Pollutants, Human Health and the Environment
The book is dedicated to Charlotte Rich, a PhD student and friend of many of the authors of the book, who died tragically in 2007 and Michael Gillott who died of cancer. Michael had the utmost respect for scientific research and the quest for truth. He persuaded us of the need to inform a wide audience, including health professionals, of the hazards and risks of pollutants.
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Over the past century a large number of man-made chemical substances have been widely dispersed into our environment, both by accident and design, raising concerns about their adverse effects on human health and the environment. There is no doubt that there has been a worrying increase in health problems that are related, at least in part, to these substances and their release from manufacturing processes, spills, inadequate handling, storage and use, and careless after-use disposal.

Over the past 30 years or so there has been a plethora of legislation at international and national levels aimed at controlling and regulating the production, use and disposal of chemicals. The introduction of the REACH (Registration, Evaluation, Authorisation and restriction of Chemicals) by the EU in 2007 has provided the innovative concept that the burden of proof is now on manufacturers to provide evidence of the safety of their products before supplying them: a practical example of the so-called precautionary principle. It also provides rules for the phasing out and substitution of the most dangerous substances already in circulation, though this, unfortunately, is likely to be a protracted process. The main objective of REACH is to improve risk management of industrial chemicals by banning their manufacture or importation into Europe.

One concern about the REACH legislation is that it will drive research and development and manufacturing involving chemicals to parts of the world where legislation is weak or non-existent. This would be highly unsatisfactory, since chemical pollution is a global issue. For example, the use of arsenical groundwater to grow crops in south-east Asia has resulted in health warnings on rice, especially warning against feeding it to babies in the USA and Europe; persistent organic pollutants such as pesticides and plasticisers, despite being used mainly at low latitudes, are accumulating in marine fish and mammals in polar and sub-polar regions; and the manufacture of pharmaceuticals in countries where environmental legislation fails to require the clean-up of waste water has the potential to increase the antibiotic resistance of pathogens.

This book provides a balanced view of the risks and benefits of several groups of substances: essential, toxic trace and radioactive elements; synthetic organic agricultural and industrial chemicals and pharmaceuticals; and particulates and nanomaterials. Most of these substances are important to modern industrialised societies but can have adverse impacts on the environment and human health. It also deals with risk reduction and the future role of chemicals in achieving sustainable development. The issue of sustainability in a world of finite resources is likely to become ever more important in considering the use of chemicals in the twenty-first century.

The book uses a risk-based approach for industrial and other chemicals. It includes a discussion of the potential use of chemicals in sustainable development and suggests that in the future there will be more emphasis on green chemistry and biomimicry, which involve learning from nature, with industry developing clean cycles of production that are built on natural processes whereby waste from one process is feed for another – the cradle-to-cradle concept.

The book includes well-researched material, with references to the latest published work. It is written in accessible English and provides an excellent introduction to anyone wishing to know more about the increasingly important subject of chemicals in the environment. The information it contains will be particularly useful to everyone affected by recent legislation including the REACH legislation.

The Earl of Selborne GBE FRS
Chair of the Council of Science and Technology
Global warming has captured the attention of the world’s media, public and politicians, but although the dramatically increasing carbon footprint we are leaving is important it is not the only effect of the twenty-first century on the environment. Many diverse pollutants have the potential to cause lasting damage to our environment. Processes that may make a quick buck today could cause untold difficulties for our successors who will inherit the challenge. We have already seen an inexorable rise in cancer incidence in the world. Although an increasingly aging population is the main driver, there is no doubt that other more subtle influences are at work. The incidence of cancer acts as a litmus test for the deleterious effects of the environment and lifestyle changes on our bodies.

It was announced this year that the lifetime risk of breast cancer in the UK has gone up from 1 in 9 women to 1 in 8 women, a rate comparable to that in the USA. Rates of this and many other types of cancer have risen dramatically since reliable cancer registries were first developed in the 1950s and they are projected to continue to do so in the future – according to the World Cancer Research Fund Report, cancer rates worldwide could increase by a further 50 per cent to 15 million new cases a year by the year 2020. Rates of many other chronic diseases, from Alzheimer’s disease to Parkinsonism, as well as mood disorders such as anxiety and depression are also increasing across the globe following industrialisation and development, at a time when many of the costs of health care are becoming unaffordable. It is clear that we must learn how to reduce the risks of such diseases and prevent the human and economic toll that they are taking on society worldwide.

Effective prevention requires a detailed understanding of the pathogenesis of disease and the dissection of the positive and negative drivers that influence the process. Cancer is a disease of cumulative somatic mutations leading to disruption in cellular growth control. It is not surprising that many pollutants can influence this process. By understanding the detailed factors involved in the aetiology of cancer it may be possible to devise public-health strategies to minimise the overall burden of disease. Furthermore, our increasing knowledge of the molecular mechanisms involved in the interplay of the environment and our genetic background may make it possible to personalise prevention strategies at some future point. This individualisation is far more likely to achieve wider compliance amongst the population rather than bland generic messages.

While it is widely acknowledged that the causes of many chronic conditions are multifactorial, attention is being directed increasingly to the role of chemicals in our diet and the wider environment – especially following the understanding of the important role of epigenetics in disease progression. For example, the role of endocrine-disrupting substances in hormone-dependent cancer, asbestiform particulates in mesothelioma and particulates generally in chronic lung disease has been established. It has even been suggested that chemicals found in certain plastics, as well as in cigarette smoke, may increase the risks of obesity or diabetes. It is clear that greater efforts should be made to reduce exposure and hence the risk to human health of potentially hazardous substances beyond those associated simply with smoking and alcohol consumption.

This book, Pollutants, Human Health and the Environment, equips health professionals with an up-to-date knowledge of hazardous substances to help them limit the risks to human health and prevent many chronic diseases. It explains clearly the difference between hazard and risk, and goes on to discuss groups of hazardous substances, chapter by chapter. It includes discussions of many controversial issues, including: toxic trace elements such as arsenic, cadmium and mercury; radiation and radioactive substances such as naturally occurring radon gas and other natural and artificial fission products; industrial chemicals such as benzene and trichloroethylene; pesticides and pharmaceuticals, which enter water supplies and the wider environment; and particulates, including asbestos. It also includes chapters on engineered nanomaterials, essential and beneficial trace elements such as selenium, copper and zinc, and natural oestrogens.

The book contains some striking information, for example: the numbers of people at risk of skin or bladder cancer from increased exposure to arsenic; the number of conditions for which there is evidence that selenium deficiency is a cause or a factor; the fact that the greatest exposure of US citizens to ionising radiation is from medical diagnostics and treatment and that this is 500 times the dose from the nuclear industry; the increased amount of oestrogen in our food because of changes in farming practices; our increased exposure to neurotoxic substances used as pesticides or preservatives; and the fact that mercury levels have increased by a factor of four over the last 100 years.

This book is highly recommended to all health professionals who wish to play an effective role in reducing the risk to human health of chemicals in the environment. For the sake of our children’s children we all need to understand the footprint we are leaving for them. The knowledge, understanding and information in this book are the key to developing effective action plans across the globe.

Karol Sikora

Professor of Cancer Medicine, Hammersmith Hospital and Dean, University of Buckingham Medical School
Tribute

Professor Stanley Bowie FRS, 1917 to 2008

Stanley Hay Umphray Bowie FRS, FRSE FEng, FIMM

Stanley Bowie was a scientist of international standing who, as Chief Geochemist, established and led the highly successful Geochemical Division of the British Geological Survey (BGS, formerly the Institute of Geological Sciences, IGS), which became a model for similar divisions in geological surveys throughout the world. He and his staff made major contributions in isotope geology, fluid-inclusion studies, trace-element geochemistry (including high-resolution geochemical mapping), ore mineralogy, economic geology and analytical chemistry. The first inductively coupled plasma mass spectrometer was developed by Alan Gray of the University of Surrey and Alan Date in the IGS with funding from the European Commission, negotiated by Stanley. Later he was involved in further instrument development, including the portable XRF analyser and the first towed seabed gamma spectrometer.

A Shetlander by birth, Stanley Bowie graduated in 1941 with a first-class honours degree in geology from the University of Aberdeen where he had also studied chemistry and physics. He was awarded the Mitchell prize for the best Honours Geology student and the Senior Kilgour Research Scholarship.

In January 1942, during the Second World War, he joined the Meteorological Branch of the Royal Air Force and was commissioned flying officer a year later. He was stationed with Bomber Command in East Anglia, which was later the base for the first American B17 squadron stationed in Britain.

In 1946 he joined the Geological Survey of Great Britain (GSGB) with the Special Investigations Unit (renamed the Atomic Energy Division, AED, in 1951). This was the Unit which had been responsible for advising the British Government on the availability of uranium supplies for the Manhattan Project during the war and subsequently provided geological information for the UK’s atomic-weapons and nuclear-energy programmes. It was Britain’s knowledge and ownership of uranium reserves that ensured that the country remained in the American-led nuclear club after 1945. Stanley worked on autoradiography studies of uranium and thorium minerals in thin and polished sections and, in collaboration with the Atomic Energy Research Establishment (AERE) at Harwell, began a programme of instrument development for uranium exploration that helped to develop Geiger-Müller counters for use in uranium exploration, borehole logging and aero-radiometric surveys. He also developed an index of radioactive minerals, which remained classified until 1976.

In 1955 Stanley was promoted to Chief Geologist of the AED and represented the UK at international conferences on atomic energy, helping to develop advanced radiometric instrumentation. He also developed, with Ken Taylor, a new system of opaque-mineral identification based on the measurement of indentation hardness and reflectance – a major advance over previous complex systems – which gave Britain an important lead in economic geology. He used the system to investigate and document uranium deposits throughout the free world, and it remained in use by most ore mineralogists until the advent of the electron microprobe.

In 1968 he was appointed Chief Geochemist, in charge of the analytical, mineralogy and isotope-geology units as well as the field geochemistry programmes. From 1968 to 1973 he led a uranium reconnaissance programme on behalf of the UKAEA using many of the instrumental methods developed earlier in his career as well as newer geochemical methods based on the delayed-neutron method of analysis. In 1970 he was appointed by NASA as a principal investigator for returned lunar samples.
His work with Peter Simpson on the ore mineralogy of these samples and with Clive Rice on the distribution of uranium using fission-track analysis made an important contribution to understanding the lunar surface.

In 1972 he obtained funding for a programme of systematic geochemical mapping of Great Britain. This programme developed, for the first time, quantitative reproducible methods for the preparation of geochemical maps of similar standing to gravity, magnetic and other geophysical maps prepared by geological surveys. Led by Dr Jane Plant, this programme became the model for geochemical databases worldwide, and many of the sampling, analytical, quality-control and quality-assurance techniques and the methods of data processing form the basis of recommendations of the IUGS/IAGC Task Group on ‘Global Geochemical Baselines’ initiated by Dr Arthur Darnley, a former colleague.

In 1975 he established and led a Royal Society Working Party on Environmental Geochemistry and Health, which included other notable scientists such as Professor John Webb of Imperial College, Dr Colin Mills of the Rowett Institute of Nutrition and Health and Dr Gerry Shaper of the Royal Free and University College Medical School. The Working Party was in contact with national coordinating committees in the USA and USSR, the Academies of Science of the five Scandinavian countries and individuals elsewhere. The proceedings of a Royal Society discussion meeting in 1977 entitled ‘Environmental Geochemistry and Health’ continue to be regarded as a key scene-setting volume covering geochemistry and the health of man, animals and plants. He collaborated in 1990 with Dr Cameron Bowie (not a relative) of the Somerset Health Authority in a book on radon and health and in a paper on the same topic, published in the Lancet in 1991.

In 1959 he was awarded the Silver Medal of the Royal Society of Arts. In 1963 he was elected a Fellow of the Royal Society of Edinburgh. He was elected a Fellow of the Royal Society in 1976 and in the same year he became president of the Institution of Mining and Metallurgy.

In 1984 a new platinum-group mineral was named bowieite by the United States Geological Survey in recognition of Stanley’s contribution to ore mineralogy. He was visiting Professor at Strathclyde University until 1985 and visiting Professor at Imperial College from 1985 until 1989, and he served on the Commission of Ore Mineralogy of the International Mineralogical Association until 1987.

This book is a tribute to Professor Stanley Bowie FRS, honouring him as one of the pioneers of geochemistry as applied in the real world, recognising especially his role in establishing high-quality geochemical mapping, researching radioactivity and radio-elements in the Earth’s crust, and applying these studies to the exploration and development of mineral resources and to the improvement of human health.
The Editors

Professor Jane Plant CBE, DSc, FRSM, FRSE, FRSA, FRGS, FIMMM, FGS, CEng, CGeol holds the Anglo American chair of Geochemistry at Imperial College London. She was formerly Chief Geochemist and later Chief Scientist of the British Geological Survey. She has been awarded seven honorary doctorates and many prizes and distinctions for her contribution to science, including the prestigious Lord Lloyd of Kilgerran Award of the Foundation of Science and Technology, the Coke Medal of the Geological Society and the Tetleman Fellowship of Yale University. She formerly chaired the Government’s Advisory Committee on Hazardous Substances and was a member of the Royal Commission on Environmental Pollution 2000–2006. She is presently on the Council of the UK All Party Parliamentary and Scientific Committee and the College of Medicine and is patron of several cancer charities, including the famous Penny Brohn Centre in Bristol.

She supervises and undertakes research in environmental geochemistry with particular reference to human health. She is an international expert on environmental pollution, specialising in understanding and modelling the sources and behaviour of essential, beneficial and toxic trace elements, radioelements and radioactivity in the environment. She established the world-renowned geochemical baseline programme of the UK (G-BASE), and subsequently co-led the global geochemical baseline International Union of Geological Sciences/International Association of GeoChemistry (IUGS/IAGC) Programme with Dr David Smith of the United States Geological Survey.

Professor Plant is the author of the internationally best-selling book *Your Life in Your Hands*, on overcoming breast cancer, and several other books on health including ones on osteoporosis and prostate cancer. Her latest popular health book, entitled *Beating Stress, Anxiety and Depression*, was published in 2009 and has been described as ground-breaking. These popular health books aim to empower sufferers by making available the latest scientific information on diet and lifestyle, as well as conventional medical treatments. She has played a leading role in developing this volume in order to help to communicate to others the significant health problems caused by chemicals in the environment.

Dr Nikolaos Voulvoulis is a Reader in Environmental Technology, leader of the Environmental Quality Research theme at the Centre for Environmental Policy and Director of the world-renowned MSc in Environmental Technology at Imperial College London. He supervises and undertakes research in the area of environmental analysis and assessment for environmental quality management. This focuses on the development of methods for assessing emerging environmental contaminants and their sources, pathways and fate in the environment, with emphasis on waste and waste-water-treatment processes. He is an international expert in environmental pollution by hazardous substances such as biocides, pesticides, endocrine-disrupting chemicals and pharmaceuticals, and on the associated policy and management issues. His research activities also involve the development and application of environmental-analysis tools, multi-criteria assessment, risk management and sustainability assessment. This research aims to develop methodologies that establish the influence of different parameters of environmental quality, process performance, and indicators of effects. His research has been having an impact on environmental decision-making and policy on environmental quality, climate change and human health nationally and internationally. Through surveys, environmental monitoring, modelling, laboratory experiments and lab-scale trials, he delivers high-quality research that has been published in some of the top journals in the field.

Dr Voulvoulis engages in a number of high-profile external teaching and research activities. Through such activities, he has developed strong links with industry, regulators, research organisations and NGOs. He is a member of the Steering Group of the Global Contaminated Land Network of the Chartered Institution of Water and Environmental Management and Director of the Opal Soil Centre responsible for the National Soil and Earthworm survey. This survey was recently included as an example of a science-based education programme and data-collection method in the European Atlas of Soil Biodiversity launched by the European Commission’s Joint Research Centre in September 2010 as part of the International Year of Biodiversity. In addition, he has recently been in charge of the evaluation of over 1000 environmental projects that were co-financed by the Instrument for Structural Policies
Pre-Accession or Cohesion Fund by the European Commission, assessing the effectiveness of these projects and their contribution to the acquis communautaire in the field of the environment – specifically in the fields of water quality and management and waste collection and treatment.

Professor Kristín Vala Ragnarsdóttir is the Dean of Engineering and Natural Sciences at the University of Iceland. She was a Professor of Environmental Geochemistry and Environmental Sustainability at the University of Bristol, UK until 2008. Educated in Geochemistry and Petrology at the University of Iceland, Reykjavík (BSc) and Geochemistry at Northwestern University, Evanston, Illinois (MS, PhD) she changed her focus a decade ago from Earth Sciences to cross-disciplinary Sustainability Science. Her research pertains to sustainability in its widest context, including nature protection, economics, society and the wellbeing of citizens. She is currently developing a framework for the establishment of sustainable communities. Vala is also working on soil-sustainability indicators for land management and undertaking a comparative study of the relative fertility of conventionally versus organically managed land to ensure future food security. Her activities also include the establishment of a framework for a sustainable financial system and natural-resource use, and she is investigating the factors involved in complex multi-factorial disease development. Previously she studied the behaviour of pollutants in the natural environment and the link between environment and health.

Professor Ragnarsdóttir was a member of the Scientific Advisory Board for Framework 7 Environment Programme from 2006 to 2008. She has been a member of grant research panels for the EC (Brussels), NERC (UK), NSF (USA) and ESA (Netherlands). Vala is a past Director of the Geochemical Society and was a member of the Board of the European Association for Geochemistry and the Geological Society of Great Britain. She was the chair of the Schumacher Society and is a current board member of the Balaton Group. Professor Ragnarsdottir is a past Associate Editor of *Geochimica Cosmochimica Acta, Chemical Geology* and *Geochemical Transactions*. She is a current Guest Editor of *Solutions*.
Contributors

E. Louise Ander
British Geological Survey, Kingsley Dunham Centre, Keyworth, Nottingham, NG12 5GG

Aldo R. Boccaccini
University of Erlangen-Nuremberg, Department of Materials Science and Engineering, 91058 Erlangen, Germany

Pamela Castle
Former Chair of the Environmental Law Foundation

Mark R. Cave
British Geological Survey, Kingsley Dunham Centre, Keyworth, Nottingham, NG12 5GG

Ho-Sik Chon
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ

Alexandra Collins
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ

Jason Dassyne
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ

Edward Derbyshire
Centre for Quaternary Research, Royal Holloway, University of London, Egham, Surrey, TW20 0EX

Danelle Dhaniram
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ

Mustafa B. A. Djamgoz
Department of Life Sciences, Imperial College London, Prince Consort Road, London SW7 2AZ

Sally Donovan
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ

Richard M. Evans
Centre for Toxicology, School of Pharmacy, University of London, 29–39 Brunswick Square, London

Claire J. Horwell
Institute of Hazard, Risk and Resilience, Department of Earth Sciences, Durham University, Science Laboratories, South Road, Durham, DH1 3LE

Timothy P. Jones
School of Earth and Ocean Sciences, Main Building, Cardiff University, Cardiff, Wales, CF10 3YE

Qin-Tao Liu
Current address: Dow Corning (China) Holding Co., Ltd.

Olwenn V. Martin
Institute for the Environment, Brunel University, Kingston Lane, Uxbridge, Middlesex, UB8 3PH

Rebecca McKinlay
Centre for Toxicology, University of London School of Pharmacy, 29–39 Brunswick Square, London

Superb K. Misra
Natural History Museum, Mineralogy, London SW7 5BD

Christopher J. Oates
Applied Geochemistry Solutions, 49 School Lane, Gerrards Cross, Buckinghamshire, SL9 9AZ

Dieudonné-Guy Ohandja
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ

Richard Owen
University of Exeter Business School, Streatham Court, Rennes Drive, Exeter, EX4 4PU; European Centre for Environment and Human Health, Peninsula College of Medicine and Dentistry, Royal Cornwall Hospital, Truro, TR1 3HD

Jilang Pan
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ

Xiyu Phoon
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ
CONTRIBUTORS

Jane A. Plant
Centre for Environmental Policy and Department of Earth Science and Engineering, Imperial College London, Prince Consort Road, London SW7 2AZ

K. Vala Ragnarsdottir
Faculty of Earth Sciences, School of Engineering and Natural Sciences, Askja, University of Iceland, Reykjavik 101, Iceland

Khareen Singh
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ

Barry Smith
Intelliscience Ltd, 38A Station Rd, Nottingham, NG4 3DB

Teresa D. Tetley
Section of Pharmacology and Toxicology, National Heart and Lung Institute, Imperial College London, London SW3 6LY

Andrew Thorley
National Heart and Lung Institute, Imperial College London, London SW3 6LY

James Treadgold
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ

Eugenia Valsami-Jones
Natural History Museum, Mineralogy, London SW7 5BD

Nikolaos Voulvoulis
Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ
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Introduction

Jane A. Plant¹*, Nikolaos Voulvoulis² and K. Vala Ragnarsdottir³

¹Centre for Environmental Policy and Department of Earth Science and Engineering, Imperial College London, Prince Consort Road, London SW7 2AZ
²Centre for Environmental Policy, Imperial College London, Prince Consort Road, London SW7 2AZ
³Faculty of Earth Sciences, School of Engineering and Natural Sciences, Askja, University of Iceland, Reykjavik 101, Iceland

Corresponding author, email jane.plant@imperial.ac.uk

This book is concerned with current and emerging issues associated with the impact of pollutants on human health and the environment. Public concern and the media are presently focused on just one pollutant – carbon dioxide – and this has distracted attention from the wider issue of chemicals in the environment generally. Carbon dioxide has become the centre of international political and media attention because of its high levels in the atmosphere, mainly as a result of the burning of fossil fuels, and its potential to cause global climate change and acidification of the oceans – changes that could not be corrected on human timescales. However, this is just one of the many significant changes in the chemistry of the Earth System caused by the burgeoning human population and the increasing demand for material goods. The changes made to the Earth by humans are now so great that many scientists propose that a new geological period known as the Anthropocene be added to the geological column (Crutzen and Stoermer, 2000). Some people date the beginning of this period at 8000 years ago, with the start of forest clearing and settled agriculture; others date it at 1784, the date of Stevenson’s steam engine, ‘The Rocket’. In this book we examine the impact that humanity has had on the chemistry of the Earth System during the Anthropocene, beginning from 8000 years ago (van Andel, 1995; Ruddiman, 2005).

There are many books dealing with aspects of environmental geochemistry: the carbon cycle, for example, or heavy metals such as lead or mercury, or the chemistry of particular environmental compartments such as soil or water. This book is different. It aims to give information on the many different groups of substances that are having an impact on the surface environment with the potential to cause harm to human health.

The contribution of chemicals to improvements in life expectancy, human health and material living standards for most people in Western-style democracies is widely acknowledged. Over the past 50 years, however, there has been mounting unease about the widespread use of synthetic chemicals and their risk to human health, including via environmental pathways (RCEP, 2003). There is also concern about the damaging effects of chemicals on the ecosystems upon which, ultimately, all life on Earth depends. Society’s fears about the risks of chemicals and other hazardous substances in the environment have been propagated by the media and, in the late twentieth century, by an information-technology revolution that has globalised communication (Gardner, 2008). Hazardous substances in the environment, from carbon dioxide, radioactivity and oil pollution to arsenic and polychlorinated biphenyls (PCBs), now occupy a central place in social, economic and political debates and developments worldwide (Beck, 1986) and will probably continue to do so in the future.

Concerns about chemicals in the environment date back at least to the early 1920s, when scientists and doctors such as Sir Robert McCarrison and Sir Albert Howard in the UK and Rudolf Steiner in Austria expressed concern about the change from traditional biologically based to chemical-based agriculture. One of the pioneers of organic farming in the UK was Lady Eve Balfour, and in 1946 she helped to found the Soil Association. The first half of the twentieth century had witnessed a growing number of early warnings of the hazards and risks of chemicals (EEA, 2001), but it was the publication in 1962 of Rachel Carson’s landmark book Silent Spring that is widely credited with launching the modern environmental movement (Glausiusz, 2007). The book catalysed public concern about synthetic chemicals in the environment and their unintended effects by documenting the damaging effects of pesticides, particularly on birds; it stated, for example, that DDT caused eggshell thinning, reproductive problems and death. Carson also accused the chemical industry of spreading disinformation, and public officials of accepting industry claims uncritically.

The concerns about the effects of pesticides raised by Carson have since been confirmed and extended to include a wide range of other health effects, from carcinogenicity, mutagenicity and reprotoxicity to endocrine disruption, genotoxicity, neurotoxicity...
and immunotoxicity. These effects have been observed in both wildlife and humans (Ecobichon and Joy, 1993; Colborn et al., 1997; Cadbury, 1998; Ecobichon, 2001; Stenersen, 2004). Carson also highlighted the broader ecological impacts of pesticides, which are increasingly recognised as a major factor affecting biodiversity and habitat loss. In the short term they can have toxic effects on directly exposed organisms, as well as causing changes in habitat and the food chain over the long term (Isenring, 2010).

In the US, Carson’s work led to the establishment of the Environmental Protection Agency in December 1970 – a model which has been followed by many individual countries as well as the European Union. Such agencies have developed formal processes of risk assessment and risk-management practices, as discussed throughout this book, as well as improved regulation.

We now know that there are many other chemicals of concern besides pesticides. In 1965, Clair Patterson, the scientist who in 1956 used mass spectrometry to establish that the Earth was 4.5 billion years old, published Contaminated and Natural Lead Environments of Man, which drew attention to the increased levels of lead from industrial sources in the environment and food chain (see Chapter 4). His work led to a ban on the use of tetraethyl lead (TEL) in gasoline (petroleum) in the US in 1986 and later on the use of lead in food cans. By the late 1990s, lead levels in the blood of Americans were reported to have dropped by up to 80 per cent (CDC, 1997; HHS, 1997). Lead in petrol and food cans is now banned in most of the developed world, but manufacturers of TEL initially shifted their sales to developing countries and leaded petrol was not banned in some African countries until 2006, 20 years later than in the US.

Concern about the chronic effects of toxic trace elements other than lead on the environment and human health has also mounted. For example, according to some authorities mercury levels in the biosphere have increased by up to four times since the 1880s (Monteiro and Furness, 1997). A high proportion of mercury emissions is from the burning of fossil fuels, especially in China and India. This contaminates the oceans, bioaccumulating up the food chain, so that foods such as fatty meat and dairy produce are contaminated. There is also increasing evidence of widespread chemical contamination of human tissues with synthetic chemicals, in some cases to levels higher than those known to cause adverse health effects in other species, even in remote populations of the globe (such as the Inuit of Northern Canada and Greenland and inhabitants of remote islands such as the Faroes) (Walsh et al., 2008). Chemical pollution is a global issue, crossing national boundaries.

Problems caused by chemicals can arise quite unexpectedly, despite the development of extensive toxicity testing and risk-assessment procedures based on sophisticated science and technology. Many such problems – such as the ozone-depleting action of the chlorofluorocarbons (CFCs) and the endocrine-disrupting effects of the birth-control pill released to the aquatic environment – had not been anticipated (RCEP, 2003; Jobling and Owen, 2010; EEA, 2001). The often unpredictable and uncertain nature of innovation and technology and the wider impacts of new developments (Collingridge, 1980; RCEP, 2008) are also demonstrated by the identification in 1992 of the transgenerational endocrine-disrupting properties of common chemicals, including some plasticisers, detergents, pesticide formulations and pharmaceuticals. These studies, led by the distinguished biologist Theo Colborn (TEDX, 2010), showed that such substances were being transferred from top predator females to their offspring, resulting in chemically induced alterations in sexual and functional development. The problems resulting from exposure to low doses and/or low ambient levels of endocrine-disrupting chemicals have been described in Our Stolen Future (Colborn et al., 1997) and The Feminization of
Nature (Cadbury, 1996). The risks and benefits of natural oestrogens are considered in Chapter 9.

In addition to the chronic effects of chemicals, there have also been several high-profile chemical accidents such as those at Flixborough in the UK in 1974, Seveso in Italy in 1976, Bhopal in India in 1984, Toulouse in France in 2001, and at the Hertfordshire Oil Storage Terminal at Buncesfield in the UK in 2005.

Concerns about pollution are not restricted to chemicals. Fears of radioactivity (see Chapter 5) date back to the bombing of Hiroshima and Nagasaki in 1945 at the end of the Second World War. Radiation has been and continues to be an emotive issue (Fisk, 1996), and psychometric studies of public perception of risk have shown that the dangers associated with radioactivity are the most dreaded and least understood hazards (Slovic, 1987).

There have been several serious nuclear accidents in the former Soviet Union, which have resulted in major explosions and/or the release of radioactive materials, with many deaths and hundreds of thousands of people evacuated. The most recent accident, at the uncontained civil nuclear reactor at Chernobyl in the Ukraine in 1986, caused severe local contamination as well as regional contamination, which continues to affect livestock farming in parts of Europe. There is concern about further explosions at the site. Outside the countries of the former Warsaw Pact and Soviet Union, however, there had been only two partial meltdowns in civil nuclear power stations until the Fukushima incident in March 2011 (see Chapter 5). The first, in the Fermi Reactor in Michigan in 1966 required the reactor to be repaired; the other, the Three Mile Island accident in Pennsylvania, led to the permanent shutdown of the reactor. The high level of safety of Western reactors, however, meant there were no deaths or serious injuries.

The Three Mile Island accident ended nuclear power development in the US for more than 30 years. The irrationality of our approach to risk (Gardner, 2008) is perhaps best illustrated by our concerns about radioactivity. For example, the average US citizen now receives approximately 50 per cent of their dose of ionising radiation from medical diagnostics compared to 0.1 per cent from the nuclear industry, and yet public concern continues to focus on nuclear energy and nuclear waste repositories.

There have been many environmentally damaging accidental oil spills, some of which were associated with considerable loss of human life. These include the Torrey Canyon disaster in 1967, the Amoco Cadiz spill in 1978 and, most recently, the BP deep-drilling rig accident in the Gulf of Mexico in 2010. In July 1988, the Piper Alpha North Sea production platform exploded, and the resulting fire destroyed the platform and killed 167 men. It has recently been revealed that more than 5 tonnes of highly toxic carcinogenic PCBs were lost in the disaster, and other oil spills have resulted in the release of toxic chemicals. These accidents highlight the dangers to the Earth’s fragile ecosystems of servicing our heavily energy- and resource-dependent societies.

As well as chemicals and radiation, environmental exposure to many other substances, including particulates (see Chapter 10), is now known to cause health problems. For example, asbestiform minerals used in the construction industries are associated with lung disease including mesothelioma, a serious type of cancer. More than 3000 people a year now die in the UK as a result of inhaling asbestos dust, and one US government estimate suggests it may eventually kill 5.4 million Americans (Tang et al., 2010). Silica dust has long been known to cause silicosis and other lung diseases in miners and quarry workers. It has also been suggested that airborne particulate matter may cause between 100 000 and 300 000 deaths in Europe each year (Tainio et al., 2007). These findings have given rise to concerns about the potential effects of manufactured or engineered nanoparticles such as carbon nanotubes (see Chapter 11), one of the most important groups of emerging contaminants (Royal Society and Royal Academy of Engineering, 2004).

Looking back over the past two and a half centuries since the industrial revolution, and especially since the second world war, there has been a change in the developed world from the industrial societies of the ‘dark satanic mills’ and smogs where risks of pollution were more localised and immediately perceptible, to a globalised, risk-conscious and even risk-averse society which is no longer simply concerned with ‘making nature useful but with problems resulting from techno-economic development itself’ (Beck, 1986). Such risks are now known to be more complex and to have global impacts, although they are often hidden and undetectable through the senses (Gardner, 2008).

Many of the risks may remain poorly understood, or unknown.

Over the past 50 years, the rapid development of disciplines such as toxicology, ecotoxicology and environmental chemistry have played an important role in improving our understanding of the hazards, sources, behaviour, fate and health effects of pollutants in the environment. Environmental data in these and related disciplines, including geochemical databases, on regional, national and international scales, are increasingly available in the scientific literature and beyond. These developments have been accompanied by great improvements in analytical methods, such as inductively coupled plasma (ICP) mass spectrometry for inorganic elements and isotopes, and gas chromatography for separating chemicals in complex environmental matrices with low detection limits. New disciplines have also emerged, including genomics, proteomics, metabolomics and now epigenetics, which help to explain the biological effects of hazardous substances. These fundamental scientific developments have been matched, notably through organisations such as the Organisation for Economic Co-operation and Development, by the development of standardised tests for the hazardous effects of chemicals and effluents entering the environment. Many of these issues and developments are discussed in the chapters which follow.

The book is based on a risk-analysis framework, using the source–pathway–target model of organisations such as the US Environmental Protection Agency and the European Environment Agency. This model is explained in Chapters 1 and 2. Each of the remaining chapters follows a similar format, to ensure comprehensiveness of cover and easy cross reference. The introduction to each chapter describes the relevance and discovery of the health and environmental impacts of the substances to be discussed. This is followed by sections on the known hazardous properties; the principal natural and artificial sources; the main environmental pathways and exposure routes; the effects on
receptors, especially humans, with particular reference to health; and methods of risk reduction. The final chapter of the book proposes ways in which human impact on the chemistry of the Earth System can be reduced. The book as a whole provides an integrated overview of the impact of chemicals in the environment, but each of the chapters can stand alone as a complete work in its own right. Each chapter has been peer reviewed.

The book is intended for students of the environment, especially environmental geochemistry, engineering and medical geography, and also for health professionals, for whom it provides essential information for epidemiological studies and public health. One of our main aims is to engage the medical profession in the increasingly important issue of the chemical disturbance of the environment by humanity, threatening not only our own health but also that of future generations and the Earth itself.

References


1

The scientific appraisal of hazardous substances in the environment

Olwenn V. Martin¹* and Jane A. Plant²

¹Institute for the Environment, Brunel University, Kingston Lane, Uxbridge, Middlesex, UB8 3PH
²Centre for Environmental Policy and Department of Earth Science and Engineering, Imperial College London, Prince Consort Road, London SW7 2AZ
*Corresponding author, email olwenn.martin@brunel.ac.uk

1.1 Introduction

This book describes a wide range of non-living toxins that are present in the environment and are potentially harmful to human health or the environment. It covers both organic and inorganic substances, natural as well as manufactured substances. It deals mostly with substances that are hazardous because of their chemical properties, but also includes some where the hazard derives from their physical properties, for example, particulates and nanoparticles. This chapter summarises some important concepts of basic toxicology and environmental epidemiology relevant to an understanding of the possible effects of pollutants in the environment.

A common misconception is that chemicals made by nature are intrinsically good and, conversely, those manufactured by man are bad (Ottoboni, 1991). However, there are many examples of toxic compounds produced by algae or other micro-organisms, venomous animals and plants. There are even examples of environmental harm resulting from the presence of relatively benign natural compounds, either in unexpected places or in unexpected quantities. It is therefore of prime importance to define what is meant by ‘chemical’ when referring to chemical hazards in this chapter and the rest of this book. The correct term for a chemical compound to which an organism may be exposed, whether of natural or synthetic origins, is xenobiotic, i.e. a substance foreign to an organism (the term has also been used for transplants). A xenobiotic can be defined as a chemical which is found in an organism but which is not normally produced or expected to be present in it. It can also cover substances that are present in much higher concentrations than are usual.

1.2 Fundamental concepts of toxicology

Toxicology is the science of poisons. A poison is commonly defined as ‘any substance that can cause an adverse effect as a result of a physicochemical interaction with living tissue’ (Duffus, 2006). The use of poisons is as old as the human race, as a method of hunting or warfare as well as murder, suicide or execution. The evolution of this scientific discipline cannot be separated from the evolution of pharmacology, or the science of cures. Theophrastus Philippius Aureolus Bombastus von Hohenheim, more commonly known as Paracelsus (1493–1541), a physician contemporary of Copernicus, Martin Luther and da Vinci, is widely considered as the father of toxicology. He challenged the ancient concepts of medicine based on the balance of the four humours (blood, phlegm, yellow and black bile) associated with the four elements and believed that illness occurred when an organ failed and poisons accumulated. This use of chemistry and chemical analogies was particularly offensive to the contemporary medical establishment. He is famously credited with the quotation that still underlies present-day toxicology.
In other words, all substances are potential poisons, since all can cause injury or death following excessive exposure. Conversely, this statement implies that all chemicals can be used safely if handled with appropriate precautions and exposure is kept below a defined limit below which risk is considered tolerable (Duffus, 2006). The concepts of tolerable risk and adverse effect illustrate the value judgements embedded in an otherwise scientific discipline relying on observable, measurable empirical evidence. What is considered abnormal or undesirable is dictated by society rather than science. Any change from the normal state is not necessarily an adverse effect even if statistically significant. An effect may be considered harmful if it causes damage, irreversible change or increased susceptibility to other stresses, including infectious disease. The stage of development or state of health of the organism may also have an influence on the degree of harm.

### 1.2.1 Routes of exposure

Toxicity will vary depending on the route of exposure. There are three routes by which exposure to environmental contaminants may occur:

- **Ingestion.**
- **Inhalation.**
- **Skin adsorption.**

In addition, direct injection may be used in testing for toxicity. Toxic and pharmaceutical agents generally produce the most rapid response and greatest effect when given intravenously, directly into the bloodstream. A descending order of effectiveness for environmental exposure routes would be inhalation, ingestion and skin adsorption.

Oral toxicity is most relevant for substances that might be ingested with food or drinks. It could be argued that this is generally under an individual’s control, but people often don’t know what chemicals there are in their food or water and are not well informed about the current state of knowledge about their harmful effects.

Inhalation of gases, vapours, dusts and other airborne particles is generally involuntarily (with the notable exception of smoking). The destination of inhaled solid particles depends upon their size and shape. In general, the smaller the particle, the further into the respiratory tract it can go. A large proportion of airborne particles breathed through the mouth or cleared by the cilia of the lungs can enter the gut.

Dermal exposure generally requires direct and prolonged contact with the skin. The skin acts as a very effective barrier against many external toxicants, but because of its large surface area (1.5–2 m²) some of the many and diverse substances it comes in contact with may elicit topical or systemic effects (Williams and Roberts, 2000). If dermal exposure is often most relevant in occupational settings, it may nonetheless be pertinent in relation to bathing waters (ingestion is also an important route of exposure in this context). The use of cosmetics raises the same questions regarding the adequate communication of current knowledge about potential effects as those related to food.

### 1.2.2 Duration of exposure

The toxic response will also depend on the duration and frequency of exposure. The effect of a single dose of a chemical may be severe whilst the same total dose given at several intervals may have little or no effect: the effect of drinking four beers in one evening, for example, is very different from that of drinking four beers in four days. Exposure duration is generally divided into four broad categories: acute, sub-acute, sub-chronic and chronic. Acute exposure to a chemical usually refers to a single exposure event or repeated exposures over a duration of less than 24 hours. Sub-acute exposure to a chemical refers to repeated exposures for 1 month or less, sub-chronic exposure to continuous or repeated exposures for 1 to 3 months or approximately 10 per cent of the lifetime of an experimental species, and chronic exposure to continuous or repeated exposures for more than 3 months, usually 6 months to 2 years in rodents (Eaton and Klaassen, 2001). Chronic exposure studies are designed to assess the cumulative toxicity of chemicals with potential lifetime exposure in humans. The same terms are used in real-life situations, though it is generally very difficult to ascertain with any certainty the frequency and duration of exposure.

For acute effects, the time component of the dose is not important, as it is the high dose that is responsible for the effects. However, the fact that acute exposure to agents that are rapidly absorbed is likely to induce immediate toxic effects does not rule out the possibility of delayed effects, and these are not necessarily similar to those associated with chronic exposure (e.g. latency between the onset of certain cancers and exposure to a carcinogenic substance). The effect of exposure to a toxic agent may depend on the timing of exposure. In other words, long-term effects as a result of exposure to a toxic agent during a critically sensitive stage of development may differ markedly from those seen if an adult organism is exposed to the same substance. Acute effects are almost always the result of accidents, or, less commonly, criminal poisoning or self-poisoning (suicide). Chronic exposure to a toxic agent is generally associated with long-term low-level chronic effects, but this does not preclude the possibility of some immediate (acute) effects after each administration. These concepts are closely related to the mechanisms of metabolic degradation and excretion of ingested substances, as illustrated in Figure 1.1.

### 1.2.3 Mechanisms of toxicity

The interaction of a foreign compound with a biological system is two-fold: there is the effect of the organism on the compound...
Toxicokinetics relate to the delivery of the compound to its site of action, including absorption (transfer from the site of administration into the general circulation), distribution (via the general circulation into and out of the tissues), and elimination (from general circulation by metabolism or excretion). The target tissue refers to the tissue where a toxicant exerts its effect, and is not necessarily where the concentration of the toxic substance is highest. Many halogenated compounds such as polychlorinated biphenyls (PCBs) or flame retardants such as polybrominated diphenyl ethers (PBDEs) are known to bioaccumulate in body fat. Whether such sequestration processes are actually protective to the individual organisms (by lowering the concentration of the toxicant at the site of action) is not clear (O’Flaherty, 2000). In an ecological context however, such bioaccumulation may serve as an indirect route of exposure for organisms at higher trophic levels, thereby potentially contributing to biomagnification through the food chain.

Absorption of any compound that has not been intravenously injected will entail transfer across membrane barriers before it reaches the systemic circulation, and the efficiency of absorption processes is highly dependent on the route of exposure.

It is also important to note that distribution and elimination, although often considered separately, take place simultaneously. Elimination itself comprises two kinds of processes, excretion and biotransformation, which also take place simultaneously. Elimination and distribution are not independent of each other, as effective elimination of a compound will prevent its distribution in peripheral tissues, whilst, conversely, wide distribution of a compound will impede its excretion (O’Flaherty, 2000). Kinetic models attempt to predict the concentration of a toxicant at the target site from the administered dose. The ultimate toxicant, i.e. the chemical species that induces structural or functional alterations resulting in toxicity, may be the compound administered (parent compound), but it can also be a metabolite of the parent compound generated by biotransformation processes, i.e. toxication rather than detoxication (Timbrell, 2000; Gregus and Klaassen, 2001). The liver and kidneys are the most important excretory organs for non-volatile substances, whilst the lungs excrete volatile compounds and gases. Other routes of excretion include the skin, hair, sweat, nails and milk. Milk may be a major route of excretion for lipophilic chemicals due to its high fat content (O’Flaherty, 2000).

Toxicodynamics is the study of toxic response at the site of action, including the reactions with and binding to cell constituents, and the biochemical and physiological consequences of these actions. Such consequences may therefore be manifested and observed at the molecular or cellular levels, at the target organ or on the whole organism. Therefore, although toxic responses have a biochemical basis, the study of toxic response is generally subdivided, either depending on the organ on which toxicity is observed, including hepatotoxicity (liver), nephrotoxicity (kidney), neurotoxicity (nervous system), pulmonotoxicity (lung) or depending on the type of toxic response, including teratogenicity (abnormalities of physiological development), immunotoxicity (immune system impairment), mutagenicity (damage of genetic material), carcinogenicity (cancer causation).
or promotion). The choice of the toxicity endpoint to observe in experimental toxicity testing is therefore of critical importance. In recent years, rapid advances of biochemical sciences and technology have resulted in the development of bioassay techniques that can contribute invaluable information regarding toxicity mechanisms at the cellular and molecular level. However, the extrapolation of such information to predict effects in an intact organism for the purpose of risk assessment is still in its infancy (Gundert-Remy et al., 2005).

1.2.4 Dose–response relationships

The theory of dose–response relationships is based on the assumptions that (1) the activity of a substance depends on the dose an organism is exposed to (i.e. all substances are inactive below a certain threshold and active over that threshold), and (2) dose–response relationships are monotonic (i.e. the response rises with the dose). Toxicity may be detected either as an all-or-nothing phenomenon such as the death of the organism or as a graded response such as the hypertrophy of a specific organ. Dose–response relationships for all-or-nothing (quantal) responses are typically S-shaped and this reflects the fact that sensitivity of individuals in a population generally exhibits a normal or Gaussian distribution (bell-shaped curve). When plotted as a cumulative frequency distribution, a sigmoid dose–response curve is observed (Figure 1.2).

Studying dose response and developing dose–response models are central to determining ‘safe’ and ‘hazardous’ levels. The simplest measure of toxicity is lethality, and determination of the median lethal dose, the LD50 is usually the first toxicological test performed with new substances. The LD50 is the dose at which a substance is expected to cause the death of half of the experimental animals and it is derived statistically from dose–response curves (Eaton and Klaassen, 2001). LD50 values are the standard for comparison of acute toxicity between chemical compounds and between species. Some values are given in Table 1.1. It is important to note that the higher the LD50, the less toxic the substance.

Similarly, the EC50, the median effective dose, is the quantity of the chemical that is estimated to have an effect in 50 per cent of the organisms. However, median doses alone are not very informative, as they do not convey any information on the shape of the dose–response curve. This is best illustrated by Figure 1.3. While toxicant A seems (always) more toxic than toxicant B on the basis of its lower LD50, toxicant B will start affecting organisms at lower doses (lower threshold) while the steeper slope for the dose–response curve for toxicant A means that once individuals become overexposed (exceed the threshold dose) the increase in response occurs over much smaller increments in dose.

1.2.4.1 Low dose responses

The classic paradigm for extrapolating dose–response relationships at low doses is based on the concept of threshold for non-carcinogens, whereas for carcinogens it is assumed that there is no threshold and a linear relationship is hypothesised (Figures 1.4 and 1.5).

The NOAEL (No Observed Adverse Effect Level) is the exposure level at which there is no statistically or biologically significant increase in the frequency or severity of adverse effects between exposed population and its appropriate control. The NOEL for the most sensitive test species and the most sensitive indicator of toxicity is usually employed for regulatory purposes. The LOAEL (Lowest Observed Adverse Effect Level) is the lowest exposure level at which there is a statistically or biologically significant increase in the frequency or severity of adverse effects between exposed population and its appropriate control. The main criticism of NOAEL and LOAEL is that they are dependent on study design, i.e. the dose groups selected and the number of individuals in each group. Statistical methods of deriving the concentration that produces a specific effect ECx.

### Table 1.1 Acute LD50 of some well-known substances (adapted from Eaton and Klaassen, 2001)

<table>
<thead>
<tr>
<th>Agent</th>
<th>LD50, mg/kg body weight</th>
</tr>
</thead>
<tbody>
<tr>
<td>Ethyl alcohol</td>
<td>10,000</td>
</tr>
<tr>
<td>Sodium chloride</td>
<td>4,000</td>
</tr>
<tr>
<td>Ferrous sulphate</td>
<td>1,500</td>
</tr>
<tr>
<td>Morphine sulphate</td>
<td>900</td>
</tr>
<tr>
<td>Phenobarbital sodium</td>
<td>150</td>
</tr>
<tr>
<td>Strychnine sulphate</td>
<td>2</td>
</tr>
<tr>
<td>Nicotine</td>
<td>1</td>
</tr>
<tr>
<td>Dioxin (TCDD)</td>
<td>0.001</td>
</tr>
<tr>
<td>Botulimum toxin</td>
<td>0.000001</td>
</tr>
</tbody>
</table>