Content

Editors' Introduction	9
Extended Editors' Introduction	11
Guide to Good Manufacturing Practice (GMP) for Medicinal Products for Human and Veterinary Use	
Introduction of the European Commission	12
Commission Directive 2003/94/EC of 8 October 2003 laying down the principles and guidelines of good manufacturing practice in respect of medicinal products for human use and investigational medicinal products for human use	14
Commission Directive 91/412/EEC of 23 July 1991 laying down the principles and guidelines of good manufacturing practice for veterinary medicinal products	2 3
Part I – Basic Requirements for Medicinal Products	29
Chapter 1: Pharmaceutical Quality System (In operation since 31 January 2013)	29
Chapter 2: Personnel (In operation since 16 February 2014)	35
Chapter 3: Premises and Equipment (In operation since 1 March 2015)	40
Chapter 4: Documentation (In operation since 30 June 2011)	44
Chapter 5: Production (In operation since 1 March 2015)	52
Chapter 6: Quality Control (In operation since 1 October 2014)	61
Chapter 7: Outsourced Activities (In operation since 31 January 2013)	67
Chapter 8: Complaints, Quality Defects and Product Recall (In operation since 1 March 2015)	70
Chapter 9: Self Inspection	75

Part II – Basic Requirements for Active Substances used as Starting Materials (In operation since 1 September 2014)	2
Table of Contents	
1 Introduction	••••
2 Quality Management	
3 Personnel	
4 Buildings and Facilities	
5 Process Equipment	
6 Documentation and Records	·
7 Materials Management	!
8 Production and In-Process Controls	!
9 Packaging and Identification Labelling of APIs and Intermediate	s :
10 Storage and Distribution	••••
11 Laboratory Controls	1
12 Validation	10
13 Change Control	10
14 Rejection and Re-Use of Materials	1
15 Complaints and Recalls	1
16 Contract Manufacturers (Including Laboratories)	1
17 Agents, Brokers, Traders, Distributors, Repackers, and Relabeller	s 1
18 Specific Guidance for APIs Manufactured by Cell Culture/Fermentation	1
19 APIs for Use in Clinical Trials	1
20 Glossary	1
Commission Delegated Regulation (EU) No 1252/2014 of 28 May 201	4
supplementing Directive 2001/83/EC with regard to principles and guidelines of GMP for active substances for medicinal products for human use	1

Part III – GMP related documents	131
Site Master File	132
ICH guideline Q9 on quality risk management (September 2015)	138
ICH guideline Q10 on pharmaceutical quality system (September 2015	5) 156
MRA Batch Certificate (1 June 2011)	173
Template for the 'written confirmation' for active substances exported to the European Union for medicinal products for human use, in accordance with Article 46b(2)(b) of Directive 2001/83/EC (Version 2 January 2013)) ''
Guideline on setting health based exposure limits for use in risk identification in the manufacture of different medicinal products in shared facilities (1 June 2015)	179
Guidelines of 19 March 2015 on the formalised risk assessment for ascertaining the appropriate good manufacturing practice for excipients of medicinal products for human use (Text with EEA relevance) (2015/C 95/02)	
Annexes	
Annex 1. Manufacture of Sterile Medicinal Products (corrected versi (In operation since 1 March 2009)	on) 193
Annex 2. Manufacture of Biological active substances and Medicinal Products for Human Use (In operation since 31 January 2013)	209
Annex 3. Manufacture of Radiopharmaceuticals (In operation since 1 March 2009)	237
Annex 4. Manufacture of Veterinary Medicinal Products other than Immunological Veterinary Medicinal Products	244
Annex 5. Manufacture of Immunological Veterinary Medicinal Products	246
Annex 6. Manufacture of Medicinal Gases (In operation since 31 July 2010)	255
Annex 7. Manufacture of Herbal Medicinal Products (In operation since 1 September 2009)	265
Annex 8. Sampling of Starting and Packaging Materials	270
Annex 9. Manufacture of Liquids, Creams and Ointments	272
Annex 10. Manufacture of Pressurised Metered Dose Aerosol Preparations for Inhalation	273

Annex 11.	Computerised Systems (In operation since 30 June 2011)	275
Annex 12.	Use of Ionising Radiation in the Manufacture of Medicinal Products	280
Annex 13.	Manufacture of Investigational Medicinal Products (In operation since 31 July 2010)	286
Annex 14.	Manufacture of Medicinal Products derived from Human Blood or Human Plasma (In operation since 30 November 2011)	304
Annex 15.	Qualification and Validation (Revised 30 March 2015 – In operation since 1 October 2015)	317
Annex 16.	Certification by a Qualified Person and Batch Release (Revised 12 October 2015 – Coming into operation from 15 April 2016)	. 330
Annex 17.	Parametric Release	339
Annex 18.	(Covered by Part II)	
Annex 19.	Reference and Retention Samples	342
Annex 20.	(Covered by Part III/Q9)	
Glossary .		346