Specifically toxic and hazardous substances

D.1 Penicillins

D.2 Cephalosporins

D.3 Hormones

D.4 Cytostatics

D.5 Others (describe)

E. Packaging only

E.1 Liquid dosage forms

E.2 Semi-solid dosage forms

E.3 Solid dosage forms

F. Contract manufacturing (kind of products)

Company reported upon is:

F.1 Acceptor

F.2 Giver

G. Drugs for clinical trials

H. Others
Including products not subjected to registration/licensing by the Competent Authorities

(e.g. veterinary products, cosmetics, health/dietary supplements, etc)

GUIDANCE NOTES

Subject: PREPARATION OF A SITE MASTER FILE

REVISION HISTORY

Rev No.	DCC Ref No.	Author	Date of Revision	Remarks of Revision
001	-	-	01 Feb 1999	New.
002	-	-	05 May 1999	-
003	-	-	03 Aug 2001	Change of organisation name
004	-	Terrance Ong	01 Jan 2003	1) Change to include requirement that the site master file must be ring-bound and to delete requirement that each chapter should start on a new sheet. 2) Change of C1 to C9 to include the Guidance Notes for each Requirement and to reflect changes in the corresponding PIC/S document - Explanatory Notes for Industry on the Preparation of a Site Master File. 3) Update document format. 4) Add in reference document.
005	205	Terrance Ong	01 Mar 2004	1) Change 3D HSA logo to 2D HSA logo. 2) Change Centre name to Centre for Drug Administration 3) Detached Revision History from Guidance Notes.
006	253	Terrance Ong	06 Sep2004	1) Re-located Revision History to the main document.