

Overview of Low-Level Radiation Exposure Assessment: Biodosimetry

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The capability to make diagnostic assessments of radiation exposure is needed to support triage of radiation casualties and medical treatment decisions in military operations. At the International Conference on Low-Level Radiation Injury and Medical Countermeasures session on biodosimetry in the military, participants reviewed the field of biodosimetry, covering a wide range of biological endpoints. Participants evaluated early changes associated with exposure to ionizing radiation, including chromosomal and DNA damage, gene expression, and associated proteins, and DNA mutations. The use and development of advanced monitoring and diagnostic technologies compatible with military operations was emphasized. Conventional radiation bioassays require a substantial amount of time between when the sample is taken and when the data can be provided for decision making. These "bench-top" bioassays are evaluated in laboratories that are not in the field; these laboratories routinely assess dose exposures of 25 cGy (photon equivalent levels). Detection thresholds can be reduced approximately fivefold by the addition of significant and time-consuming efforts. Alternative real-time bioassays that can be measured in field laboratories or with handheld detection devices show promise as screening and clinical diagnostic tools, but they require further development and validation studies.

Introduction

Biomarkers can be subdivided according to their applications.¹ Markers of exposure and dose can be used to measure and record acute, accidental or occupational exposure when limited or no physical measurements are available. Markers of risk or susceptibility can help predict which individuals are at greater risk for development of spontaneous disease. However, at present, no radiation-specific biomarkers of risk or susceptibility exist. Although markers of disease responses, the trace cellular or molecular changes that occur during development of disease, they may not reflect past radiation exposure.

The chromosome-aberration-based bioassay is the gold standard for radiation bioassays.² The lymphocyte dicentric assay (Fig. 1A) for radiation exposure assessment (see references cited in review articles³⁻⁵) has been the subject of intense scien-

tific interest since its introduction in 1962 by Bender.⁶ Lloyd and colleagues determined its threshold (>2 cGy X-ray) at low radiation doses on the basis of a multiple laboratory collaboration (the study involving the measurements of more than 250,000 metaphase spreads of cells).⁷ The commonly accepted threshold for typical applications of the lymphocyte-dicentric radiation bioassay is a 25 cGy photon-equivalent dose, which is based on the measurement of approximately 1,000 cells per sample.² Variations in the background dicentric levels in the general population demonstrate the effect of confounding factors on the threshold detection level of chromosome-aberration-based radiation bioassays. Figure 1B illustrates the signal (i.e., induction by ionization) to noise (i.e., background levels) characteristics of low radiation doses for the lymphocyte-dicentric bioassay in prior and chronic low-dose rate exposure scenarios. The occurrence of persistent chromosome aberrations (i.e., reciprocal translocations) using fluorescence *in situ* hybridization methodology is the recommended endpoint for radiation exposure assessment.⁸ Further threshold increases for radiation dose assessment by scoring persistent reciprocal translocation chromosome aberrations would be expected due to interindividual variability and age-dependent progressive increases (1×10^{-4} per 1,000 cells) in background reciprocal translocation levels.⁹

Military operational and medical needs require forward field assessment of individual radiation exposures. Figure 2 illustrates a conventional battle field triage system in terms of medical care and casualty evacuation.¹⁰ The application of conventional chromosome-aberration-based bioassays in forward deployable military operations is limited by the fact that, in a battle situation, a large number of metaphase spreads would need to be scored by experienced radiation-cytogenetic laboratorians. Instead, the military needs alternative dose and exposure bioassays that use technologies that are compatible with military operations. The bioassays must be sensitive enough to detect doses over appropriate dose ranges and, equally important, must provide rapid biomarker detection so that timely decisions can be made in the field. The biomarkers must be linked to both the radiation dose received and to the potential clinical outcome from the dose.

The conference session on military biodosimetry evaluated several classes of radiation-dose assessment biomarkers as alternatives to the conventional chromosome-aberration-based bioassay. The candidate classes included radiation-induced DNA damage, chromosome aberrations and DNA mutations detected in telophase cells, changes in gene expression, and fluid phase measurement of DNA mutations. The subject of DNA mutagenesis, measured by cytological and fluid phase mitochondrial DNA isolation assays, is only briefly covered due to space limitations. Use of advanced monitoring and diagnostic technol-

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Assessment Of Risk From Low Level Exposure To Radiation And Chemicals

Michael Rajnik



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Energy and Located at Los Alamos National Laboratory ,1999 Energy Research Abstracts ,1990 **Los Alamos National Laboratory Continued Operation Site-Wide** ,1999 **Risk assessment and risk management in regulatory decision-making** United States. Presidential/Congressional Commission on Risk Assessment and Risk Management,1997 **Current Catalog** National Library of Medicine (U.S.), First multi year cumulation covers six years 1965 70 **Risk Analysis and Governance in EU Policy Making and Regulation** Bernardo Delogu,2016-05-02 This book provides an easy but comprehensive and rigorous access to the main concepts terminology methods and procedures of risk analysis intended for all those involved in the EU policy and regulatory decision making on risks It establishes a common ground of knowledge which enables a more informed dialogue on risks a closer collaboration between decision makers and scientists and a better appraisal of the potential and limits of risk science The book also brings together in an accessible way much multidisciplinary knowledge which had been dispersed over many technical documents and specialist books The EU is in the front line of health safety and environmental risk management GMOs food safety hazardous chemicals climate change radiation hazards are just a few of the popular risk issues addressed by the EU through policy and regulatory measures The risk analysis paradigm including risk assessment management and communication has been at the core of the EU decision making for a long time already EU Institutions strive for a science based approach to risk management Nevertheless the dialogue and collaboration on risk issues between policy makers stakeholders and scientists are still difficult and the potential and limits of science in support of decision making as well as the basic concepts of risk analysis are not fully understood outside the narrow specialist circles **Environmental Health Perspectives** ,1994 **Final Report: Risk assessment and risk management in regulatory decision-making** United States. Presidential/Congressional Commission on Risk Assessment and Risk Management,1997 **Toxicology of industrial compounds** H. Thomas,1995-11-21 A large number of chemical compounds are constantly being introduced and produced both to assist and ease modern human life Among these chemicals industrial compounds represent a particular fraction of chemicals which are not intended for use in biological systems but to which humans may be accidentally exposed whether in the workplace by product application or via the environment Industrial chemicals as opposed to pharmaceuticals and agrochemicals are in many cases subjected only to a very basic examination of handling safety and may lack further toxicity testing This implies that essentially nothing is known about their bioavailability metabolism excretion and toxicological properties unless problems arise Covering the investigation of industrial chemicals this book focuses on their individual structure biological fate potential toxicity to mammals and the molecular mechanisms possibly underlying their adverse effects by highlighting the use and significance of experimental toxicology Special emphasis is placed on mechanistic aspects in the safety assessment of industrial compounds in addition to current regulatory and legal considerations *Hayes' Principles and Methods of Toxicology* A. Wallace Hayes,Claire L. Kruger,2014-10-10 Hayes Principles and Methods of Toxicology has long

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