

# Center For Drug Evaluation and Research List of Guidance Documents

Guidance documents represent the Agency's current thinking on a particular subject. They do not create or confer any rights for or on any person and do not operate to bind FDA or the public. An alternative approach may be used if such approach satisfies the requirements of the applicable statutes, regulations, or both. For information on a specific guidance document, please contact the originating office (see footnote 1 in recent guidances), or contact the Division of Drug Information in the Office of Training and Communications.

Office of Training and Communications  
Division of Drug Information  
10903 New Hampshire Avenue  
Silver Spring, MD 20993

Telephone: 301-796-3400  
Fax: 301-847-8714  
Internet (I): <http://www.fda.gov/cder/guidance/index.htm>  
E-mail: [druginfo@fda.hhs.gov](mailto:druginfo@fda.hhs.gov)

## Table of Contents (by Subject Category)

Advertising .....	Page 3
Biopharmaceutics .....	Page 4
Biosimilarity.....	Page 5
Chemistry .....	Page 5
Clinical Antimicrobial .....	Page 9
Clinical Medical .....	Page 12
Clinical Pharmacology .....	Page 19
CMC - Microbiology .....	Page 20
Combination Products (Drug/Device/Biologic) .....	Page 20
Current Good Manufacturing Practices/Compliance .....	Page 21
Drug Safety .....	Page 24
Electronic Submissions .....	Page 25
Generic Drug .....	Page 27
Good Review Practices .....	Page 29
ICH .....	Page 29
IND .....	Page 38
Industry Letters .....	Page 38
Labeling .....	Page 39
Modernization Act .....	Page 41
OTC .....	Page 43
Pharmacology/Toxicology .....	Page 44
Procedural .....	Page 46
Small Entity Compliance Guides .....	Page 51
User Fee .....	Page 52

**Advertising****Issued Date**

Aerosol Steroid Product Safety Information in Prescription Drug Advertising and Promotional Labeling (I)	1/12/1998
Consumer-Directed Broadcast Advertisements (I)	8/9/1999
Industry-Supported Scientific and Educational Activities (I)	12/3/1997
Product Name Placement, Size, and Prominence in Advertising & Promotional Labeling	1/25/2012

**Advertising Draft****Issued Date**

"Help-Seeking" and Other Disease Awareness Communications by or on Behalf of Drug and Device Firms (I)	2/10/2004
Accelerated Approval Products -- Submission of Promotional Materials (I)	3/26/1999
Brief Summary: Disclosing Risk Information in Consumer-Directed Print Advertisements(I)	2/10/2004
Direct-to-Consumer Television Advertisements -- FDAAA DTC Television Ad Pre-Dissemination Review Program	3/13/2012
Presenting Risk Information in Prescription Drug and Medical Device Promotion (I)	5/27/2009
Promoting Medical Products in a Changing Healthcare Environment; Medical Product Promotion by Healthcare Organizations or Pharmacy Benefits Management Companies (PBMs) (I)	1/5/1998

**Biopharmaceutics****Issued Date**

Bioanalytical Method Validation (I)	5/23/2001
Bioavailability and Bioequivalence Studies for Orally Administered Drug Products - General Considerations (Revised) (I)	3/19/2003
Corticosteroids, Dermatologic (topical) In Vivo (I)	6/2/1995
Dissolution Testing of Immediate Release Solid Oral Dosage Forms (I)	8/25/1997
Extended Release Oral Dosage Forms: Development, Evaluation, and Application of In Vitro/In Vivo Correlations (I)	9/26/1997
Food-Effect Bioavailability and Fed Bioequivalence Studies (I)	1/31/2003
Metaproterenol Sulfate and Albuterol Metered Dose Inhalers In Vitro (I)	6/27/1989
Statistical Approaches to Establishing Bioequivalence (I)	2/2/2001
Waiver of In Vivo Bioavailability and Bioequivalence Studies for Immediate Release Solid Oral Dosage Forms Based on a Biopharmaceutics Classification System (I)	8/31/2000

**Biopharmaceutics Draft****Issued Date**

Bioavailability and Bioequivalence Studies for Nasal Aerosols and Nasal Sprays for Local Action (I)	4/3/2003
---	----------

## **Biosimilarity Draft**

Quality Considerations in Demonstrating Biosimilarity to a Reference Protein Product	2/15/2012
Scientific Considerations in Demonstrating Biosimilarity to a Reference Product	2/15/2012
Guidance for Industry on Biosimilars: Q & As Regarding Implementation of the BPCI Act of 2009	2/15/2012

## **Chemistry, Manufacturing, and Controls (CMC)**

## **Issued Date**

Botanical Drug Products (I)	6/9/2004
Changes to an Approved Application for Specified Biotechnology and Specified Synthetic Biological Products (I)	7/24/1997
Changes to an Approved NDA or ANDA (Revised) (I)	4/8/2004
Changes to an Approved NDA or ANDA: Questions and Answers (I)	1/22/2001
Changes to an Approved New Drug Application or Abbreviated New Drug Application; Specifications -Use of Enforcement Discretion for Compndial Changes (I)	11/22/2004
Container Closure Systems for Packaging Human Drugs and Biologics (I)	7/7/1999
Demonstration of Comparability of Human Biological Products Including Therapeutic Biotechnology Derived Products (I)	3/26/1996
Development of New Stereoisomeric Drugs (I)	5/1/1992

Drug Master Files (I)	9/1/1989
Drug Master Files for Bulk Antibiotic Drug Substances (I)	11/29/1999
Environmental Assessment of Human Drug and Biologics Applications (I)	7/27/1998
Format and Content for the CMC Section of an Annual Report (I)	9/1/1994
Incorporation of Physical-Chemical Identifiers into Solid Oral Dosage Form Drug Products for Anticounterfeiting (I)	10/11/2011
IND Meetings for Human Drugs and Biologics; Chemistry, Manufacturing, and Controls Information (I)	5/25/2001
INDs for Phase 2 and 3 Studies; Chemistry, Manufacturing, and Controls Information (I)	5/20/2003
Limiting the Use of Certain Phthalates as Excipients in CDER-Regulated Products	12/6/2012
Monoclonal Antibodies Used as Reagents in Drug Manufacturing (I)	3/29/2001
Nasal Spray and Inhalation Solution, Suspension, and Spray Drug Products -- Chemistry, Manufacturing, and Controls Documentation (I)	7/5/2002
NDAs: Impurities in Drug Substances (I)	2/25/2000
Orally Disintegrating Tablets (I)	12/16/2008
PAC-ALTS: Postapproval Changes - Analytical Testing Laboratory Sites (I)	4/28/1998
Residual Drug in Transdermal and Related Drug Delivery Systems (I)	8/16/2011

Residual Solvents in Drug Products Marketed in the United States	11/25/2009
Size of Beads in Drug Products Labeled for Sprinkle	5/2/2012
Submitting Documentation for the Manufacturing of and Controls for Drug Products* (I)	2/1/1987
Submitting Samples and Analytical Data for Methods Validation* (I)	2/1/1987
Submitting Supporting Documentation in Drug Applications for the Manufacture of Drug Products (I)	2/1/1987
SUPAC-IR Immediate-Release Solid Oral Dosage Forms: Scale-Up and Post-Approval Changes: Chemistry, Manufacturing and Controls, In Vitro Dissolution Testing, and In Vivo Bioequivalence Documentation (I)	11/30/1995
SUPAC-IR Questions and Answers (I)	2/18/1997
SUPAC-IR/MR: Immediate Release and Modified Release Solid Oral Dosage Forms, Manufacturing Equipment Addendum (I)	2/26/1999
SUPAC-MR: Modified Release Solid Oral Dosage Forms: Scale-Up and Postapproval Changes: Chemistry, Manufacturing, and Controls, In Vitro Dissolution Testing, and In Vivo Bioequivalence Documentation (I)	10/6/1997
SUPAC-SS - Nonsterile Semisolid Dosage Forms; Scale-Up and Postapproval Changes: Chemistry, Manufacturing, and Controls; In Vitro Release Testing and In Vivo Bioequivalence Documentation (I)	6/13/1997
Tablet Scoring: Nomenclature, Labeling, and Data for Evaluation	3/13/2013
The Sourcing and Processing of Gelatin to Reduce the Potential Risk Posed by Bovine Spongiform (I)	12/20/2000
Validation of Chromatographic Methods -- Reviewer's Guidance (I)	11/1/1994

**Chemistry, Manufacturing, and Controls (CMC) Draft****Issued Date**

Analytical Procedures and Methods Validation (I)	8/30/2000
Assay Development for Immunogenicity Testing of Therapeutic Proteins (I)	12/4/2009
CMC Postapproval Manufacturing Changes Reportable in Annual Reports (I)	6/25/2010
Comparability Protocols - Chemistry, Manufacturing, and Controls Information (I)	2/25/2003
Drugs, Biologics, and Medical Devices Derived From Bioengineered Plants for Use in Humans and Animals (I)	9/12/2002
Interpreting Sameness of Monoclonal Antibody Products Under the Orphan Drug Regulations (I)	7/26/1999
Liposome Drug Products: Chemistry, Manufacturing, and Controls; Human Pharmacokinetics and Bioavailability; and Labeling Documentation (I)	8/21/2002
Metered Dose Inhalers (MDI) and Dry Powder Inhalers (DPI) Drug Products (I)	11/19/1998
Non-Penicillin Beta-Lactam Risk Assessment: A CGMP Framework	3/15/2011
Regulatory Classification of Pharmaceutical Co-Crystals	12/1/2011
SUPAC-SS: Nonsterile Semisolid Dosage Forms Manufacturing Equipment Addendum (I)	1/5/1999

**Clinical Antimicrobial****Issued Date**



Acute Bacterial Exacerbations of Chronic Bronchitis in Patients With Chronic Obstructive Pulmonary Disease: Developing Antimicrobial Drugs for Treatment	10/1/2012
Acute Bacterial Otitis Media: Developing Drugs for Treatment	10/2/2012
Acute Bacterial Sinusitis: Developing Drugs for Treatment	10/9/2012
Antiretroviral Drugs Using Plasma Human Immunodeficiency Virus Ribonucleic Acid Measurements -Clinical Considerations for Accelerated and Traditional Approval (I)	11/1/2002
Antiviral Product Development -Conducting and Submitting Virology Studies to the Agency	6/5/2006
Antibacterial Drug Products: Use of Noninferiority Trials to Support Approval	11/29/2010
Clinical Development and Labeling of Anti-Infective Drug Products (I)	10/26/1992
Clinical Evaluation of Anti-Infective Drugs (Systemic) (I)	9/1/1977
Influenza: Developing Drugs for Treatment and/or Prophylaxis	4/13/2011
Role of HIV Drug Resistance Testing in Antiretroviral Drug Development (I)	10/31/2007

### **Clinical Antimicrobial Draft**

### **Issued Date**

Acute Bacterial Meningitis; Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Acute Bacterial Skin and Skin Structure Infections: Developing Drugs for Treatment	7/22/1998

Acute or Chronic Bacterial Prostatitis; Developing Antimicrobial Drugs for Treatment (I)	8/27/2010
Antiviral Product Development — Conducting and Submitting Virology Studies to the Agency Guidance for Submitting HCV Resistance Data -Attachment to Guidance	
Bacterial Vaginosis; Developing Antimicrobial Drugs for Treatment (I)	10/15/2007
Catheter-Related Bloodstream Infections - Developing Antimicrobial Drugs for Treatment (I)	10/18/1999
Chronic Hepatitis C Virus Infection: Developing Direct-Acting Antiviral Agents for Treatment	9/14/2010
Community Acquired Pneumonia; Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Community-Acquired Bacterial Pneumonia: Developing Drugs for Treatment (I)	3/20/2009
Complicated Intra-Abdominal Infections: Developing Drugs for Treatment	10/1/2012
Complicated Urinary Tract Infections: Developing Drugs for Treatment	2/24/2012
Developing Antimicrobial Drugs -General Considerations for Clinical Trials (I)	7/22/1998
Developing Antimicrobial Drugs to Treat Inhalational Anthrax (Post-Exposure) (I)	3/18/2002
Empiric Therapy of Febrile Neutropenia; Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Evaluating Clinical Studies of Antimicrobials in the Division of Anti-Infective Drug Products (I)	2/17/1997
Helicobacter pylori-Associated Duodenal Ulcer Disease in Adults: Developing Drugs for Treatment (I)	10/5/2009

Hospital-Acquired Bacterial Pneumonia and Ventilator-Associated Bacterial Pneumonia: Developing Drugs for Treatment	11/29/2010
Lyme Disease; Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Microbiological Data for Systemic Antibacterial Drug Products - Development, Analysis, and Presentation (I)	9/17/2009
Neglected Tropical Diseases of the Developing World: Developing Drugs for Treatment or Prevention	8/23/2011
Nosocomial Pneumonia - Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Secondary Bacterial Infections of Acute Bronchitis - Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Smallpox (Variola) Infection: Developing Drugs for Treatment or Prevention (I)	11/23/2007
Streptococcal Pharyngitis and Tonsillitis; Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Uncomplicated and Complicated Skin and Skin Structure Infections; Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Uncomplicated Gonorrhea -- Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Uncomplicated Urinary Tract Infections - Developing Antimicrobial Drugs for Treatment (I)	7/22/1998
Vaccinia Virus -- Developing Drugs to Mitigate Complications From Smallpox Vaccination (I)	3/9/2004
Vaginal Microbicides: Development for the Prevention of HIV Infection	11/23/2012
Vulvovaginal Candidiasis - Developing Antimicrobial Drugs for Treatment (I)	7/22/1998

<b><u>Clinical Medical</u></b>	<b><u>Issued Date</u></b>
Acceptance of Foreign Clinical Studies (I)	3/13/2001
Antianxiety Drugs -- Clinical Evaluation (I)	9/1/1977
Antidepressant Drugs -- Clinical Evaluation (I)	9/1/1977
Antiepileptic Drugs (adults and children) -- Clinical Evaluation (I)	1/1/1981
Available Therapy (I)	7/23/2004
Calcium DTPA and Zinc DTPA Drug Products -- Submitting a New Drug Application (I)	8/13/2004
Cancer Drug and Biological Products - Clinical Data in Marketing Applications (I)	10/5/2001
Chronic Cutaneous Ulcer and Burn Wounds - Developing Products for Treatment (I)	6/2/2006
Clinical and Statistical Sections of an Application --Format and Content* (I)	7/1/1988
Clinical Development Programs for Drugs, Devices, and Biological Products for the Treatment of Rheumatoid Arthritis (RA) (I)	2/17/1999
Clinical Endpoints for the Approval of Cancer Drugs and Biologics (I)	5/16/2007
Collection of Race and Ethnicity Data in Clinical Trials for FDA Regulated Products (I)	9/19/2005

Content and Format of Investigational New Drug Applications (INDs) for Phase 1 Studies of Drugs, Including Well-Characterized, Therapeutic, Biotechnology-Derived Products (I)	11/20/1995
Developing Medical Imaging Drug and Biological Products, Part 1: Conducting Safety Assessments (I)	6/22/2004
Developing Medical Imaging Drug and Biological Products, Part 2: Clinical Indications (I)	6/22/2004
Developing Medical Imaging Drug and Biological Products, Part 3: Design, Analysis, and Interpretation of Clinical Studies (I)	6/22/2004
Development and Use of Risk Minimization Action Plans (I)	3/29/2005
Development of Vaginal Contraceptive Drugs (NDA) (I)	3/2/1988
Diabetes Mellitus -- Evaluating Cardiovascular Risk in New Antidiabetic Therapies to Treat Type 2 Diabetes (I)	12/19/2008
Establishing Pregnancy Exposure Registries (I)	9/23/2002
Evaluating the Risks of Drug Exposure in Human Pregnancies	4/28/2005
Evaluation of the Effects of Orally Inhaled and Intranasal Corticosteroids on Growth in Children (I)	3/6/2007
Exocrine Pancreatic Insufficiency Drug Products-Submitting New Drug Applications (I)	4/14/2006
FDA Approval of New Cancer Treatment Uses for Marketed Drug and Biological Products (I)	2/2/1999
FDA Requirements for Approval of Drugs to Treat Non-Small Cell Lung Cancer (I)	1/29/1991
Formatting, Assembling and Submitting New Drug and Antibiotic Applications* (I)	2/1/1987

General Anesthetics -- Clinical Evaluation (I)	5/1/1982
General Considerations for the Clinical Evaluation of Drugs (I)	12/1/1978
General Considerations for the Clinical Evaluation of Drugs in Infants and Children (I)	9/1/1977
Good Pharmacovigilance Practices and Pharmacoepidemiologic Assessment (I)	3/29/2005
Guidance for Institutional Review Boards, Clinical Investigators, and Sponsors: Exception from Informed Consent Requirements for Emergency Research	4/4/2011
Hypnotic Drugs -- Clinical Evaluation (I)	9/1/1977
IND Exemptions for Studies of Lawfully Marketed Drug or Biological Products for the Treatment of Cancer (Revised) (I)	1/15/2004
Integration of Dose-Counting Mechanisms Into Metered-Dose Inhaler Drug Products (I)	3/13/2003
Internal Radioactive Contamination - Development of Decorporation Agents (I)	3/2/2006
Irritable Bowel Syndrome -- Clinical Evaluation of Products for Treatment	5/31/2012
Investigational New Drug Applications for Positron Emission Tomography (PET) Drugs	12/4/2012
Levothyroxine Sodium Tablets -- In Vivo Pharmacokinetic and Bioavailability Studies and In Vitro Dissolution Testing (I)	3/8/2001
Local Anesthetics -- Clinical Evaluation (I)	5/1/1982
MDI and DPI Drug Products -- Clinical Development and Programs (I)	9/19/1994

Oncologic Drugs Advisory Committee Discussion on FDA Requirements for Approval of New Drugs for Treatment of Colon and Rectal Cancer (I)	4/19/1988
Patient-Reported Outcome Measures: Use in Medical Product Development to Support Labeling Claims (I)	12/9/2009
Pediatric Use Supplements --Content and Format (I)	5/24/1996
Postmarketing Adverse Experience Reporting for Human Drugs and Licensed Biological Products; Clarification of What to Report (I)	8/27/1997
Postmarketing Reporting of Adverse Drug Experiences (I)	3/1/1992
Premarketing Risk Assessment (I)	3/29/2005
Providing Clinical Evidence of Effectiveness for Human Drug and Biological Products (I)	5/15/1998
Prussian Blue for Treatment of Internal Contamination With Thallium or Radioactive Cesium (I)	2/4/2003
Psychoactive Drugs in Infants and Children -- Clinical Evaluation (I)	7/1/1979
Study and Evaluation of Gender Differences in the Clinical Evaluation of Drugs (I)	7/22/1993
Study of Drugs Likely to be Used in the Elderly (I)	11/1/1989
Submission of Abbreviated Reports and Synopses in Support of Marketing Applications (I)	9/13/1999
Summary for New Drug and Antibiotic Applications -- Format and Content* (I)	2/1/1987
Systemic Lupus Erythematosus - Developing Drugs for Treatment (I)	6/22/2010

The Radioactive Drug Research Committee: Human Research Without An Investigational New Drug Application (I)

8/2/2010

<b><u>Clinical Medical Draft</u></b>	<b><u>Issued Date</u></b>
Abuse-Deterrent Opioids-Evaluation and Labeling	1/14/2013
Acne Vulgaris: Developing Drugs for Treatment (I)	9/19/2005
Allergic Rhinitis: Clinical Development Programs for Drug Products (I)	6/21/2000
Alzheimer's Disease: Developing Drugs for the Treatment of Early Stage Disease	2/8/2013
Assessment of Abuse Potential of Drugs (I)	1/27/2010
Chronic Obstructive Pulmonary Disease: Developing Drugs for Treatment (I)	11/9/2007
Clinical Development Programs for Drugs, Devices, and Biological Products Intended for the Treatment of Osteoarthritis (OA) (I)	7/15/1999
Clinical Trial Endpoints for the Approval of Non-Small Cell Lung Cancer Drugs and Biologics	6/16/2011
Clinical Trial Sponsors on the Establishment and Operation of Clinical Trial Data Monitoring Committees (I)	11/20/2001
Codevelopment of Two or More Unmarketed Investigational Drugs for Use in Combination	12/14/2010
Combination Products Timeliness of Premarket Reviews (I)	5/4/2004



Computerized Systems Used in Clinical Trials (I)	10/4/2004
Determining the Extent of Safety Data Collection Needed in Late Stage Premarket and Postapproval Clinical Investigations	2/109/2012
Developing Products for Weight Management Revision 1 (I)	2/15/2007
Development of Parathyroid Hormone for the Prevention and Treatment of Osteoporosis (I)	6/14/2000
Diabetes Mellitus: Developing Drugs and Therapeutic Biologics for Treatment and Prevention (I)	3/3/2008
Drugs, Biologics, and Medical Devices Derived from Bioengineered Plants for Use in Humans and Animals	9/12/2002
Enrichment Strategies for Clinical Trials to Support Approval of Human Drugs and Biological Products	12/14/2012
Estrogen and Estrogen/ Progestin Drug Products to Treat Vasomotor Symptoms and Vulvar and Vaginal Atrophy Symptoms - Recommendations for Clinical Evaluation (I)	1/31/2003
Exercise-Induced Bronchospasm (EIB) - Development of Drugs to Prevent EIB (I)	2/20/2002
Gingivitis: Development and Evaluation of Drugs for Treatment or Prevention (I)	6/28/2005
Immunogenicity Assessment for Therapeutic Protein Products	2/11/2013
Inhalation Drug Products Packaged in Semipermeable Container Closure Systems (I)	7/26/2002
Investigational New Drug Applications (INDs)-Determining Whether Human Research Studies Can Be Conducted Without an IND (I)	10/14/2010
Malaria: Developing Drug and Nonvaccine Biological Products for Treatment and Prophylaxis (I)	6/7/2007

OTC Treatment of Herpes Labialis with Antiviral Agents (I)	3/8/2000
Pathologic Complete Response in Neoadjuvant Treatment of High-Risk Early-Stage Breast Cancer: Use as an Endpoint to Support Accelerated Approval	5/30/2012
Pediatric Oncology Studies in Response to a Written Request (I)	6/21/2000
Qualification Process for Drug Development Tools (I)	10/25/2010
Sinusitis: Designing Clinical Development Programs of Nonantimicrobial Drugs for Treatment (I)	11/22/2006
Standards for Clinical Trial Imaging Endpoints	8/18/2011
Suicidal Ideation and Behavior: Prospective Assessment of Occurrence in Clinical Trials	8/15/2012
The Use of Clinical Holds Following Clinical Investigator Misconduct (I)	9/2/2004

### **Clinical Pharmacology**

### **Issued Date**

Clinical Pharmacogenomics: Premarket Evaluation in Early-Phase Clinical Studies and Recommendations for Labeling	1/28/2013
Exposure-Response Relationships - Study Design, Data Analysis, and Regulatory Applications (I)	5/6/2003
Format and Content of the Human Pharmacokinetics and Bioavailability Section of an Application (I)	2/1/1987
Pharmacokinetics in Patients With Impaired Hepatic Function; Study Design, Data Analysis, and Impact on Dosing and Labeling (I)	5/30/2003

Pharmacokinetics in Patients with Impaired Renal Function: Study Design, Data Analysis, and Impact on Dosing and Labeling (I)	5/15/1998
---	-----------

Population Pharmacokinetics (I)	2/10/1999
---------------------------------	-----------

### **Clinical Pharmacology Draft**

### **Issued Date**

Clinical Lactation Studies - Study Design, Data Analysis and Recommendations for Labeling	2/8/2005
---	----------

Drug Interaction Studies--Study Design, Data Analysis, Implications for Dosing, and Labeling Recommendations	2/21/2012
--	-----------

General Considerations for Pediatric Pharmacokinetic Studies for Drugs and Biological Products (I)	
--	--

Pharmacokinetics in Pregnancy - Study Design, Data Analysis, and Impact on Dosing and Labeling (I)	11/1/2004
--	-----------

### **CMC Microbiology**

### **Issued Date**

Submission Documentation for Sterilization Process Validation Applications for Human and Veterinary Drug Products (I)	11/1/1994
---	-----------

### **CMC Microbiology Draft**

### **Issued Date**

Submission of Documentation in Applications for Parametric Release of Human and Veterinary Drug Products Terminally Sterilized by Moist Heat Processes (I)	8/5/2008
--	----------

**Combination Products (Drug/Device/Biologic)**

**Issued Date**

Application User Fees for Combination Products

4/21/2005

**Combination Products (Drug/Device/Biologic) Draft**

**Issued Date**

Coronary Drug-Eluting Stents-Nonclinical and Clinical Studies (I)

3/27/2008

**Current Good Manufacturing Practices/Compliance**

**Issued Date**

A Review of FDA's Implementation of the Drug Export Amendments of 1986 (I)

5/1/1990

Bar Code Label Requirements - Questions and Answers (Revised Aug 2011)

10/5/2006

Compressed Medical Gases (I)

12/1/1989

Computerized Systems Used in Clinical Trials (I)

5/10/1999

Current Good Manufacturing Practice for Phase 1 Investigational Drugs (I)

7/15/2008

Current Good Manufacturing Practice for Positron Emission Tomography Drug Products (I)

12/10/2009

Dosage Delivery Devices for Orally Ingested OTC Liquid Drug Products	5/5/2011
Expiration Dating and Stability Testing of Solid Oral Dosage Form Drugs Containing Iron (I)	6/27/1997
Formal Dispute Resolution: Scientific and Technical Issues Related to Pharmaceutical Current Good Manufacturing Practices (I)	1/12/2006
Good Laboratory Practice Regulations -- Questions and Answers (I)	6/1/1981
Guidance for Hospitals, Nursing Homes, and Other Health Care Facilities (I)	4/6/2001
Investigating Out of Specification (OOS) Test Results for Pharmaceutical Production (I)	10/12/2006
Marketed Unapproved Drugs; Compliance Policy Guide (I)	9/19/2011
Media Fills for Validation of Aseptic Preparations for Positron Emission Tomography	4/11/2012
Nuclear Pharmacy Guideline Criteria for Determining When to Register as a Drug Establishment (I)	5/1/1984
Part 11, Electronic Records, Electronic Signatures - Scope and Application	9/5/2003
PET Drugs — Current Good Manufacturing Practice (CGMP)	8/4/2011
Pharmaceutical Components at Risk for Melamine Contamination (I)	8/7/2009
Pharmacy Compounding -- Compliance Policy Guide (I)	6/7/2002
Possible Dioxin/PCB Contamination of Drug and Biological Products (I)	8/23/1999

Preparation of Investigational New Drug Products (Human and Animal)	11/1/1992
Prescription Drug Marketing Act Regulations for Donation of Prescription Drug Samples to Free Clinics (I)	3/14/2006
Process Analytical Technology -- A Framework for Innovative Pharmaceutical Manufacturing and Quality Assurance (I)	10/4/2004
Process Validation: General Principles and Practices	1/25/2011
Pyrogen and Endotoxins Testing: Questions and Answers	6/28/2012
Quality Systems Approach to Pharmaceutical Current Good Manufacturing Practice Regulations (I)	10/2/2006
Sterile Drug Products Produced by Aseptic Processing (I)	10/4/2004
Street Drug Alternatives (I)	4/3/2000
Testing of Glycerin for Diethylene Glycol (I)	5/2/2007
The Use of Mechanical Calibration of Dissolution Apparatus 1 and 2 - Good Manufacturing Practice (CGMP) (I)	1/27/2010

**Current Good Manufacturing Practices/Compliance Draft**

**Issued Date**

Comparability Protocols -- Protein Drug Products and Biological Products -- Chemistry, Manufacturing, and Controls Information (I)	9/5/2003
Current Good Manufacturing Practices for Combination Products (I)	10/4/2004

Current Good Manufacturing Practices for Medical Gases (3rd Revision) (I)	5/6/2003
Expiration Dating of Unit-Dose Repackaged Drugs: Compliance Policy Guide	5/31/2005
Heparin for Drug and Medical Device Use: Monitoring Crude Heparin for Quality	2/13/2012
Manufacturing, Processing or Holding of Active Pharmaceutical Ingredients (I)	4/17/1998
Powder Blends and Finished Dosage Units--Stratified In-Process Dosage Unit Sampling and Assessment (I)	11/7/2003

### **Drug Safety**

### **Issued Date**

Drug Safety Information--Food and Drug Administration's Communication to the Public (I)	3/7/2007
Drug-Induced Liver Injury: Premarketing Clinical Evaluation (I)	7/30/2009
Medication Guides--Distribution Requirements and Inclusion of Medication Guides in Risk Evaluation and Mitigation Strategies	11/17/2011
Postmarketing Adverse Event Reporting for Medical Products and Dietary Supplements During an Influenza Pandemic	2/24/2012
Postmarketing Studies and Clinical Trials--Implementation of Section 505(o)(3) of the Federal Food, Drug, and Cosmetic Act	4/1/2011
Safety Reporting Requirements for INDs and BA/BE Studies- Small Entity Compliance Guide	12/20/2012
Safety Reporting Requirements for INDs (Investigational New Drug Applications) and BA/BE (Bioavailability/Bioequivalence) Studies	12/20/2012

### **Drug Safety Draft**

### **Issued Date**

Best Practices for Conducting and Reporting Pharmacoepidemiologic Safety Studies Using Electronic Healthcare Data Sets	2/16/2011
Classifying Significant Posmarketing Drug Safety Issues	3/9/2012
Drug Safety Information -- FDA's Communication to the Public	3/9/2012
Format and Content of Proposed Risk Evaluation and Mitigation Strategies (REMS), REMS Assessments, and Proposed REMS Modifications (I)	10/1/2009
Safety Considerations for Product Design to Minimize Medication Errors	12/13/2012
Safety Labeling Changes; Implementation of the Federal Food, Drug, and Cosmetic Act	4/13/2011

### **Electronic Submissions**

### **Issued Date**

Indexing Structured Product Labeling (I)	6/2/2008
Providing Regulatory Submissions in Electronic Format -- Content of Labeling (I)	4/21/2005
Providing Regulatory Submissions in Electronic Format -Human Pharmaceutical Product Applications and Related Submissions Using the eCTD Specifications (R2)	6/1/2008
Regulatory Submissions in Electronic Format; General Considerations (I)	1/28/1999



**Electronic Submissions Draft****Issued Date**

Compliance Policy on Reporting Drug Sample Distribution Information	4/3/2012
Providing Regulatory Submissions in Electronic Format--General Considerations (I)	10/22/2003
Providing Regulatory Submissions in Electronic Format — Certain Human Pharmaceutical Product Applications and Related Submissions Using the eCTD Specifications	1/3/2013
Providing Regulatory Submissions in Electronic Format - Postmarketing Expedited Safety Reports (I)	5/4/2001
Providing Regulatory Submissions in Electronic Format - Postmarketing Individual Case Safety Reports (I)	6/12/2008
Providing Regulatory Submissions in Electronic Format - Prescription Drug Advertising and Promotional Labeling (I)	1/31/2001
Providing Regulatory Submissions in Electronic Format--Receipt Date (I)	6/5/2007
Providing Submissions in Electronic Format -- Standardized Study Data	2/21/2012
Providing Submissions in Electronic Format -- Summary Level Clinical Site Data for CDER	12/19/2012
Specifications for Preparing and Submitting Summary Level Clinical Site Data for CDER's Inspection Planning	12/19/2012

**Generic Drug****Issued Date**

180-Day Exclusivity When Multiple Abbreviated New Drug Applications Are Submitted on the Same Day (I)	8/1/2003
Abbreviated New Drug Applications: Impurities in Drug Products	11/29/2010
Alternate Source of Active Pharmaceutical Ingredients in Pending ANDAs (I)	12/12/2000
ANDAs: Impurities in Drug Substances; Chemistry, Manufacturing and Controls Information (I)	7/15/2009
ANDAs: Pharmaceutical Solid Polymorphism; Chemistry, Manufacturing and Controls Information (I)	7/9/2007
Court Decisions, ANDA Approvals, and 180-Day Exclusivity Under the Hatch-Waxman Amendments to the Federal Food, Drug, and Cosmetic Act (I)	3/30/2000
Handling and Retention of Bioavailability and Bioequivalence Testing Samples (I)	5/26/2004
Individual Product Bioequivalence Recommendations - List of Product Bioequivalence Recommendations (I)	6/11/2010
Letter announcing that the OGD will now accept the ICH long-term storage conditions as well as the stability studies conducted in the past (I)	8/18/1995
Letter describing efforts by the CDER & the ORA to clarify the responsibilities of CDER chemistry review scientists and ORA field investigators in the new & abbreviated drug approval process in order to reduce duplication or redundancy in the process (I)	10/14/1994
Letter on incomplete Abbreviated Applications, Convictions Under GDEA, Multiple Supplements, Annual Reports for Bulk Antibiotics, Batch Size for Transdermal Drugs, Bioequivalence Protocols, Research, Deviations from OGD Policy (I)	4/8/1994
Letter on the provision of new information pertaining to new bioequivalence guidelines and refuse-to-file letters (I)	7/1/1992
Letter on the provision of new procedures and policies affecting the generic drug review process (I)	3/15/1989
Letter on the request for cooperation of regulated industry to improve the efficiency and effectiveness of the generic drug review process, by assuring the completeness and accuracy of required information and data submissions (I)	11/8/1991

Letter on the response to 12/20/84 letter from the Pharmaceutical Manufacturers Association about the Drug Price Competition and Patent Term Restoration Act (I)	3/26/1985
Letter to all ANDA and AADA applicants about the Generic Drug Enforcement Act of 1992 (GDEA), and the Office of Generic Drugs intention to refuse-to-file incomplete submissions as required by the new law (I)	1/15/1993
Letter to regulated industry notifying interested parties about important detailed information regarding labeling, scale-up, packaging, minor/major amendment criteria, and bioequivalence requirements (I)	8/4/1993
Major, Minor, and Telephone Amendments to Abbreviated New Drug Applications (I)	12/21/2001
Revising ANDA Labeling Following Revision of the RLD Labeling (I)	4/25/2000
Submission of Summary Bioequivalence Data for Abbreviated New Drug Applications (I)	5/6/2011
Variations in Drug Products that May Be Included in a Single ANDA (I)	1/27/1999

### **Generic Drug Draft**

### **Issued Date**

Initial Completeness Assessments for Type II API DMFs Under GDUFA	10/2/2012
Listed Drugs, 30-Month Stays, and Approval of ANDAs and 505 (b)(2) Applications Under Hatch Waxman, as Amended by the Medicare Prescription Drug Improvement, and Modernization Act of 2003 - Questions and Answers (I)	11/4/2004
Generic Drug User Fee Amendments of 2012: Questions and Answers	8/27/2012
Self-Identification of Generic Drug Facilities, Sites, and Organizations	8/27/2012
ANDAs: Stability Testing of Drug Substances and Products	9/24/2012

**Good Review Practices****Issued Date**

Pharmacology/Toxicology Review Format (I)

5/10/2001

**ICH - Efficacy****Issued Date**

E1A - The Extent of Population Exposure to Assess Clinical Safety: for Drugs Intended for Long Term Treatment of Non-Life-Threatening Conditions (I)

3/1/1995

E2A - Clinical Safety Data Management: Definitions and Standards for Expedited Reporting (I)

3/1/1995

E2B - Data Elements for Transmission of Individual Case Safety Reports (I)

1/15/1998

E2B(M) - Data Elements for Transmission of Individual Case Safety Reports (Revised) (I)

4/3/2002

E2B(M): Data Elements for Transmission of Individual Case Safety Reports -- Questions and Answers (Revision 2) (I)

3/9/2005

E2C - Clinical Safety Data Management: Periodic Safety Update Reports for Marketed Drugs (I)

5/19/1997

E2C Addendum - Clinical Safety Data Management: Periodic Safety Update Reports for Marketed Drugs (I)

2/5/2004

E2E - Pharmacovigilance Planning (I)

4/1/2005

E2F Development Safety Update Report (I)

8/22/2011

E3 - Structure and Content of Clinical Study Reports (I)	7/17/1996
E3 Structure and Content of Clinical Study Reports - Questions and Answers (R1)	1/25/2013
E4 - Dose-Response Information to Support Drug Registration (I)	11/9/1994
E5 - Ethnic Factors in the Acceptability of Foreign Clinical Data (I)	6/10/1998
E6 - Good Clinical Practice: Consolidated Guideline (I)	5/9/1997
E7 - Studies in Support of Special Populations: Geriatrics (I)	8/2/1994
E7 Studies in Support of Special Populations; Geriatrics; Questions and Answers	2/21/2012
E8 - General Considerations for Clinical Trials (I)	12/24/1997
E9 - Statistical Principles for Clinical Trials (I)	9/16/1998
E10 - Choice of Control Group and Related Issues in Clinical Trials (I)	5/14/2001
E11 - Clinical Investigation of Medicinal Products in the Pediatric Population (I)	12/15/2000
E14 - Clinical Evaluation of QT/QTc Interval Prolongation and Proarrhythmic Potential for Non Antiarrhythmic Drugs (I)	10/20/2005
E14 Clinical Evaluation of QT/QTc Interval Prolongation and Proarrhythmic Potential for Non Antiarrhythmic Drugs. Q&As (I)	11/18/2008
E15 - Pharmacogenomics Definitions and Sample Coding (I)	4/8/2008

<b><u>ICH - Joint Safety/Efficacy (Multidisciplinary)</u></b>	<b><u>Issued Date</u></b>
Companion Document for M2: eCTD Specification Questions & Answers and Change Requests (I)	8/1/2006
M2 - Electronic Common Technical Document Specification (eCTD) (I)	4/2/2003
M3(R2) - Nonclinical Safety Studies for the Conduct of Human Clinical Trials and Marketing Authorization for Pharmaceuticals (I)	1/21/2010
M3(R2)Nonclinical Safety Studies for the Conduct of Human Clinical Trials and Marketing Authorization for Pharmaceuticals: Questions and Answers (R1)	2/25/2013
M4 - Common Technical Document for the Registration of Pharmaceuticals for Human Use - Granularity Annex (I)	10/17/2005
M4 - Organization of the Common Technical Document (CTD) (I)	10/16/2001
M4 - The CTD -- Efficacy Questions and Answers (Revised) (I)	12/22/2004
M4 - The CTD -- General Questions and Answers (Revised) (I)	12/22/2004
M4 - The CTD - Quality Questions and Answers/Location Issues (I)	6/9/2004
M4 - The CTD -- Safety Questions and Answers (I)	2/4/2003

<b><u>ICH - Quality</u></b>	<b><u>Issued Date</u></b>
ICH Q3C Maintenance Procedures for the Guidance for Industry Q3C Impurities: Residual Solvents - Final Recommendation for the Revision of the Permitted Daily Exposure for Cumene According to the Maintenance Procedures for Q3C Impurities: Residual Solvents	2/23/2012
Q1A(R2) - Stability Testing of New Drug Substances and Products (I)	11/21/2003
Q1B - Photostability Testing of New Drug Substances and Products (I)	5/16/1997
Q1C - Stability Testing for New Dosage Forms (I)	5/9/1997
Q1D - Bracketing and Matrixing Designs for Stability Testing of New Drug Substances and Products (I)	1/16/2003
Q1E - Evaluation of Stability Data (I)	6/8/2004
Q2A - Text on Validation of Analytical Procedures (I)	3/1/1995
Q2B - Validation of Analytical Procedures: Methodology (I)	5/9/1997
Q3A(R) - Impurities in New Drug Substances (I)	6/6/2008
Q3B(R) - Impurities in New Drug Products (I)	7/31/2006
Q3C - Impurities: Residual Solvents (I)	12/24/1997
Q3C Impurities: Residual Solvents: Maintenance Procedures for the Guidance for Industry Q3C (ICH Q3C Maintenance Procedures for the Guidance for Industry Q3C Impurities: Residual Solvent)	2/11/2002
Q3C Tables and List	2/22/2012

Q4B: Annex 1: Residue on Ignition/Sulphated Ash General Chapter (I)	2/21/2008
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions - Annex 8: Sterility Test General Chapter (I)	12/22/2009
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions; Annex 2 on Test for Extractable Volume of Parenteral Preparations General Chapter (I)	1/9/2009
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions; Annex 3 on Test for Particulate Contamination: Subvisible Particles General Chapter (I)	1/9/2009
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions; Annex 4A: Microbiological Examination of Non-Sterile Products: Microbial Enumeration Tests General Chapter (I)	4/8/2009
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions; Annex 4B: Microbiological Examination of Non-Sterile Products: Tests for Specified Micro-organisms General Chapter (I)	4/8/2009
Q4B Evaluation and Recommendation of Pharmacopoeial Texts; Annex 4C: Microbiological Examination of Non-Sterile Products: Acceptance Criteria for Pharmaceutical Preparations and Substances for Pharmaceutical Use General Chapter(I)	4/8/2009
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions- Annex 5: Disintegration Test General Chapter (I)	12/23/2009
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions Annex 7(R2) Dissolution Test General Chapter	6/23/2011
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions - Annex 8: Sterility Test General Chapter (I)	12/22/2009
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions- Annex 9: Tablet Friability General Chapter (I)	4/5/2010
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions; Annex 10: Polyacrylamide Gel Electrophoresis General Chapter (I)	4/12/2010
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the International Conference on Harmonisation Regions; Annex 11: Capillary Electrophoresis General Chapter (I)	9/3/2010
Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the International Conference on Harmonisation Regions; Annex 12 on Analytical Sieving General Chapter (I)	9/2/2010



Q4B: Evaluation and Recommendation of Pharmacopoeial Texts for Use in the International Conference on Harmonisation Regions (I)	2/21/2008
Q5A - Viral Safety Evaluation of Biotechnology Products Derived From Cell Lines of Human or Animal Origin (I)	9/24/1998
Q5B - Quality of Biotechnology Products: Analysis of the Expression Construct in Cells Used for Production of r-DNA Derived Protein Products (I)	2/23/1996
Q5C - Quality of Biotechnological Products: Stability Testing of Biotechnology/Biological Products (I)	7/10/1996
Q5D - Quality of Biotechnological/Biological Products: Derivation and Characterization of Cell Substrates Used for Production of Biotechnological/Biological Products (I)	9/21/1998
Q5E - Comparability of Biotechnological/Biological Products Subject to Changes in Their Manufacturing Process (I)	6/30/2005
Q6A - Specifications: Test Procedures and Acceptance Criteria for New Drug Substances and New Drug Products: Chemical Substances (I)	12/29/2000
Q6B - Test Procedures and Acceptance Criteria for Biotechnological/Biological Products (I)	8/18/1999
Q7A - Good Manufacturing Practice for Active Pharmaceutical Ingredients (I)	9/25/2001
Q8 (R2) - Pharmaceutical Development (I)	11/19/2009
Q8, Q9, and Q10 Questions and Answers (I)	11/1/2011
ICH Q8, Q9, & Q10 Questions and Answers -- Appendix: Q&As from Training Sessions (Q8, Q9, & Q10 Points to Consider)	7/25/2012
Q9 - Quality Risk Management (I)	6/2/2006
Q10 Pharmaceutical Quality System (I)	4/8/2009

<b><u>ICH - Safety</u></b>	<b><u>Issued Date</u></b>
S1A - The Need for Long-Term Rodent Carcinogenicity Studies of Pharmaceuticals (I)	3/1/1996
S1B - Testing for Carcinogenicity in Pharmaceuticals (I)	2/23/1998
S1C(R2) - Dose Selection for Carcinogenicity Studies of Pharmaceuticals	9/17/2008
S2A - Specific Aspects of Regulatory Genotoxicity Tests for Pharmaceuticals (I)	4/24/1996
S2B - Genotoxicity: Standard Battery Testing (I)	11/21/1997
S2(R1) Genotoxicity Testing and Data Interpretation for Pharmaceuticals Intended for Human Use	6/7/2012
S3A - Toxicokinetics: The Assessment of Systemic Exposure in Toxicity Studies (I)	3/1/1995
S3B - Pharmacokinetics: Repeated Dose Tissue Distribution Studies (I)	3/1/1995
S4A - Duration of Chronic Toxicity Testing in Animals (Rodent and Nonrodent Toxicity Testing) (I)	6/25/1999
S5A - Detection of Toxicity to Reproduction for Medicinal Products (I)	9/22/1994
S5B - Detection of Toxicity to Reproduction for Medicinal Products: Addendum on Toxicity to Male Fertility (I)	4/5/1996

S6(R1) Preclinical Safety Evaluation of Biotechnology-Derived Pharmaceuticals	5/18/2012
S7A - Safety Pharmacology Studies for Human Pharmaceuticals (I)	7/13/2001
S7B - Nonclinical Evaluation of the Potential for Delayed Ventricular Repolarization (QT Interval Prolongation) by Human Pharmaceuticals (I)	10/20/2005
S8 - Immunotoxicity Studies for Human Pharmaceuticals (I)	4/13/2006
S9 Nonclinical Evaluation for Anticancer Pharmaceuticals (I)	3/8/2010
S10 Photosafety Evaluation of Pharmaceuticals	2/4/2013

### **ICH Draft - Efficacy**

### **Issued Date**

E12A Principles for Clinical Evaluation of New Antihypertensive Drugs (I)	8/9/2000
E2B(R3) Electronic Transmission of Individual Case Safety Reports Implementation Guide — Data Elements and Message Specification; and Appendix to the Implementation Guide — Backwards and Forwards Compatibility	10/19/2011
E2C(R2) Periodic Benefit-Risk Evaluation Report	4/11/2012
E2D - Postapproval Safety Data Management: Definitions and Standards for Expedited Reporting (I)	9/15/2003

### **ICH Draft - Joint Safety/Efficacy (Multidisciplinary)**

### **Issued Date**

M5 - Data Elements and Standards for Drug Dictionaries (I)	9/1/2005
--	----------

Submitting Marketing Applications According to the ICH/CTD Format: General Considerations (I)	9/1/2001
---	----------

### **ICH Draft - Quality**

### **Issued Date**

Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions - Annex 6: Uniformity of Dosage Units General Chapter (I)	2/17/2009
---	-----------

Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions - Annex 13: Bulk Density and Tapped Density of Powders General Chapter (I)	7/14/2010
--	-----------

Q4B Evaluation and Recommendation of Pharmacopoeial Texts for Use in the ICH Regions - Annex 14: Bacterial Endotoxins Test General Chapter (I)	7/19/2010
---	-----------

### **INDs**

### **Issued Date**

Content and Format of INDs for Phase 1 Studies of Drugs Including Well-Characterized, Therapeutic, Biotechnology-Derived Products (I)	10/4/2000
--	-----------

### **Industry Letters**

### **Issued Date**

Continuation of a series of letters communicating interim and informal generic drug policy and guidance. Availability of Policy and Procedure Guides, and further operational changes to the generic drug review program (I)	3/2/1998
--	----------

Fifth of a series of letters providing informal notice about the Act, discussing the statutory mechanism by which ANDA applicants may make modifications in approved drugs where clinical data is required (I)	4/10/1987
--	-----------

Fourth of a series of letters providing informal notice to all affected parties about policy developments and interpretations regarding the Act. Three year exclusivity provisions of Title I (I)	10/31/1986
Implementation of the Drug Price Competition and Patent Term Restoration Act. Preliminary Guidance (I)	10/11/1984
Implementation Plan USP injection nomenclature (I)	10/2/1995
Seventh of a series of letters about the Act providing guidance on the "180-day exclusivity" provision of section 505(j)(4)(B)(iv) of the FD&C (I)	7/29/1988
Sixth of a series of informal notice letters about the Act discussing 3- and 5-year exclusivity provisions of sections 505(c)(3)(D) and 505(j)(4)(D) of the FD&C Act (I)	4/28/1988
Supplement to 10/11/84 letter about policies, procedures and implementation of the Act (Q & A format) (I)	11/16/1984
Third of a series of letters regarding the implementation of the Act (I)	5/1/1985
Year 2000 Letter from Dr. Janet Woodcock (I)	10/19/1998

<b><u>Labeling</u></b>	<b><u>Issued Date</u></b>
Adverse Reactions Section of Labeling for Human Prescription Drug and Biological Products; Content and Format (I)	1/24/2006
Clinical Studies Section of Labeling for Human Prescription Drug and Biological Products; Content and Format (I)	1/24/2006
Content and Format of the Dosage and Administration Section of Labeling for Human Prescription Drug and Biological Products (I)	3/23/2010
Content and Format for Geriatric Labeling (I)	10/5/2001

Contents of a Complete Submission for the Evaluation of Proprietary Names (I)	2/8/2010
Hypertension Indication: Drug Labeling for Cardiovascular Outcome Claims	3/15/2011
Labeling for Human Prescription Drug and Biological Products - Determining Established Pharmacologic Class for Use in the Highlights of Prescribing Information (I)	10/19/2009
Labeling for Human Prescription Drug and Biological Products - Implementing the PLR Content and Format Requirements	2/25/2013
Updating Labeling for Susceptibility Test Information in Systemic Antibacterial Drug Products and Antimicrobial Susceptibility Testing Devices (I)	7/2/2009
Warnings and Precautions, Contraindications, and Boxed Warning Sections of Labeling for Human Prescription Drug and Biological Products - Content and Format (I)	10/11/2011

### **Labeling Draft**

### **Issued Date**

Clinical Pharmacology Section of Labeling for Human Prescription Drug and Biological Products—Content and Format (I)	3/3/2009
Labeling for Combined Oral Contraceptives (I)	3/5/2004
Noncontraceptive Estrogen Drug Products for the Treatment of Vasomotor Symptoms and Vulvar and Vaginal Atrophy Symptoms — Recommended Prescribing Information for Health Care Providers and Patient Labeling (I)	11/16/2005
Pediatric Information Incorporated Into Human Prescription Drug and Biological Products Labeling Good Review Practice	2/28/2013
Public Availability of Labeling Changes in "Changes Being Effectuated" Supplements (I)	9/20/2006
Referencing Discontinued Labeling for Listed Drugs in Abbreviated New Drug Applications (I)	10/26/2000

<b><u>Modernization Act</u></b>	<b><u>Issued Date</u></b>
Changes to an Approved NDA or ANDA	4/2004
Classifying Resubmissions in Response to Action Letters	5/14/1998
Fast Track Drug Development Programs - Designation, Development, and Application Review & Appendix 2	7/22/2004
Formal Dispute Resolution: Appeals Above the Division Level	2/2000
Formal Meetings With Sponsors and Applicants for PDUFA Products	5/19/2009
Implementation of Section 120 of the Food and Drug Administration Modernization Act of 1997- Advisory Committees	10/1998
Implementation of Section 126 of the Food and Drug Administration Modernization Act of 1997 - Elimination of Certain Labeling Requirements	7/1998
Information Program on Clinical Trials for Serious or Life-Threatening Diseases and Conditions	3/2002
National Uniformity for Nonprescription Drugs - Ingredient Listing for OTC Drugs	4/1998
Providing Clinical Evidence of Effectiveness for Human Drug and Biological Products	5/14/1998
Qualifying for Pediatric Exclusivity Under Section 505A of the Federal Food, Drug, and Cosmetic Act	9/1999
Qualifying for Pediatric Exclusivity Under Section 505A of the Federal Food, Drug, and Cosmetic Act - 17 Frequently Asked Questions on Pediatric Exclusivity (505A), The Pediatric "Rule," and Their Interaction	7/27/1999

Repeal of Section 507 of the Federal Food, Drug and Cosmetic Act	5/1/1998
Reports on the Status of Postmarketing Study Commitments — Implementation of Section 130 of the Food and Drug Administration Modernization Act of 1997	2/15/2006
Standards for Prompt Review of Efficacy Supplements	5/15/1998
Submission of Abbreviated Reports and Synopses in Support of Marketing Applications	8/1998
Submitting and Reviewing Complete Responses to Clinical Holds	10/2000

### **Modernization Act Draft**

### **Issued Date**

Information Program on Clinical Trials for Serious or Life-Threatening Diseases and Conditions	1/2004
--	--------

### **OTC**

### **Issued Date**

Enforcement Policy on Marketing OTC Combination Products (CPG 71320.16) (I)	5/1/1984
General Guidelines for OTC Combination Products (I)	11/28/1978
Label Comprehension Studies for Nonprescription Drug Products (I)	8/3/2010
Labeling and Effectiveness Testing: Sunscreen Drug Products for Over-The-Counter Human Use — Small Entity Compliance Guide	12/6/2012



Labeling for Bronchodilators: Cold, Cough, Allergy, Bronchodilator, And Antiasthmatic Drug Products for Over-the-Counter Human Use (Small Entity Compliance Guide)	11/15/2012
Labeling of Nonprescription Human Drug Products Marketed Without an Approved Application as Required by the Dietary Supplement and Nonprescription Drug Consumer Protection Act: Questions and Answers	9/1/2009
Labeling OTC Human Drug Products -- Updating Labeling in ANDAs (I)	2/22/2001
Labeling Over-the-Counter Human Drug Products; Updating Labeling In Reference Listed Drugs and Abbreviated New Drug Applications (I)	10/18/2002
Labeling OTC Human Drug Products Using a Column Format (I)	12/19/2000
Labeling OTC Human Drug Products; Small Entity Compliance Guide (I)	5/13/2009
Labeling Over-the-Counter Human Drug Products; Questions and Answers	1/5/2009
Organ-Specific Warnings: Internal Analgesic, Antipyretic, and Antirheumatic Drug Products for Over-the-Counter Human Use -Small Entity Compliance Guide (I)	8/17/2010
Postmarketing Adverse Event Reporting for Nonprescription Human Drug Products Marketed without an Approved Application (I)	7/14/2009
Time and Extent Applications (I)	9/28/2011
Topical Acne Drug Products for Over-the-Counter Human Use--Revision of Labeling and Classification of Benzoyl Peroxide as Safe and Effective; Small Entity Compliance Guide	6/22/2011
Upgrading Category III Antiperspirants to Category I (43 FR 46728 - 46731) (I)	10/10/1978

**OTC Draft**

**Issued Date**

Labeling OTC Human Drug Products - Submitting Requests for Exemptions and Deferrals (I)	12/19/2000
Organ-Specific Warnings: Internal Analgesic, Antipyretic, and Antirheumatic Drug Products for Over-the-Counter Human Use — Labeling for Products That Contain Acetaminophen	7/5/2012
Self-Selection Studies for Nonprescription Drug Products	9/16/2011

### **Pharmacology/Toxicology**

### **Issued Date**

Carcinogenicity Study Protocol Submissions (I)	5/23/2002
Estimating the Maximum Safe Starting Dose in Initial Clinical Trials for Therapeutics in Adult Healthy Volunteers (I)	7/22/2005
Exploratory IND Studies (I)	1/17/2006
Format and Content of the Nonclinical Pharmacology/ Toxicology Section of an Application (I)	2/1/1987
Immunotoxicology Evaluation of Investigational New Drugs (I)	11/1/2002
Nonclinical Pharmacology/Toxicology Department of Topical Drugs Intended to Prevent the Transmission of Sexually Transmitted Diseases (STD) and/or the Development of Drugs Intended to Act as Vaginal Contraceptives (I)	10/16/1996
Nonclinical Evaluation of Late Radiation Toxicity of Therapeutic Radiopharmaceuticals	11/25/2011
Nonclinical Safety Evaluation of Drug or Biologic Combinations (I)	3/15/2006
Nonclinical Safety Evaluation of Pediatric Drug Products (I)	2/15/2006

Nonclinical Studies for the Safety Evaluation of Pharmaceutical Excipients	5/19/2005
Photosafety Testing (I)	5/7/2003
Recommended Approaches to Integration of Genetic Toxicology Study Results (I)	1/4/2006
Reference Guide for the Nonclinical Toxicity Studies of Antiviral Drugs Indicated for the Treatment of N/A Non-Life Threatening Disease: Evaluation of Drug Toxicity Prior to Phase I Clinical Studies (I)	2/1/1989
Reproductive and Developmental Toxicities -- Integrating Study Results to Assess Concerns	9/22/2011
Safety Testing of Drug Metabolites (I)	2/15/2008
Single Dose Acute Toxicity Testing for Pharmaceuticals - Revised (I)	8/26/1996

### **Pharmacology/Toxicology Draft**

### **Issued Date**

Animal Models--Essential Elements to Address Efficacy Under the Animal Rule (I)	1/21/2009
Genotoxic and Carcinogenic Impurities in Drug Substances and Products: Recommended Approaches (I)	12/16/2008
Nonclinical Safety Evaluation of Reformulated Drug Products and Products Intended for Administration by an Alternate Route (I)	3/7/2008
Statistical Aspects of the Design, Analysis, and Interpretation of Chronic Rodent Carcinogenicity Studies of Pharmaceuticals (I)	5/8/2001

<b><u>Procedural</u></b>	<b><u>Issued Date</u></b>
180-Day Generic Drug Exclusivity Under the Hatch-Waxman Amendments to the Federal Food, Drug, and Cosmetic Act (I)	7/14/1998
Centralized IRB Review Proceedings in Multicenter Clinical Trials	3/16/2006
Citizen Petitions and Petitions for Stay of Action Subject to Section 505(q) of the Federal Food, Drug, and Cosmetic Act	6/8/2011
Court Decisions, ANDA Approvals, and 180-Day Exclusivity Under the Hatch-Waxman Amendments to the Federal Food, Drug, and Cosmetic Act	3/27/2000
Disclosure of Materials Provided to Advisory Committees in Connection with Open Advisory Committee Meetings Convened by the Center for Drug Evaluation and Research Beginning on January 1, 2000 (I)	11/30/1999
Drug Products Containing Ensulizole, Hypromellose, Meradimate, Octinoxate, and Octisalate - Labeling Enforcement Policy (I)	6/3/2003
Emergency Use Authorization of Medical Products: Availability	7/26/2007
End-of-Phase 2A Meetings (I)	9/21/2009
Fast Track Drug Development Programs: Designation, Development, and Application Review (I)	11/18/1998
FDA Export Certificate (I)	7/12/2004
FDA Oversight of PET Drug Products -- Questions and Answers	12/4/2012
Fixed Dose Combinations and Co-Packaged Drug Products for Treatment of HIV (I)	10/18/2006
Formal Dispute Resolution: Appeals Above the Division Level	3/13/2013

Formal Meetings Between the FDA and Sponsors or Applicants (I)	5/14/2009
Good Review Management Principles for PDUFA Products (I)	3/31/2005
Guidance for Clinical Trial Sponsors Establishment and Operation of Clinical Trial Data Monitoring Committees Contains Nonbinding Recommendations	3/16/2006
Implementation of Section 120 of the Food and Drug Administration Modernization Act of 1997- Elimination of Certain Labeling Requirements (I)	11/2/1998
Implementation of Section 126 of the FDA Modernization Act of 1997 - Elimination of Certain Labeling Requirements, (I)	7/21/1998
Information Program on Clinical Trials for Serious or Life-Threatening Diseases and Conditions (I)	3/18/2002
Integrated Summaries of Effectiveness and Safety: Location Within the Common Technical Document (I)	4/21/2009
Levothyroxine Sodium Products - Enforcement of August 14, 2001, Compliance Date and Submission of New Applications (I)	7/13/2001
Medication Guides - Adding a Toll-Free Number for Reporting Adverse Events (I)	6/8/2009
National Uniformity for Nonprescription Drugs Ingredient Labeling for OTC Drugs (I)	4/9/1998
PET Drug Applications — Content and Format for NDAs and ANDAs; Fludeoxyglucose F 18 Injection; Ammonia N 13 Injection; Sodium Fluoride F 18 Injection	8/31/2011
PET Drug Applications - Content and Format for NDAs and ANDAs: Attachment I: Sample formats for chemistry, manufacturing, and controls (CMC) sections_2011	8/31/2011
Pharmacogenomic Data Submissions (I)	3/23/2005
Planning for the Effects of High Absenteeism to Ensure Availability of Medically Necessary Drug Products	3/15/2011

Potassium Iodide (KI) in Radiation Emergencies - Questions and Answers (I)	12/23/2002
Potassium Iodide as a Thyroid Blocking Agent in Radiation Emergencies (I)	12/10/2001
Potassium Iodide Tablets Shelf Life Extension for Federal Agencies and State and Local Governments (I)	3/8/2004
Qualifying for Pediatric Exclusivity Under Section 505A of the Federal Food, Drug, and Cosmetic Act - Revised (I)	10/1/1999
Refusal to File (I)	7/12/1993
Repeal of Section 507 of the Federal Food, Drug, and Cosmetic Act (I)	6/15/1998
Reports on the Status of Postmarketing Studies - Implementation of Section 130 of the Food and Drug Administration Modernization Act of 1997 (I)	2/16/2006
Special Protocol Assessment (I)	5/17/2002
Standards for the Prompt Review of Efficacy Supplements, Including Priority Efficacy Supplements (I)	5/15/1998
Submitting and Reviewing Complete Responses to Clinical Holds (Revised) (I)	10/26/2000
The Leveraging Handbook; an Agency Resource for Effective Collaborations - Guidance for FDA Staff (I)	6/19/2003
"Toll-Free Number Labeling and Related Requirements for Over-the-Counter and Prescription Drugs Marketed With Approved Applications" Small Entity Compliance Guide	6/15/2012
Useful Written Consumer Medication Information (CMI) (I)	7/18/2006
Women and Minorities Guidance Requirements	7/20/1998

<b><u>Procedural Draft</u></b>	<b><u>Issued Date</u></b>
Applications Covered by Section 505(b)(2) (I)	12/8/1999
Certification Process of Designated Medical Gases	12/18/2012
Dear Healthcare Provider Letters: Improving Communication of Important Safety Information	11/12/2010
Disclosing Information Provided to Advisory Committees in Connection with Open Advisory Committee Meetings Related to the Testing or Approval of New Drugs and Convened by CDER, Beginning January 1, 2000 (I)	12/22/1999
Disclosure of Conflicts of Interest for Special Government Employees Participating in FDA Product Specific Advisory Committees	2/14/2002
Electronic Source Data in Clinical Investigations	11/20/2012
Enforcement Policy -- OTC Sunscreen Drug Products Marketed Without an Approved Application	6/14/2011
Financial Disclosure by Clinical Investigators: Guidance for Clinical Investigators, Industry, and FDA Staff	5/24/2011
Forms for Registration of Producers of Drugs and Listing of Drugs in Commercial Distribution (I)	5/15/2001
How to Comply with the Pediatric Research Equity Act (I)	9/7/2005
Independent Consultants for Biotechnology Clinical Trial Protocols (I)	5/7/2003
Information Program on Clinical Trials for Serious or Life-Threatening Diseases and Conditions (I)	1/27/2004

Integrated Summary of Effectiveness (I)	8/28/2008
Notification to FDA of Issues that May Result in a Prescription Drug or Biological Product Shortage	2/27/2012
Oversight of Clinical Investigations — A Risk-Based Approach to Monitoring	8/29/2011
Pharmacogenomic Data Submissions -Companion Guidance (I)	8/29/2007
Postmarketing Safety Reporting for Human Drug and Biological Products Including Vaccines (I)	3/12/2001
Responding to Unsolicited Requests for Off-Label Information About Prescription Drugs and Medical Devices	12/27/2011
Submission of Patent Information for Certain Old Antibiotics (I)	12/3/2008
Submitting Debarment Certification Statements (I)	10/2/1998
Target Product Profile--A Strategic Development Process Tool (I)	3/30/2007
The Use of Clinical Holds Following Clinical Investigator Misconduct (I)	8/27/2002
Tropical Disease Priority Review Vouchers (I)	10/20/2008
Use of Histology in Biomarker Qualification Studies	12/29/2011

### **Small Entity Compliance Guides**

### **Issued Date**



Labeling and Effectiveness Testing: Sunscreen Drug Products for Over-The-Counter Human Use — Small Entity Compliance Guide	12/6/2012
Labeling for Bronchodilators: Cold, Cough, Allergy, Bronchodilator, And Antiasthmatic Drug Products for Over-the-Counter Human Use (Small Entity Compliance Guide)	11/15/2012
Labeling OTC Human Drug Products; Small Entity Compliance Guide	5/13/2009
Organ-Specific Warnings: Internal Analgesic, Antipyretic, and Antirheumatic Drug Products for Over-the-Counter Human Use — Small Entity Compliance Guide	8/17/2010
PET Drugs--Current Good Manufacturing Practice (CGMP); Small Entity Compliance Guide	8/4/2011
Safety Reporting Requirements for INDs and BA/BE Studies- Small Entity Compliance Guide	12/20/2012
Sterility Requirements for Aqueous-Based Drug Products for Oral Inhalation; Small Entity Compliance Guide	11/7/2001
"Toll-Free Number Labeling and Related Requirements for Over-the-Counter and Prescription Drugs Marketed With Approved Applications" Small Entity Compliance Guide	6/15/2012
Topical Acne Drug Products for Over-the-Counter Human Use--Revision of Labeling and Classification of Benzoyl Peroxide as Safe and Effective; Small Entity Compliance Guide	6/22/2011

<b><u>User Fee</u></b>	<b><u>Issued Date</u></b>
Applicability of User Fees to (1) Applications Withdrawn Before Filing, or (2) Applications the Agency Has Refused to File and That Are Resubmitted or Filed Over Protest (Attachment F)	7/12/1993
Application, Product, and Establishment Fees: Common Issues and Their Resolution (Revised) (Attachment D) (I)	12/16/1994
Classifying Resubmissions in Response to Action Letters (I)	5/14/1998

Fees-Exceed-the-Costs Waivers Under the Prescription Drug User Fee Act (I)	8/25/1999
Information Request and Discipline Review Letters Under the Prescription Drug User Fee Act (I)	11/21/2001
Submitting Separate Marketing Applications and Clinical Data for Purposes of Assessing User Fees (I)	1/3/2005
User Fee Waivers for Fixed Dose Combination Products and Co-Packaged Human Immunodeficiency Virus Drugs for the President's Emergency Plan for Acquired Immunodeficiency Syndrome Relief (I)	2/8/2007
User Fee Waivers, Reductions, and Refunds for Drug and Biological Products	9/26/2011