Benomyl

Methyl Benomilo. I-(butylcarbamoyl)benzimidazol-2-ylcarbamate.

 $C_{14}H_{18}N_4O_3 = 290.3$ CAS - 17804-35-2.

Profile

Benomyl is a fungicide used for the treatment and control of fungal plant diseases

♦ References.

- 1. WHO. Benomyl. Environmental Health Criteria 148. Geneva: WHO, 1993. Ávailable at: http://www.inchem.org/documents/ehc/ehc/48.htm (accessed 23/04/04)
- WHO. Benomyl health and safety guide. IPCS Health and Safety Guide 81. Geneva: WHO, 1993. Available at: http:// www.inchem.org/documents/hsg/hsg/hsg81_e.htm (accessed 23/04/04)

Toxicity. Although experimental evidence in animals has suggested a possible link between benomyl and congenital eye defects (anophthalmia) the association could not be confirmed in humans.1-

- 1. Gilbert R. "Clusters" of anophthalmia in Britain. BMJ 1993;
- Bianchi F, et al. Clusters of anophthalmia. BMJ 1994; **308:** 205. Kristensen P, Irgens LM. Clusters of anophthalmia. BMJ 1994; **309:** 205.
- 4. Castilla EE. Clusters of anophthalmia. BMJ 1994; 308: 206.

Benzyl Benzoate

Bencilo, benzoato de; Bensylbensoat; Bentsyylibentsoaatti; Benzil Benzoat; Benzil-benzoát; Benzilbenzoatas; Benzoato de bencilo; Benzoato de Benzilo; Benzoesäurebenzylester; Benzyl Benz.; Benzyl-benzoát; Benzyle, benzoate de; Benzylis benzoas; Benzylu

 C_6H_5 .CO.O.C H_2 . $C_6H_5 = 212.2$. CAS — 120-51-4. ATC — P03AX01. ATC Vet — QP53AX11.

Pharmacopoeias. In Eur. (see p.vii), Int., Jpn, and US. Ph. Eur. 6.2 (Benzyl Benzoate). Colourless or almost colourless crystals, or a colourless or almost colourless oily liquid. Fp. is not below 17°. Practically insoluble in water; miscible with alcohol, with dichloromethane, and with fatty and essential oils. Store in well-filled airtight containers. Protect from light.

USP 31 (Benzyl Benzoate). A clear, colourless, oily liquid with a slight aromatic odour. Practically insoluble in water and in glycerol; miscible with alcohol, with chloroform, and with ether. Store at a temperature not exceeding 40° in well-filled airtight containers. Protect from light.

Adverse Effects and Treatment

Benzyl benzoate is irritant to the eyes and mucous membranes and it may be irritant to the skin. Hypersensitivity reactions have been reported. If ingested, benzyl benzoate may cause stimulation of the CNS and convulsions. Systemic symptoms have been reported on excessive topical use. For poisoning associated with topical use the skin should be washed. Appropriate symptomatic measures should also be instituted.

Uses and Administration

Benzyl benzoate is an acaricide used in the treatment of scabies (p.2035) although other treatments are generally preferred. A 25% emulsion is applied to the whole body, usually from the neck down (although the BNF considers that application should be extended to the scalp, neck, face, and ears). If the application is thorough, one treatment may suffice, although the possibility of failure is lessened by a second application within 5 days. Alternatively, three applications at 12-hour intervals, without bathing, may be made, followed by bathing 12 hours after the last application. The BNF recommends one application to the whole body, repeated, without bathing, on the next day, and washed off

24 hours later; a third application may sometimes be necessary. Benzyl benzoate is not generally recommended for infants and children, but if used the application should be diluted to minimise the risk of irritation, although this also reduces efficacy.

Benzyl benzoate has also been used as a pediculicide.

Benzyl benzoate is also used as a solubilising agent.

Preparations

BP 2008: Benzyl Benzoate Application; USP 31: Benzyl Benzoate Lotic

Proprietary Preparations (details are given in Part 3)

Austral.: Ascabiol; Benzemul; Braz.: Acarsan; Bencocan; Benzibel†; Benzein†; Benzocax; Benzoben†; Benzocan†; Benzolato†; Benzolina†; Benzolan†; Benzolina†; Benzolato†; Samezan†; Samezan*; Samezan* Samilab; Samodex; Scabenzil; Scabioid; Zilaben; Ger.: Acaril†; Acarosan†; Antiscabiosum; Gr.: Benzogal; Irl.: Ascabiol†; Israel: Scabiex; Ital.: Mom Lozione Preventiva; Mex.: Ansar; Hastilan; Pol.: Novoscabin; Port.: Acar ilbial; Neo-Acarina†; Piozil; **S.Afr.:** Ascabiol; **UK:** Ascabiol; **Venez.:** Benzalcor; Benzo-Bencil; Benzodit†; Niostal†.

Multi-ingredient: Arg.: Anusol Duo S; Anusol-A; Amecrem†; Bencil Multi-ingredient: Arg.: Anusol Duo S; Anusol-A; Amecrem; Bencii Scab; Detebencii; Hevabencii; Perbel; Permecii; Sapucai; Scabioderm: Austral.: Anusoi; Belg.: Pulmex; Pulmex Baby; Braz.: Anusol-HC; Fr.: Allerbiocid S; Ascabio; Sanytol; Hong Kong: Anusol-HC; Hung.: Novascabin; H.I.: Anugesic-HC; Anusol-HC; Ital.: Antiscabbia Camidio il a DDT Terapeutico; Antiscabbia CM; Dekar 2; Prurex; Skab 2; Malaysia: Anucare; Anusol; NZ: Anusol; Pol.: Cetriscabin; S.Afr.: Anugesic; Singapore: Anusol; Spain: Tulgrasum Cicatrizante; Yacutir; Swed.: Tenutex; Thai: Anusol; UK: Anugesic-HC; Anusol-HC, Plus HC; Sudocrem; USA: Anumed; Anumed HC; Hemril; Venez.: Kertyol.

Bioallethrin (BAN)

Allethrin I; Bioaletrina; Depallethrin. (RS)-3-Allyl-2-methyl-4-oxocyclopent-2-enyl (1R,3R)-2,2-dimethyl-3-(2-methylprop-1-enyl)cyclopropanecarboxylate.

 $C_{19}H_{26}O_3 = 302.4$. CAS — 584-79-2. ATC — P03AC02. ATC Vet - QP53AC02.

Bioallethrin is a pyrethroid insecticide (see Pyrethrum Flower, p.2049). It is used topically, with the synergist piperonyl butoxide (p.2049), in the treatment of pediculosis (p.2034). It is also used in anti-mosquito devices and for the control of household insect pests.

◊ References.

- WHO, Allethrins. Environmental Health Criteria 87. Geneva: WHO, 1989. Available at: http://www.inchem.org/documents/ehc/ehc/ehc87.htm (accessed 23/04/04)
 WHO. Allethrins health and safety guide. IPCS Health and Safety Guide 24. Geneva: WHO, 1989. Available at: http://www.inchem.org/documents/hsg/hsg/hsg024.htm (accessed 22/04/04) 23/04/04)

Preparations

Proprietary Preparations (details are given in Part 3)

Multi-ingredient: Arg.: Limpacid; Para Piojicida; Scabioderm; Austral.: Paralice†, Belg.: Para: Braz.: Samapen†, Canad.: Para†; Fr.: Para Special Poux; Ger.: Jacutin N; Spregal; Israel: Monocide; Ital.: Cruzzy; Neth.:

Brodifacoum

Brodifacum; WBA-8119. 3-[3-(4'-Bromobiphenyl-4-yl)-1,2,3,4tetrahydro-I-naphthyl]-4-hydroxycoumarin.

 $C_{31}H_{23}BrO_3 = 523.4.$ - 56073-10-0.

Brodifacoum is an anticoagulant rodenticide. It is reported to be effective in warfarin-resistant strains of rodents.

◊ References.

- 1. WHO. Anticoagulant rodenticides. Environmental Health Criteria 175. Geneva: WHO, 1995. Available at: http://www.inchem.org.documents/ehc/ehc/ehc175.htm (accessed 23/04/04)
- WHO. Brodifacoum health and safety guide. IPCS Health and Safety Guide 93. Geneva: WHO, 1995. Available at: http:// www.inchem.org/documents/hsg/hsg/hsg093.htm (accessed

Toxicity. Brodifacoum, a second-generation anticoagulant rodenticide, inhibits prothrombin synthesis to cause bleeding that may be occult.1 It is absorbed from the gastrointestinal tract; dermal absorption is possible. Poisons containing 100 mg in each kg of bait are not hazardous to man; more concentrated forms are particularly hazardous and their availability should be restricted. Baits, which should be prepared only by trained personnel, should contain a suitable marker-dye.

There have been reports of poisoning with brodifacoum.²⁻¹⁰

- 1. WHO. Safe use of pesticides: ninth report of the WHO expert committee on vector biology and control. WHO Tech Rep Ser 720 1985. Available at: http://libdoc.who.int/trs/WHO_TRS_720.pdf (accessed 21/07/08)

 2. Watts RG, et al. Accidental poisoning with a superwarfarin compound (brodifacoum) in a child. Pediatrics 1990; 86: 883–7.
- 3. Ross GS, et al. An acquired hemorrhagic disorder from long-acting rodenticide ingestion. Arch Intern Med 1992; 152: 410–12.
- Kruse JA, Carlson RW. Fatal rodenticide poisoning with brodi-facoum. Ann Emerg Med 1992; 21: 331–6.
- Tecimer C, Yam LT. Surreptitious superwarfarin poisoning with brodifacoum. South Med J 1997; 90: 1053–5.
- Corke PJ. Superwarfarin (brodifacoum) poisoning. Anaesth Intensive Care 1997; 25: 707–9.
- 7. La Rosa FG. et al. Brodifacoum intoxication with marijuana
- La Rosa F4, et al. Brodifacoum intoxication with marijuana smoking, Arch Pathol Lab Med 1997; 121: 67–9.
 Miller MA, et al. Rapid identification of surreptitious brodifa-coum poisoning by analysis of vitamin K-dependent factor ac-tivity. Am J Emerg Med 2006; 24: 383.
- Olmos V, López CM. Brodifacoum poisoning with toxicokinetic data. Clin Toxicol 2007; 45: 487–9.
- Kapadia P, Bona R. Acquired deficiency of vitamin K-dependent clotting factors due to brodifacoum ingestion. Conn Med 2008; 72: 207–9.

Bromadiolone

Bromadiolona. 3-[3-(4'-Bromobiphenyl-4-yl)-3-hydroxy-1-phenylpropyl]-4-hydroxycoumarin.

 $C_{30}H_{23}BrO_4 = 527.4$ CAS — 28772-56-7

Profile

Bromadiolone is an anticoagulant rodenticide.

♦ References.

- WHO. Anticoagulant rodenticides. Environmental Health Criteria 175. Geneva: WHO, 1995. Available at: http://www.inchem.org/ documents/ehc/ehc/ehc175.htm (accessed 23/04/04)
- 2. WHO. Bromadiolone health and safety guide. *IPCS Health and Safety Guide 94*. Geneva: WHO, 1995. Available at: http://www.inchem.org/documents/hsg/hsg/hsg094.htm (accessed

Toxicity. Bromadiolone, a second-generation anticoagulant rodenticide, inhibits prothrombin synthesis to cause bleeding that may be occult.1 It is absorbed from the gastrointestinal tract; dermal absorption is possible. Poisons containing 100 mg in each kg of bait are not hazardous to man; more concentrated forms are particularly hazardous and their availability should be restricted. Baits, which should be prepared only by trained personnel, should contain a suitable marker-dye.

There have been reports of poisoning with bromadiolone.²⁻⁵

- 1. WHO. Safe use of pesticides: ninh report of the WHO expert committee on vector biology and control. WHO Tech Rep Ser 720 1985. Available at: http://libdoc.who.int/trs/WHO_TRS_720.pdf (accessed 21/07/08)
- 2. Greeff MC, et al. "Superwarfarin" (bromodialone) poisoning in two children resulting in prolonged anticoagulation. Lancet 1987; ii: 1269
- 3. Chow EY, et al. A case of bromadiolone (superwarfarin) ingestion. CMAJ 1992; 147: 60-2.
- 4. Grobosch T, et al. Acute bromadiolone intoxication. J Anal Toxicol 2006; 30: 281-6.
- Lo VM, et al. Bromadiolone toxicokinetics: diagnosis and treatment implications. Clin Toxicol 2008; 1–8.

Bromophos

Bromofós; Bromofos; OMS-658. O-4-Bromo-2,5-dichlorophenyl O,O-dimethyl phosphorothioate.

 $C_8H_8BrCl_2O_3PS = 366.0.$ CAS - 2104-96-3. ATC Vet — QP53AF11.

Profile

Bromophos is an organophosphorus insecticide (p.2047) used in veterinary practice for the control of ectoparasites in the environment. It has also been used as an agricultural insecticide.

Butopyronoxyl

Butopiroxinilo; Indalone. Butyl 3,4-dihydro-2,2-dimethyl-4-oxo-2H-pyran-6-carboxylate.

 $C_{12}H_{18}O_4 = 226.3.$ CAS — 532-34-3.

Profile

Butopyronoxyl has been used as an insect repellent.

Carbamate Insecticides

Insecticidas del grupo de los carbamatos.

Description. The carbamate insecticides are N-substituted esters of carbamic acid.

 WHO. Carbamate pesticides: a general introduction. Environmental Health Criteria 64. Geneva: WHO, 1986. Available at: http:// www.inchem.org/documents/ehc/ehc/ehc64.htm (accessed 24/07/08)

Adverse Effects

As for Organophosphorus Insecticides, p.2047.

The carbamates are cholinesterase inhibitors, differing from the organophosphorus insecticides in that the inhibition they produce is generally less intense and more rapidly reversible. In addition, they do not appear to enter the CNS as readily and severe central effects are therefore uncommon.

Treatment of Adverse Effects

If substantial amounts of carbamate insecticides have been ingested the use of gastric lavage should be considered if the patient presents within 1 hour. Contaminated clothing should be removed and the skin washed with soap and water. Treatment is largely symptomatic and supportive and includes atropine, but this may not always be necessary due to the rapidly reversible nature of the cholinesterase inhibition produced. Pralidoxime is thought to be generally unnecessary; some animal studies have suggested that it may increase the toxicity of carbamates

♦ References.

- 1. WHO. Safe use of pesticides: fourteenth report of the WHO expert committee on vector biology and control. WHO Tech Rep Ser 813 1991. Available at: http://libdoc.who.int/trs/ WHO_TRS_813.pdf (accessed 21/07/08)
- Proudfoot A, ed. Pesticide poisoning: notes for the guidance of medical practitioners. 2nd ed. London: DoH, The Stationery Office, 1996.

Carbaryl (BAN)

Carbaril (pINN); Carbarilo; Carbarilum; Karbaril; OMS-29. I-Naphthyl methylcarbamate.

Карбарил

 $C_{12}H_{11}NO_2 = 201.2.$ CAS — 63-25-2. ATC Vet - OP53AE01.

Pharmacopoeias. In Br.

BP 2008 (Carbaryl). A white to off-white or light grey powder which darkens on exposure to light. Very slightly soluble in water; soluble in alcohol and in acetone. Store at a temperature not exceeding 25°. Protect from light.

Adverse Effects and Treatment

As for Carbamate Insecticides, above. Carbaryl may be absorbed after ingestion, inhalation, or skin contamination.

Carbaryl has been reported to produce neoplasms in mice and rats and in late 1995 the UK DoH advised that it would be prudent to consider carbaryl as a potential human carcinogen; its medicinal use was limited to prescription only. However, the DoH emphasised that the risk was a theoretical one and that any risk from the intermittent use of head lice preparations was likely to be very small.

Uses and Administration

Carbaryl is a carbamate insecticide (above). It is used as a 0.5 or 1.0% lotion or shampoo in the treatment of head and pubic pediculosis (p.2034). Lotions are generally preferred to shampoos as the contact time is longer. Aqueous lotions are preferred to treat pubic lice because alcoholic lotions are irritant to excoriated skin and the genitalia; aqueous lotions may also be preferable in asthmatic subjects or children to avoid alcoholic fumes. Skin or hair treated with an alcohol-based preparation should be allowed to dry naturally.

Carbaryl is also used as a topical ectoparasiticide in veterinary practice and as an agricultural, horticultural, and household insecticide.

- 1. WHO. Carbaryl health and safety guide. *IPCS Health and Safety Guide* 78. Geneva: WHO, 1993. Available at: http://www.inchem.org/documents/hsg/hsg/hsg78_e.htm (accessed
- WHO. Carbaryl. Environmental Health Criteria 153. Geneva: WHO, 1994. Available at: http://www.inchem.org/documents/ ehc/ehc/ehc153.htm (accessed 23/04/04)

Preparations

BP 2008: Carbaryl Lotion.

Proprietary Preparations (details are given in Part 3) Israel: Hafif; UK: Carylderm.

Carbosulfan

Carbosulfán. 2,3-Dihydro-2,2-dimethylbenzofuran-7-yl (dibutylaminothio)methylcarbamate.

 $C_{20}H_{32}N_2O_3S = 380.5.$ CAS - 55285-14-8.

Carbosulfan is a carbamate insecticide (p.2037) used in agriculture and for the larvicidal treatment of rivers in the control of onchocerciasis (p.137).

Chloralose (HNN)

Alphachloralose; Chloralosane; α-Chloralose; Chloralosum; Cloralosa; Glucochloral. (R)-1,2-O-(2,2,2-Trichloroethylidene)- α -Dglucofuranose.

 $C_8H_{11}CI_3O_6 = 309.5$ CAS — 15879-93-3.

The symbol † denotes a preparation no longer actively marketed

Profile

Chloralose has general properties similar to those of cloral hydrate (p.979), of which it is a derivative. It is used as a rodenticide. It was formerly used for its hypnotic properties.

Chlordane

Chlordan; Clordano. 1,2,4,5,6,7,8,8-Octachloro-2,3,3a,4,7,7ahexahydro-4,7-methanoindene.

 $C_{10}H_6CI_8 = 409.8.$ CAS — 57-74-9.

Chlordane is a chlorinated insecticide (p.2037). Its use is limited, or even prohibited, in some countries because of toxicity due to its persistent nature.

♦ References.

- 1. Kutz FW, et al. A fatal chlordane poisoning. J Toxicol Clin Toxicol 1983; 20: 167-74.
- 2. Olanoff LS, et al. Acute chlordane intoxication. J Toxicol Clin Toxicol 1983: 20: 291-306.
- WHO. Chlordane. Environmental Health Criteria 34. Geneva: WHO, 1984. Available at: http://www.inchem.org/documents/ehc/ehc/ehc34.htm (accessed 23/04/04)
- WHO. Chlordane health and safety guide. IPCS Health and Safety Guide 13. Geneva: WHO, 1988. Available at: http://www.inchem.org/documents/hsg/hsg/hsg013.htm (accessed 2201/16).

Chlorinated Insecticides

Insecticidas clorados

Adverse Effects

Chlorinated or organochlorine insecticides form a very wide group and the toxicity of individual members varies considerably. In general these insecticides produce symptoms consistent with CNS stimulation. They may be absorbed through the respiratory and gastrointestinal tracts and through the skin.

Symptoms of acute poisoning include nausea and vomiting, paraesthesia, giddiness, tremors, convulsions, coma, and respiratory failure. Liver, kidney, and myocardial toxicity have been reported. Effects on the blood include agranulocytosis and aplastic anaemia. Symptoms may be complicated by the effects of the solvent

Chlorinated insecticides have been reported to enhance microsomal hepatic enzyme activity. Skin reactions can occur after

Polychlorinated biphenyl (PCB) and terphenyl compounds were formerly used as insecticides in many countries. They accumulate in body fat and are not readily excreted, although they are distributed into breast milk and possibly cross the placenta; because of this and because of accidental contamination they remain a cause for concern. The related polybrominated biphenyl compounds (PBB), which have no insecticidal uses, have also been absorbed by humans after accidental contamination of the food chain.

Some chlorinated insecticides have weak oestrogenic effects; it has been proposed that exposure may increase the risk of breast

Treatment of Adverse Effects

If chlorinated insecticides have been ingested gastric lavage or activated charcoal may be considered if the patient presents within 1 hour. Contaminated clothing should be removed and the skin washed with soap and water. Treatment is largely symptomatic