## **Code of Practice H: Carcinogens**

## 1. Chemical Carcinogens and Mutagens: Rules and Usage Guidance Notes

Information regarding the potential for materials to act as carcinogens can be found in the Hazard Statements: such materials are classified as H340, H341, H350, H351, H360, H361 *etc*.

Identification of chemical carcinogens causing cancer in man has been slow because there is a time-lag; exposure of young persons must be taken particularly seriously. Extra care is needed to avoid exposure of women who are pregnant, or liable to become pregnant, because in a number of cases the foetus has been shown to be especially sensitive to harmful substances ingested or inhaled by the mother. There is now a wide measure of overlap between carcinogens and <u>mutagens</u> (agents that cause inherited changes in the genetic message) and, to a lesser extent, with <u>teratogens</u> (agents that act early in pregnancy to cause malformations of the developing foetus).

Chemicals differ very greatly in carcinogenic potency, and there is no clear dividing line between very weak carcinogens, in which it is very difficult to establish activity with certainty, and non-carcinogens. Thus, expressed as daily dose per kilogramme of body weight to induce tumours in 50% of test cases, the carcinogenic dose of aflatoxin B<sub>1</sub> (a fungal metabolite) is about 1 mg, the dose of 2-naphthylamine (a human occupational carcinogen) is about 10 mg and the dose of the weakest (doubtful) agents, such as saccharin, approaches 10 g. Some chemicals have a weak potency and can be handled without significant risk so long as the usual rules of laboratory hygiene are observed. Others are very potent and should only be used, if at all, with strict precautions in special accommodation.

Besides actual potency, factors affecting the degree of hazard are the volatility of the substance, whether it forms a fine dust, its ease of being absorbed through the lungs, mouth or skin, and the duration and severity of exposure.

Please note the following guidance:

- (a) The use of high-risk chemical carcinogens that are not formally prohibited should be avoided wherever possible. If they must be used, the most stringent conditions must be applied to their acquisition, storage, use and disposal. They must be handled and used in such a way as to prevent entry of the carcinogens into the body of the researcher or of anyone else by any route.
- (b) When using materials that are known or suspected carcinogens or mutagens a specific risk assessment must be made in the worker's laboratory notebook and counter signed by the supervisor. This assessment should include detailed information regarding the handling, uses and disposal procedures for the material. The following list covers many known or suspected carcinogens, but there is no definitive list. Manufacturers and suppliers have a legal duty to provide and maintain material safety data sheets (MSDS) for chemicals which they supply. Upto-date MSDS should be consulted to check for whether chemicals to be used have been identified as suspected carcinogens. New materials prepared during the course of research should also be considered.

- (c) The need to avoid the use of benzene and carbon tetrachloride should be noted particularly. Formaldehyde shows mutagenic activity but there is no published work showing it to be carcinogenic. However, it would be prudent to minimise inhalation of formaldehyde fumes and to avoid its interaction, even in small concentrations, with hydrogen chloride. The product bis(chloromethyl)ether is a human carcinogen with high risk potential.
- (d) Any work involving carcinogenic, mutagenic or teratogenic materials should be subject to a separate risk assessment.

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J M Sanderson Departmental Safety Coordinator 6 September 2019

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p-Chloroaniline Benzidine [and its salts] A-alpha-C (2-Amino-9H-pyrido[2,3-Benzidine-based dyes Chlorodiphenyl (chlorinated biphenyls) 1-(2-Chloroethyl)-3-cyclohexyl-1b]indole) Benzo[b]fluoranthene Benzo[j]fluoranthene nitrosourea (CCNU) (Lomustine) Acetaldehyde Acetamide Benzo[k]fluoranthene 1-(2-Chloroethyl)-3-(4methylcyclohexyl)-1-nitrosourea Acetochlor Benzofuran 2-Acetylaminofluorene Benzo[a]pyrene (Methyl-CCNU) Acifluorfen Benzotrichloride Chloroform Acrolein Benzyl chloride bis(Chloromethyl) ether Acrylamide Benzyl violet 4B Chloromethyl methyl ether Acrylonitrile Beryllium and beryllium compounds 3-Chloro-2-methylpropene Actinomycin D Bis(2-chloroethyl)ether 4-Chloro-ortho-phenylenediamine Adriamycin (Doxorubicin N,N-Bis(2-chloroethyl)-2p-Chloro-o-toluidine hydrochloride) naphthylamine (Chlornapazine) Chlorophenols AF-2;[2-(2-furyl)-3-(5-nitro-2-Bischloroethyl nitrosourea (BCNU) Chlorophenoxy herbicides furyl)]acrylamide (Carmustine) Chloroprene Aflatoxins Bis(chloromethyl)ether and technical-Chlorothalonil Agaritine grade chloromethyl methyl ether Chlorozotocin Alachlor Bitumens, extracts of steam-refined Chromium Alcidine and air refined Chromium (hexavalent compounds) Aldrin Bleomycins Chrysene Allyl chloride Bracken fern C.I. Acid Red 114 Allyl glycidyl ether Bromodichloromethane C.I. Basic Red 9 monohydrochloride Allyl isothiocyanate 2-bromoethyl ether Ciclosporin (Cyclosporin A; Allyl isovalerate **Bromoform** Cyclosporine) Aluminium products 1,3-Butadiene Cinnamyl anthranilate 1,4-Butanediol dimethanesulfonate 2-Aminoanthraquinone Cisplatin (Busulfan, myleran) Citrus Red No. 2 p-Aminoazobenzene o-Aminoazotoluene [solvent yellow 3] Butylated hydroxyanisole (BHA) Clofibrate 4-Aminobiphenyl (4-aminodiphenyl) t-butyl methyl ether Coal gasification products 3-Amino-9-ethylcarbazole beta-Butyrolactone Coal-tars and pitches hydrochloride Cobalt metal powder 1-Amino-2-methylanthraquinone Cobalt [II] oxide Amitrole Cadmium and cadmium compounds Conjugated estrogens Ammonium dichromate Caffeic acid Copper acetoarsenite Analgesic mixtures containing Captafol Creosotes phenacetin Captan Crystal violet Androgenic (anabolic) steroids Carbazole para-Cresidine Aniline (and homologs) Carbon tetrachloride Cupferron ortho-Anisidine Carbon-black extracts Cycasin ortho-Anisidine hydrochloride Carrageenan, degraded Cyclamates para-anisidine Ceramic fibers (airborne particles of 1,4-cyclohexadiene anthanthrene respirable size) Cyclophosphamide (anhydrous) Antimony oxide (antimony trioxide) Cyclophosphamide (hydrated) Chlorambucil Chloramphenicol Arsenic (inorganic arsenic compounds) chlorbenzilat Arsine Chlordane D&C Orange No. 17 Chlordecone (Kepone) D&C Red No. 8 Asbestos Chlordimeform D&C Red No. 9 Auramine Chlorendic acid D&C Red No. 19 Azaserine Chlorinated camphene Dacarbazine Azathioprine Azacitidine Chlorinated Parrafins Daminozide Dantron (Chrysazin; 1,8-Azobenzene alpha-Chlorinated toluenes Azathioprine Chlormadinone aceate Dihydroxyanthraquinone) Chlornaphazine[n,n-bis(2-chloroethyl)dapsone 2-naphthylamine] Daunomycin Benz[a]anthracene Chlorodibromomethane **DCM** 

DDD (Dichlorodiphenyldichloroethane)

Chloroethane (ethyl chloride)

Benzene

DDE 3,3'-Dimethylbenzidine (ortho-Tolidine) Gallium arsenide (Dichlorodiphenyldichloroethylene) 3,3'-Dimethylbenzidine dihydrochloride Gasoline (petrol) DDT (Dichlorodiphenyltrichloroethane) Dimethylcarbamoyl chloride Glasswool fibers (airborne particles of DDVP (Dichlorvos) 1,1-Dimethylhydrazine (UDMH) respirable size) Decabromodiphenyl ether 1,2-Dimethylhydrazine Glu-P-1 (2-Amino-6methyldipyrido[1,2- a:3', 2'-N,N'-Diacetylbenzidine Dimethyl sulfate 2,4-Diaminoanisole Dimethylvinyl Chloride dlimidazole) Glu-P-2 (2-Aminodipyrido[1,2-a:3',2'-2,4-Diaminoanisole sulfate 2,4-dinitrofluorobenzene 4,4'-Diaminodiphenyl ether (4,4'-1,6-Dinitropyrene d]imidazole) Oxydianiline) 1,8-Dinitropyrene Glycidaldehyde 2,4-Diaminotoluene 2,4-Dinitrotoluene Glycidol Diaminotoluene (mixed) 2,6-Dinitrotoluene Griseofulvin o-Dianisidine-based dyes 1,4-Dioxane Gyromitrin (Acetaldehyde Dibenz[a,h]acridine 1,2-diphenylhydrazine methylformylhydrazone) Dibenz[a,j]acridine (hydrazobenzene) Dibenz[a,h]anthracene Diphenylhydantoin (Phenytoin) Н HC Blue 1 7H-Dibenzo[c,g]carbazole Diphenylhydantoin (Phenytoin), Dibenzo[a,e]pyrene sodium salt Heptachlor Direct Black 38 Dibenzo[a,h]pyrene Heptachlor epoxide Dibenzo[a,i]pyrene Direct Blue 6 Hexachlorobenzene Direct Brown 95 Hexachlorobutadiene Dibenzo[a,l]pyrene 1,2-Dibromo-3-chloropropane (DBCP) Disperse Blue 1 Hexachlorocyclohexanes 1,2-Dibromoethane Hexachlorodibenzodioxin Ε 2,3-Dibromo-1-propanol Hexachloroethane Dichloroacetylene Epichlorohydrin Hexamethylphosphoramide (HMPA) p-Dichlorobenzene Erionite Hydrazine Hydrazine dihydrobromide Estradiol 17B 3,3'-Dichlorobenzidine 3,3'-Dichlorobenzidine dihydrochloride Estrogens (not conjugated) Hydrazine sulfate 1,4-Dichloro-2-butene Estradiol-17 Hydroquinone 3,3'-Dichloro-4,4'-diaminodiphenyl Estrone Hydroxybutyric acid lactone ether Ethinylestradiol 1,1-Dichloroethane Mestranol 1,2-Dichloroethane Estrone Indeno [1,2,3-cd]pyrene Ethinylestradiol Dichloroethyl ether Indium trichloride Dichloromethane (Methylene chloride) Ethyl acrylate IQ (2-Amino-3-methylimidazo[4,5-1,2-Dichloropropane Ethyl methanesulfonate f]quinoline) 1,3-Dichloropropene Ethyl-4,4'-dichlorobenzilate Iron dextran complex Dieldrin Ethylene dibromide Isosafrole Dienestrol Ethylene dichloride (1,2-Diepoxybutane Dichloroethane) Κ Kepone (Chlordecone) Di(2-ethylhexyl)phthalate N-Ethyl-N-nitrosourea 1,2-Diethylhydrazine Ethylene imine Diethyl sulfate Ethylene oxide DES, Diethylstilbestrol Ethylene thiourea Lactofen Diglycidyl resorcinol ether (DGRE) Ethyleneimine Lasiocarpine Dihydrosafrole Lead Diisopropyl sulfate Lead acetate 3,3'-Dimethoxybenzidine (ortho-Folpet Lead phosphate Dianisidine) Formaldehyde (gas or aqueous Lindane and other 3,3'-Dimethoxybenzidine hexachlorocyclohexane isomers solution) dihydrochloride (ortho-dianisidine 2-(2-Formylhydrazino)-4-(5-nitro-2dihydrochloride) furyl) thiazole para-Dimethylaminoazobenzene Malonaldehyde Furan 4-Dimethylaminoazobenzene Furazolidone Mancozeb trans-2-[(Dimethylamino)methylimino]-Furmecyclox Maneb 5-[2-(5-nitro-2-furyl)vinyl]-1,3,4-Fusarin C Me-A-alpha-C (2-Amino-3-methyl-9Hoxadiazole pyrido[2, 3-b]indole)

Medroxyprogesterone acetate

G

7,12-Dimethylbenz(a)anthracene

MeIQ(2-Amino-3,4-5-Nitroacenaphthene Perylene dimethylimidazo[4,5-f]quinoline) 5-Nitro-o-anisidine Phenacetin MeIQx(2-Amino-3,8-Phenazopyridine hydrochloride o-Nitroanisole dimethylimidazo[4,5-f]quinoxaline) 4-Nitrobiphenyl Phenesterin Melphalan p-Nitrochlorobenzene Phenobarbital Merphalan Phenolphthalein 6-Nitrochrysene Mestranol Nitrofen Phenoxybenzamine Phenoxybenzamine hydrochloride Methoxychlor 2-Nitrofluorene N-Phenyl-b-naphthylamine 2-Methylaziridine (Propyleneimine) Nitrofurazone Methylazoxymethanol 1-[(5-Nitrofurfurylidene)amino]-2-Phenyl glycidyl ether Methylazoxymethanol acetate imidazollidinone Phenylhydrazine and its salts Methyl bromide 1-[(5-Nitrofurfurylidene)-N-[4-(5-Nitroo-Phenylphenate, sodium 3-Methylcholanthrene 2-furyl)-2 thiazolyl]acetamide 2-Phenylphenol Nitrogen mustard (Mechlorethamine) 5-Methylchrysene Phenytoin 4,4'-Methylene bis(2-chloroaniline) Nitrogen mustard hydrochloride PhiP(2-Amino-1-methyl-6-(MOCA, MBOCA) (Mechlorethamine hydrochloride) phenylimidazol[4,5-b]pyridine) 4,4'-Methylene bis(N,N-Nitrogen mustard N-oxide Polybrominated biphenyls dimethyl)benzenamine Nitrogen mustard N-oxide Polychlorinated biphenyls Methylene chloride hydrochloride Polychlorinated dibenzo-p-dioxins 4,4'-Methylene bis(2-methylaniline) 2-Nitronaphthalene Polychlorinated dibenzofurans 4,4'-Methylenedianiline (MDA) 2-Nitropropane Polycyclic aromatic hydrocarbons 4,4'-Methylenedianiline dihydrochloride 4-Nitropyrene Polygeenan Methylhydrazine and its salts N-Nitrosodi-n-butylamine Ponceau MX Methyl chloromethyl ether N-Nitrosodiethanolamine Ponceau 3R Methyl-CCNU N-Nitrosodiethylamine Potassium bromate Methyl iodide N-Nitrosodimethylamine Potassium dichromate Methyl methanesulfonate Procarbazine p-Nitrosodiphenylamine Procarbazine hydrochloride 2-Methyl-1-nitroanthraquinone N-Nitrosodiphenylamine N-Methyl-N'-nitro-N-nitrosoguanidine N-Nitrosodi-n-propylamine Procymidone (MNNG) N-Nitroso-N-ethylurea Progesterone N-Methyl-N-nitrosourea 3-(N-Nitrosomethylamino)propionitrile **Progestins** N-Methylolacrylamide 4-(N-Nitrosomethylamino)-1-(3-1,3-Propane sultone Methylthiouracil pyridyl)-1-butanone (NNK) Progargite Metiram N-Nitrosomethylethylamine beta-Propiolactone Metronidazole N-Nitroso-N-methylurea Propylene dichloride Michler's ketone N-Nitroso-N-methylurethane Propylene imine Mineral Oils, untreated and mildly N-Nitrosomethylvinylamine Propylene oxide treated N-Nitrosomorpholine Propylthiouracil Mirex N-Nitrosonornicotine Pyridinium chlorochromate Mitomycin C N-Nitrosopiperidine **MOPP** N-Nitrosopyrrolidine Radionuclides Monocrotaline N-Nitrososarcosine 5-(Morpholinomethyl)-3-[(5-nitro-Norethisterone (Norethindrone) Radon fufurylidene)-amino]-2-oxazolidinone Reserpine Mustard gas 0 Residual (heavy) fuel oils Ochratoxin A Rhodamine 101 N Oestrogen, nonstreoidal Rosin core solder Nafenopin Oestrogen, steroidal 1-Naphthylamine Oil Orange SS S 2-Naphthylamine 4,4'-Oxydianiline Saccharin 3-Naphthylamine Oxadiazon Saccharin, sodium Nickel and certain nickel compounds Oxymetholone Safrole Nickel carbonyl Oxazepam Selenium sulfide Nickel subsulfide Shale-oils Niridazole P Silica, crystalline (airborne particles of Nitrilotriacetic acid Panfuran S respirable size) Nitrilotriacetic acid, trisodium salt Pentachloroethane Sodium chromate tetrahydrate

Sodium dichromate

Pentachlorophenol

monohydrate

Sodium hexafluoroarsenate(V)

Sodium ortho-phenylphenate

Sterigmatocystin Streptozotocin

Strontium chromate

Styrene Styrene oxide Sulfallate Sulfur trioxide Sulphur trioxide N,N-

dimethylformamide complex

T

Talc containing asbestiform fibers

Terrazole

Testosterone and its esters

 $2, 3, 7, 8\hbox{-} Tetra chlorodibenzo-para-diox in$ 

(TCDD, dioxin)

1,1,2,2-Tetrachloroethane

Tetrachloroethylene

(Perchloroethylene)

p-a,a,a-Tetrachlorotoluene

3,3',5,5'-tetramethylbenzidine

Tetranitromethane Thioacetamide

4,4'-Thiodianiline

Thiourea

Thorium dioxide Titanium dioxide

Tobacco, oral use of smokeless

products

Tobacco smoke
Toluene diisocyanate

p-toluenesulphonic acid

ortho-Toluidine

ortho-Toluidine-based dyes

ortho-Toluidine hydrochloride

para-Toluidine

Toxaphene

Tremolite silicates

Treosulfan (Tresoluphan)

Trichlormethine (Trimustine

hydrochloride)

1,1,2-Trichloroethane

Trichloroethylene

2,4,6-Trichlorophenol

1,2,3-Trichloropropane

Triphenyltin hydroxide

Tris(aziridinyl)-para-benzoquinone

(Triaziquone)

Tris(1-aziridinyl)phosphine sulfide

(Thiotepa)

Tris(2-chloroethyl) phosphate

Tris(2,3-dibromopropyl)phosphate

TRIZMA base

Trp-P-1 (Tryptophan-P-1) (3-Amino-

1,4-dimethyl-5H-pyrido[4,3-b]indole)

Trp-P-2 (Tryptophan-P-2) (3-Amino-1-methyl-5H-pyrido[4,3-b]indole)
Trypan blue

U

Uracil mustard

Uranium compounds

Urethane (Ethyl carbamate)

V

Vinyl bromide

Vinyl chloride

4-Vinyl-1-cyclohexene diepoxide (Vinyl

cyclohexene dioxide) Vinylidene chloride (1,1dichloroethylene)

n-vinyl pyrollidone

Vinyl pivalate

Vinyl trichloride (1,1,2-Trichloroethane)

X

2,6-Xylidine (2,6-Dimethylaniline)

Ζ

Zinc chromate

Zineb

**Source**: NIOSH (27/6/2017) and University of Oxford Department of Chemistry MSDS web resource (pre-2011).