INTERNATIONAL CONFERENCE ON HARMONISATION OF TECHNICAL REQUIREMENTS FOR REGISTRATION OF PHARMACEUTICALS FOR HUMAN USE

ICH HARMONISED TRIPARTITE GUIDELINE

EVALUATION AND RECOMMENDATION OF PHARMACOPOEIAL TEXTS FOR USE IN THE ICH REGIONS ON RESIDUE ON IGNITION/SULPHATED ASH GENERAL CHAPTER Q4B ANNEX 1

Current *Step 4* version
dated 1 November 2007

This Guideline has been developed by the appropriate ICH Expert Working Group and has been subject to consultation by the regulatory parties, in accordance with the ICH Process. At Step 4 of the Process the final draft is recommended for adoption to the regulatory bodies of the European Union, Japan and USA.

Q4B Annex 1
Document History

Code *	History	Date
Q4B Annex 1	Approval by the Steering Committee under *Step 2* and release for public consultation.	8 June 2006

Current *Step 4* version

Q4B Annex 1	Approval by the Steering Committee under *Step 4* and recommendation for adoption to the three ICH regulatory bodies.	1 November 2007

* *Code as per the new codification system adopted by the ICH Steering Committee in November 2007*

EVALUATION AND RECOMMENDATION OF PHARMACOPOEIAL TEXTS FOR USE IN THE ICH REGIONS

ON

RESIDUE ON IGNITION/SULPHATED ASH GENERAL CHAPTER

ICH Harmonised Tripartite Guideline

Having reached *Step 4* of the ICH Process at the ICH Steering Committee meeting on 1 November 2007, this guideline is recommended for adoption to the three regulatory parties to ICH

TABLE OF CONTENTS

EVALUATION AND RECOMMENDATION OF PHARMACOPOEIAL TEXTS FOR USE IN THE ICH REGIONS ON RESIDUE ON IGNITION/SULPHATED ASH GENERAL CHAPTER

1. INTRODUCTION

This annex is the result of the Q4B process for Residue on Ignition/Sulphated Ash. The proposed texts were submitted by the Pharmacopoeial Discussion Group (PDG).

2. Q4B OUTCOME

2.1 Analytical Procedures

The ICH Steering Committee, based on the evaluation by the Q4B Expert Working Group (EWG), recommends that the official pharmacopoeial texts, Ph.Eur. 20414 Sulphated Ash, JP 2.44 Residue on Ignition Test, and USP <281> Residue on Ignition can be used as interchangeable in the ICH regions given the following:

2.1.1 Unless otherwise specified in a monograph, an appropriate sample weight is chosen, typically 1-2 g, to result in a level of residue sufficient to be accurately measurable by weight (typically 1 mg). If not specified in the monograph, the appropriate sample weight should be justified, and the sample weight and the acceptance criteria should be specified in the application dossier.

2.1.2 The muffle furnace should be appropriately calibrated to ensure compliance with regional GMP requirements.

2.2 Acceptance Criteria

The proposed texts evaluated did not contain acceptance criteria.

3. TIMING OF ANNEX IMPLEMENTATION

When this annex is implemented (incorporated into the regulatory process at ICH *Step 5*) in a region, it can be used in that region. Timing may differ for each region.

4. CONSIDERATIONS FOR IMPLEMENTATION

4.1 General consideration: When sponsors or manufacturers change their existing methods to the implemented Q4B-evaluated pharmacopoeial texts that are referenced in Section 2.1 of this annex, any change notification, variation, and/or prior approval procedures should be handled in accordance with established regional regulatory mechanisms pertaining to compendial changes.

4.2 FDA consideration: Based on the recommendation above, and in accordance with the conditions set forth in this annex, the pharmacopoeial texts referenced in Section 2.1 of this annex can be considered interchangeable. However, FDA might request that a company demonstrate that the chosen method is acceptable and suitable for a specific material or product, irrespective of the origin of the method.

4.3 EU consideration: For the European Union, the monographs of the Ph. Eur. have mandatory applicability. Regulatory authorities can accept the reference

in a marketing authorisation application, renewal or variation application citing the use of the corresponding text from another pharmacopoeia as referenced in Section 2.1, in accordance with the conditions set out in this annex, as fulfilling the requirements for compliance with the Ph. Eur. Chapter, Sulphated Ash: 20414, on the basis of the declaration of interchangeability made above.

4.4 MHLW consideration: The pharmacopoeial texts referenced in Section 2.1 of this annex can be used as interchangeable in accordance with the conditions set out in this annex. Details of implementation requirements will be provided in the notification by MHLW when this annex is implemented.

5. REFERENCES USED FOR THE Q4B EVALUATION

5.1 The PDG Stage 5B sign-off document: Japanese Pharmacopoeial Forum Volume 14, Number 4 (December 2005). (Note: the PDG cover letter published in this volume was subsequently changed based on Q4B comments.)

5.2 The pharmacopoeial references for Residue on Ignition/Sulphated Ash for this annex are:

5.2.1 European Pharmacopoeia (Ph. Eur.): Supplement 5.6 (official on January 2007) (reference Sulphated Ash 01/2007:20414);

5.2.2 Japanese Pharmacopoeia (JP): 2.44 Residue on Ignition Test as it appears in the JP Fifteenth Edition (March 31, 2006, The Ministry of Health, Labour and Welfare Ministerial Notification No. 285);

5.2.3 United States Pharmacopeia (USP): <281> Residue on Ignition official in USP 29, 2nd Supplement, August 2006.