



INTERNATIONAL PROGRAMME ON CHEMICAL SAFETY

**Harmonization Project Meeting to Update the IPCS
Document on Chemical-Specific Adjustment Factors**

Meeting Record

9-10 December 2004, Geneva, Switzerland

INTERNATIONAL PROGRAMME ON CHEMICAL SAFETY (IPCS)

**Report of the Harmonization Project Meeting to Update the Document on
Chemical-Specific Adjustment Factors**

9-10 December, Geneva, Switzerland¹

ITEM 1: WELCOME AND OPENING OF MEETING

1. The Harmonization Project Meeting to update the IPCS "Guidance Document for the Use of Data in Development of Chemical-Specific Adjustment Factors (CSAFs) for Interspecies Differences and Human Variability in Dose/Concentration-Response Assessment" was convened on 9-10 December 2004, in Geneva, Switzerland.
2. The meeting was opened by the IPCS Coordinator, Dr Tim Meredith. Dr Meredith welcomed participants (a list appears at Annex 1). He explained the purpose of the meeting was to consider the peer review comments received on the draft CSAF document and finalize it for publication. He noted that the recent Stocktake of the IPCS Harmonization Project found that the document was already in use in some countries/organizations (including Canada, the United States, JEFCA and JMPR) and had been identified for use in others (Australia, Japan, the United Kingdom and the EU). It was anticipated that finalization of the document will enable further uptake and use, however the meeting was also requested to provide advice on potential avenues to increase adoption and use of the document.

**ITEM 2: MEETING CHAIR AND ARRANGEMENTS FOR MEETING
REPORT**

3. The meeting elected Ms Bette Meek, Canada, as its Chair, and Dr Andrew Renwick, United Kingdom, as Rapporteur for the revised document (Agenda Item 5). It was agreed that the Secretariat, Ms Carolyn Vickers, would prepare a short meeting record addressing the other agenda items. Dr Meredith then turned the meeting over to the Chair.

ITEM 3: ADOPTION OF THE AGENDA

4. With the inclusion of a discussion on the issue of PBPK modelling under Item 7: Other Business, the agenda was adopted (refer to Annex 2).

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ITEM 4: INTRODUCTION TO THE DOCUMENT

5. The Chair provided an overview of the history of the document and the process by which peer review comments were received and considered. This was noted.

ITEM 5: FINALIZATION OF THE CSAF PUBLICATION

6. The meeting's revisions to the document were recorded in the revised document prepared by Andrew Renwick. Following editing, the document will be published.

ITEM 6: ADOPTION AND USE OF THE DOCUMENT

7. The meeting considered ways to facilitate adoption and use of the document by national and international risk assessment bodies. John Lipscomb reported on activities in the US to adopt formal guidance on CSAFs, which is expected to be based substantially on the IPCS guidance. Sharon Munn advised that when the document is finalized, steps could be taken to have it referenced in the EU Technical Guidance Document, thereby adopting it into EU risk assessment processes.
8. Suggestions to promote use included: publication of further journal articles or a special feature with an editorial; presentation at scientific conferences; preparation of a training module (e.g. for further education workshops or to include in University courses); and promoting the guidance to relevant regulatory and advisory committees and industry bodies. The journal articles and presentations could include experience based on promotion and use of the framework, as a way of facilitating further uptake. John Lipscomb advised he would present the guidance as part of a workshop at the 2005 Society of Toxicology meeting. Bette Meek offered assistance from Canada for a pilot project on the development of training materials for Harmonization Project products. A number of journals, conferences and organizations were specifically suggested as a starting point, for follow-up by the Secretariat in consultation with individual experts who offered advice/assistance.

ITEM 7: ANY OTHER BUSINESS

PBPK Modelling

9. The meeting was informed about discussions at the October 2004 meeting of the IPCS Harmonization Project Steering Committee (HSC) on possible new work to develop guidance on good practice in PBPK modelling. The HSC had noted that work on this topic was underway in several countries and had suggested discussions with key parties to confirm that an international activity led by IPCS would be supported, including discussing at this CSAF meeting.
10. The HSC Liaison for this work, Bette Meek, delivered a presentation providing additional background on the proposal, including the related initiatives by Canada and the US EPA. Meeting participants identified other bodies also considering the

issue, including the AEGLs Committee, EU COST (under the European Science Foundation), and possibly ECVAM. The meeting agreed that a way forward would be to convene a teleconference of potentially interested experts/partner organizations, leading to a small planning Working Group, which would prepare for an expert meeting process to develop the guidance. Sharon Munn and John Lipscomb agreed to consult internally in the EU and US, respectively, prior to the teleconference to gauge support. The timing for the work would need to be agreed among the interested parties and IPCS, and subject to resources being available.

ITEM 8: CLOSURE

11. In closing the meeting, the Chair and Secretariat thanked participants for their valuable contribution to the finalization of the document. On behalf of IPCS, Carolyn Vickers extended special thanks to Bette Meek and Andrew Renwick for their commitment to the development of the document since its inception and their assistance as Chair and Rapporteur for this December meeting.

**IPCS Harmonization Project Meeting to Update the Chemical-Specific
Adjustment Factors Document
Invited Participants List**

Dr Alan Boobis
Imperial College School of Medicine and Technology, United Kingdom

Dr Jeffrey Brent **
University of Colorado Health Sciences Center, USA

Dr Ursula Gundert-Remy
Federal Institute for Risk Assessment, Germany

Dr Akihiko Hirose
National Institute of Health Sciences, Japan

Dr John C. Lipscomb
US Environmental Protection Agency

Andrew Maier
Toxicology Excellence for Risk Assessment , USA

Ms Bette Meek, Health Canada

Dr Edward V. Ohanian**
US Environmental Protection Agency

Dr Andrew Renwick
School of Medicine, University of Southampton, United Kingdom

Dr Margareta Warholm
Karolinska Institutet, Sweden

Professor Ian Whyte**
Newcastle Mater Misericordiae Hospital, Australia

Representatives

Ms Sharon Munn
European Chemicals Bureau, Italy

Secretariat

Ms Carolyn Vickers
International Programme on Chemical Safety, Switzerland

** Invited, but unable to participate

WORLD HEALTH ORGANIZATION
INTERNATIONAL LABOUR ORGANIZATION
UNITED NATIONS ENVIRONMENT PROGRAMME

IPCS/CSAF/04.01
9-10 December 2004
Geneva

INTERNATIONAL PROGRAMME ON CHEMICAL SAFETY
Project on the Harmonization of Approaches to the Assessment of Risk
from Exposure to Chemicals

Meeting to Update the Document on Chemical-Specific Adjustment Factors
9-10 December 2004, WHO Headquarters, Geneva
Commencing at 9.00 am on 9 December in Room X.7

Annotated Agenda

1. Welcome and Opening of Meeting.

The meeting will be opened by the IPCS Coordinator, Dr Tim Meredith.

2. Meeting Chair and arrangements for meeting report.

The meeting will be invited to elect a Chair and Rapporteur.

3. Adoption of the Agenda (IPCS/CSAF/04.01)

The Chair will invite participants to adopt the agenda, amended as necessary.

4. Introduction to the Document

An brief explanation of the development of the document will be provided.

5. Finalization of the CSAF Publication.

Dr Andy Renwick will provide an overview of the comments received and how they were addressed in preparation of the revised draft document. The meeting will be asked to consider the revised draft, make any necessary amendments and finalize it for editing and publication.

6. Adoption and Use of the Document

The meeting will be asked to propose ways to facilitate adoption and use of the document by national and international risk assessment bodies.

7. Any other business.

- PBPK Modelling (proposal for Harmonization Project work).

8. Closure.