

ICH HARMONISED TRIPARTITE GUIDELINE

**STABILITY TESTING OF
NEW DRUG SUBSTANCES AND PRODUCTS
Q1A(R2)**

Current *Step 4* version
dated 6 February 2003

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.

Q1A(R2)
Document History

First Codification	History	Date	New Codification November 2005
Q1	Approval by the Steering Committee under <i>Step 2</i> and release for public consultation.	16 September 1992	Q1
Q1A	Approval by the Steering Committee under <i>Step 4</i> and recommendation for adoption to the three ICH regulatory bodies. Q1 was renamed Q1A.	27 October 1993	Q1A
Q1A(R)	Approval by the Steering Committee of the first revision under <i>Step 2</i> and release for public consultation.	7 October 1999	Q1A(R1)
Q1A(R)	Approval by the Steering Committee of the first revision under <i>Step 4</i> and recommendation for adoption to the three ICH regulatory bodies.	8 November 2000	Q1A(R1)

Current *Step 4* version

Q1A(R2)	Approval by the Steering Committee of the second revision directly under <i>Step 4</i> without further public consultation, to include consequences of the adoption of Q1F (Stability Data Package for Registration Applications in Climatic Zones III and IV), and recommendation for adoption to the three ICH regulatory bodies.	6 February 2003	Q1A(R2)
---------	---	--------------------	---------

COVER NOTE FOR REVISION OF Q1A(R)
STABILITY TESTING OF
NEW DRUG SUBSTANCES AND PRODUCTS

The purpose of this note is to outline the changes made in Q1A(R) that result from adoption of ICH Q1F “Stability Data Package for Registration Applications in Climatic Zones III and IV”. These changes are:

1. The intermediate storage condition has been changed from $30^{\circ}\text{C} \pm 2^{\circ}\text{C}/60\% \text{RH} \pm 5\% \text{RH}$ to $30^{\circ}\text{C} \pm 2^{\circ}\text{C}/65\% \text{RH} \pm 5\% \text{RH}$ in the following sections:
 - 2.1.7.1 Drug Substance - Storage Conditions - General Case
 - 2.2.7.1 Drug Product - Storage Conditions - General Case
 - 2.2.7.3 Drug products packaged in semi-permeable containers
 - 3 Glossary - “Intermediate testing”

2. $30^{\circ}\text{C} \pm 2^{\circ}\text{C}/65\% \text{RH} \pm 5\% \text{RH}$ can be a suitable alternative long-term storage condition to $25^{\circ}\text{C} \pm 2^{\circ}\text{C}/60\% \text{RH} \pm 5\%$ in the following sections:
 - 2.1.7.1 Drug Substance - Storage Conditions - General Case
 - 2.2.7.1 Drug Product - Storage Conditions - General Case

3. $30^{\circ}\text{C} \pm 2^{\circ}\text{C}/35\% \text{RH} \pm 5\% \text{RH}$ has been added as a suitable alternative long-term storage condition to $25^{\circ}\text{C} \pm 2^{\circ}\text{C}/40\% \text{RH} \pm 5\%$ and the corresponding example for the ratio of water-loss rates has been included in the following section:
 - 2.2.7.3 Drug products packaged in semi-permeable containers

Mid-stream switch of the intermediate storage condition from $30^{\circ}\text{C} \pm 2^{\circ}\text{C}/60\% \text{RH} \pm 5\% \text{RH}$ to $30^{\circ}\text{C} \pm 2^{\circ}\text{C}/65\% \text{RH} \pm 5\% \text{RH}$ can be appropriate provided that the respective storage conditions and the date of the switch are clearly documented and stated in the registration application.

It is recommended that registration applications contain data from complete studies at the intermediate storage condition $30^{\circ}\text{C} \pm 2^{\circ}\text{C}/65\% \text{RH} \pm 5\% \text{RH}$, if applicable, by three years after the date of publication of this revised guideline in the respective ICH tripartite region.

STABILITY TESTING OF NEW DRUG SUBSTANCES AND PRODUCTS

ICH Harmonised Tripartite Guideline

First Recommended for Adoption at *Step 4* of the ICH Process on 27 October 1993.

Revised under *Step 2* of the ICH Process on 7 October 1999 and Recommended for Adoption at *Step 4* of the ICH Process on 8 November 2000.

This guideline has been Revised a second time and has reached *Step 4* of the ICH Process at the ICH Steering Committee meeting on 6 February 2003. It is recommended for adoption to the three regulatory parties to ICH

TABLE OF CONTENTS

1. INTRODUCTION	1
1.1. Objectives of the Guideline	1
1.2. Scope of the Guideline.....	1
1.3. General Principles.....	1
2. GUIDELINES	1
2.1. Drug Substance	1
2.1.1. General	1
2.1.2. Stress Testing	2
2.1.3. Selection of Batches	2
2.1.4. Container Closure System.....	2
2.1.5. Specification	2
2.1.6. Testing Frequency.....	3
2.1.7. Storage Conditions.....	3
2.1.8. Stability Commitment	5
2.1.9. Evaluation.....	5
2.1.10. Statements/Labeling.....	6

Explore Litigation Insights

Docket Alarm provides insights to develop a more informed litigation strategy and the peace of mind of knowing you're on top of things.

Real-Time Litigation Alerts



Keep your litigation team up-to-date with **real-time alerts** and advanced team management tools built for the enterprise, all while greatly reducing PACER spend.

Our comprehensive service means we can handle Federal, State, and Administrative courts across the country.

Advanced Docket Research



With over 230 million records, Docket Alarm's cloud-native docket research platform finds what other services can't. Coverage includes Federal, State, plus PTAB, TTAB, ITC and NLRB decisions, all in one place.

Identify arguments that have been successful in the past with full text, pinpoint searching. Link to case law cited within any court document via Fastcase.

Analytics At Your Fingertips



Learn what happened the last time a particular judge, opposing counsel or company faced cases similar to yours.

Advanced out-of-the-box PTAB and TTAB analytics are always at your fingertips.

API

Docket Alarm offers a powerful API (application programming interface) to developers that want to integrate case filings into their apps.

LAW FIRMS

Build custom dashboards for your attorneys and clients with live data direct from the court.

Automate many repetitive legal tasks like conflict checks, document management, and marketing.

FINANCIAL INSTITUTIONS

Litigation and bankruptcy checks for companies and debtors.

E-DISCOVERY AND LEGAL VENDORS

Sync your system to PACER to automate legal marketing.