

Pesticide residues in food—1999

Toxicological evaluations

**Sponsored jointly by FAO and WHO
with the support of the International Programme
on Chemical Safety (IPCS)**

**Joint meeting of the
FAO Panel of Experts on Pesticide Residues
in Food and the Environment
and the
WHO Core Assessment Group**

Rome, 20–29 September 1999



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The preparatory work for the toxicological evaluations of pesticide residues carried out by the WHO Expert Group on Pesticide Residues for consideration by the FAO/WHO Joint Meeting on Pesticide Residues in Food and the Environment is actively supported by the International Programme on Chemical Safety (IPCS) within the framework of the Inter-Organization Programme for the Sound Management of Chemicals.

The International Programme on Chemical Safety (IPCS), established in 1980, is a joint venture of the United Nations Environment Programme (UNEP), the International Labour Organisation (ILO), and the World Health Organization (WHO). The overall objectives of the IPCS are to establish the scientific basis for assessing the risk to human health and the environment from exposure to chemicals, through international peer-review processes as a prerequisite for the promotion of chemical safety, and to provide technical assistance in strengthening national capacities for the sound management of chemicals.

The Inter-Organization Programme for the Sound Management of Chemicals (IOMC) was established in 1995 by UNEP, ILO, the Food and Agriculture Organization of the United Nations, WHO, the United Nations Industrial Development Organization, and the Organisation for Economic Co-operation and Development (Participating Organizations), following recommendations made by the 1992 United Nations Conference on the Environment and Development to strengthen cooperation and increase coordination in the field of chemical safety. The purpose of the IOMC is to promote coordination of the policies and activities pursued by the Participating Organizations, jointly or separately, to achieve the sound management of chemicals in relation to human health and the environment.

IPCS gratefully acknowledges the assistance of Mrs E. Heseltine, St Léon-sur-Vézère, France, for editing these monographs.

ISBN 92 4 1665 15 7

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* First full evaluation

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Abbreviations used

ADI	acceptable daily intake
AUC	area under the curve
BrdU	5-bromo-2'-deoxyuridine
bw	body weight
CI	confidence interval
CYP	cytochrome P450
DMBA	7,12-dimethylbenz[<i>a</i>]anthracene
DMSO	dimethyl sulfoxide
ED ₅₀	median effective dose
F	female
F ₀	parental generation
F ₁	first filial generation
F ₂	second filial generation
FIFRA	Federal Insecticide, Fungicide and Rodenticide Act (United States)
GABA	γ-aminobutyric acid
GLP	good laboratory practice
HPLC	high-performance liquid chromatography
IARC	International Agency for Research on Cancer
IC ₅₀	median inhibitory concentration
IPCS	International Programme on Chemical Safety
IU	international unit
JMPR	Joint FAO/WHO Meeting on Pesticide Residues
LC ₅₀	median lethal concentration
LD	limit of detection
LD ₅₀	median lethal dose
LOAEL	lowest-observed-adverse-effect level
M	male
MRL	maximum residue limit
NA	not analysed
NBHBA	<i>N</i> -nitrosobutyl- <i>N</i> -(4-hydroxybutyl)amine
NOAEL	no-observed-adverse-effect level
NOAEC	no-observed-effect concentration
NR	not reported
OECD	Organisation for Economic Co-operation and Development
PEG	polyethylene glycol
ppm	parts per million
QA	quality assurance
S9	9000 x g supernatant fraction of rodent liver
SD	Sprague-Dawley
SPF	specific pathogen-free
TBPC	1-[4-(1,1-dimethylethyl)phenoxy]-2-cyclohexanol
TCP	3,5,6-trichloro-2-pyridinol
TLC	thin-layer chromatography
TMRL	tolerable maximum residue limit
TPA	12- <i>O</i> -tetradecanoylphorbol 13-acetate
USEPA	US Environmental Protection Agency
w/v	weight per volume

Introduction

The toxicological monographs and monograph addenda contained in this volume were prepared by a WHO Core Assessment Group that met with the FAO Panel of Experts on Pesticide Residues in Food and the Environment in a Joint Meeting on Pesticide Residues (JMPR) in Rome, Italy, on 20–29 September 1999.

One of the substances evaluated by the Core Assessment Group at this Meeting, pyriproxyfen, was evaluated for the first time. The other 10 substances had been evaluated at earlier meetings. For four of these, chlormequat, glufosinate-ammonium, propylene thiourea, and pyrethrum extract, only information received since the previous evaluations is summarized, in 'monograph addenda'. The appropriate earlier documents should be consulted in order to obtain full toxicological profiles of these chemicals. Toxicological monographs were prepared on chlorpyrifos, dimethipin, ethoprophos, permethrin, 2-phenylphenol and its sodium salt, and propargite, summarizing new data and, where relevant, incorporating information from previous monographs and addenda. Reports and other documents resulting from previous Joint Meetings on Pesticide Residues are listed in Annex 1.

The report of the Joint Meeting has been published by the FAO as *FAO Plant Production and Protection Paper* 153. That report contains comments on the compounds considered, acceptable daily intakes established by the WHO Core Assessment Group, and maximum residue limits established by the FAO Panel of Experts. Monographs on residues prepared by the FAO Panel of Experts are published as a companion volume, as *Evaluations 1999, Part I, Residues*, in the FAO Plant Production and Protection Paper series.

The toxicological monographs and addenda contained in this volume are based on working papers that were prepared by temporary advisers before the 1999 Joint Meeting. A special acknowledgement is made to those advisers. The monographs were edited by Mrs E. Heseltine, St Léon-sur-Vézère, France.

The preparation and editing of this volume were made possible by the technical and financial contributions of the lead institutions of the International Programme on Chemical Safety (IPCS), which supports the activities of the JMPR. The designations employed and the presentation of the material in this publication do not imply the expression of any opinion whatsoever on the part of the Central Unit of the IPCS concerning the legal status of any country, territory, city or area or of its authorities, nor concerning the delimitation of its frontiers or boundaries. The mention of specific companies or of certain manufacturers' products does not imply that they are endorsed or recommended by the IPCS in preference to others of a similar nature that are not mentioned.

Any comments or new information on the biology or toxicology of the compounds included in this volume should be addressed to: Joint WHO Secretary of the Joint FAO/WHO Meeting on Pesticide Residues, International Programme on Chemical Safety, World Health Organization, Avenue Appia, 1211 Geneva 27, Switzerland.

**TOXICOLOGICAL MONOGRAPHS
AND MONOGRAPH ADDENDA**

