

unsaturated propellants, the number 1 is used as the fourth digit from the right to indicate an unsaturated double bond.

Thus for dichlorodifluoromethane (propellant 12):

- (a) First digit = 0 signifies number of C atoms = 1
- (b) Second digit = 1 signifies number of H atoms = 0
- (c) Third digit = 2 signifies number of F atoms = 2
- (d) Number of Cl atoms = 4 - (2 - 0) = 2

Under the terms of the Montreal Protocol, aimed at reducing damage to the ozone layer, the use of chlorofluorocarbons, including dichlorodifluoromethane, dichlorotetrafluoroethane, and trichloromonofluoromethane, has been prohibited from January 1996.<sup>(2-6)</sup> However, this prohibition does not apply to essential uses such as existing pharmaceutical formulations for which no alternative chlorofluorocarbon-free product is available. The EPA and FDA approved essential-use status for dichlorodifluoromethane for a sterile aerosol talc used in the treatment of malignant pleural effusion in patients with lung cancer.

## 19 Specific References

- 1 Health and Safety Executive. *EH40/2005: Workplace Exposure Limits*. Sudbury: HSE Books, 2011. <http://www.hse.gov.uk/pubns/priced/eh40.pdf> (accessed 1 March 2012).
- 2 Fischer FX, et al. CFC propellant substitution: international perspectives. *Pharm Technol* 1989; 13(9): 44, 48, 50, 52.
- 3 Kempner N. Metered dose inhaler CFC's under pressure. *Pharm J* 1990; 245: 428-429.
- 4 Dalby RN. Possible replacements for CFC-propelled metered-dose inhalers. *Med Device Technol* 1991; 2(4): 21-25.
- 5 CFC-free aerosols; the final hurdle. *Manuf Chem* 1992; 63(7): 22-23.
- 6 Mackenzie D. Large hole in the ozone agreement. *New Scientist* 1992; Nov 28: 5.

## 20 General References

- Amin YM, et al. Fluorocarbon aerosol propellants XII: correlation of blood levels of trichloromonofluoromethane to cardiovascular and respiratory responses in anesthetized dogs. *J Pharm Sci* 1979; 68: 160-163.
- Byron PR, ed. *Respiratory Drug Delivery*. Boca Raton, FL: CRC Press, 1990.
- Johnson MA. *The Aerosol Handbook*, 2nd edn. Caldwell: WE Dorland, 1982: 305-335.
- Niazi S, Chiou WL. Fluorocarbon aerosol propellants XI: pharmacokinetics of dichlorodifluoromethane in dogs following single dose and multiple dosing. *J Pharm Sci* 1977; 66: 49-53.
- Sanders PA. *Handbook of Aerosol Technology*, 2nd edn. New York: Van Nostrand Reinhold, 1979: 19-35.
- Sawyer E, et al. Microorganism survival in non-CFC propellant P134a and a combination of CFC propellants P11 and P12. *Pharm Technol* 2001; 25(3): 90-96.
- Sciarra JJ. Pharmaceutical aerosols. In: Lachman L et al. eds. *The Theory and Practice of Industrial Pharmacy*, 3rd edn. Philadelphia: Lea and Febiger, 1986: 589-618.
- Sciarra CJ, Sciarra JJ. Aerosols. In: *Remington: The Science and Practice of Pharmacy*, 21st edn. Philadelphia, PA: Lippincott, Williams and Wilkins, 2006: 1000-1017.
- Sciarra CJ, Sciarra JJ. Pressurized dispensers. In: Schlossman ML. *The Chemistry and Manufacture of Cosmetics*, vol. 1, 4th edn. Carol Stream, IL: Allured Publishing Corporation, 2009: 451-478.
- Sciarra JJ. Inc: Banker GS, Rhodes C, eds. *Modern Pharmaceutics*, 3rd edn. New York: Marcel Dekker, 1996: 547-574.
- Sciarra JJ, Stoller L. *The Science and Technology of Aerosol Packaging*. New York: Wiley, 1974: 97-130.
- Strobach DR. Alternatives to CFCs. *Aerosol Age* 1988; 32-33: 42-43.

## 21 Authors

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## 22 Date of Revision

4 May 2017.

# Chloroxylenol

## 1 Nonproprietary Names

BP: Chloroxylenol

USP-NF: Chloroxylenol

## 2 Synonyms

4-Chloro-3,5-dimethylphenol; 2-chloro-5-hydroxy-1,3-dimethylbenzene; 4-chloro-1-hydroxy-3,5-dimethylbenzene; 2-chloro-5-hydroxy-*m*-xylene; 2-chloro-*m*-xylenol; 3,5-dimethyl-4-chlorophenol; *Nipacide PX*; parachlorometaxylenol; *p*-chloro-*m*-xylenol; PCMX.

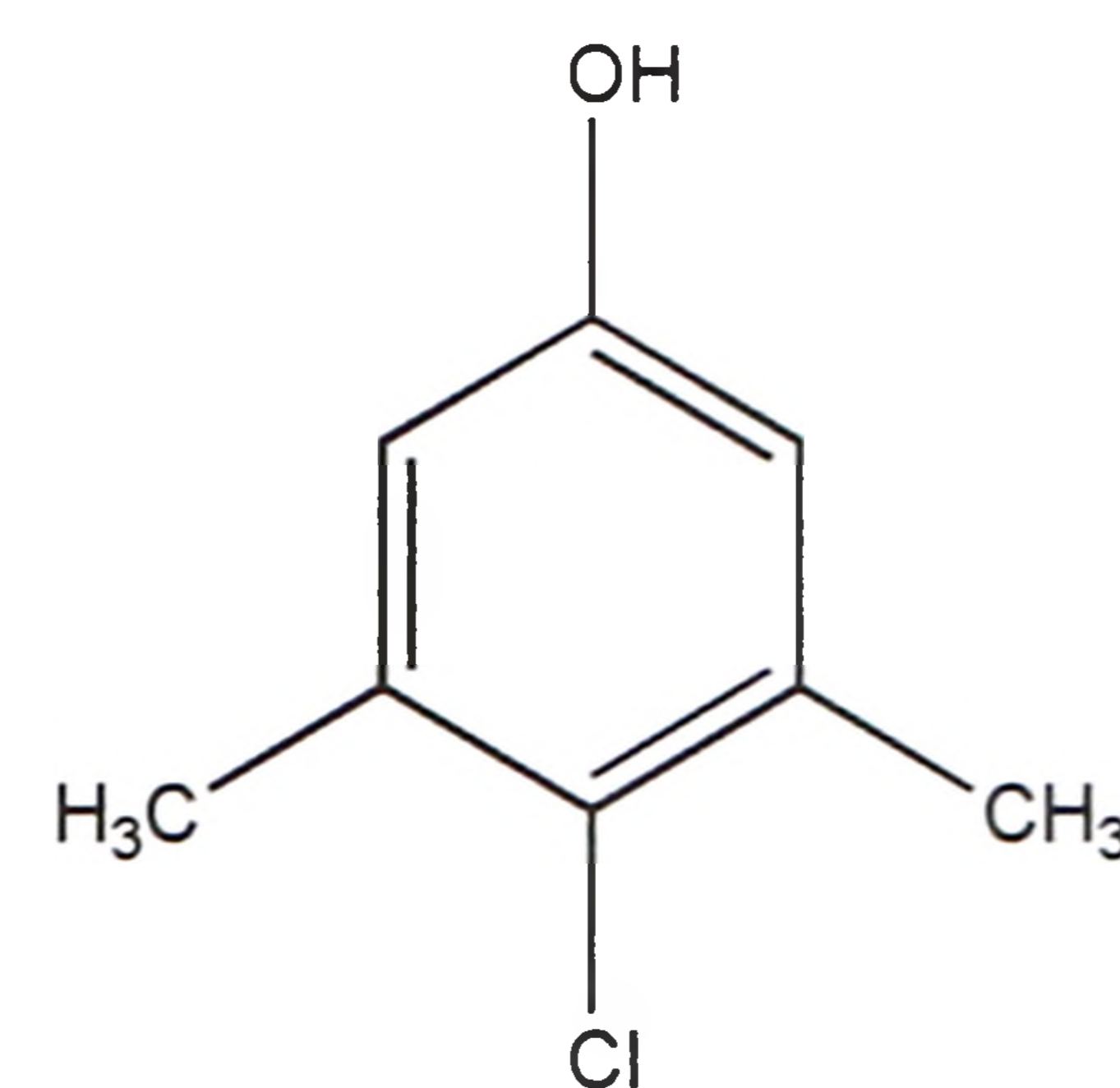
## 3 Chemical Name and CAS Registry Number

4-Chloro-3,5-xyleneol [88-04-0]

## 4 Empirical Formula and Molecular Weight

C<sub>8</sub>H<sub>9</sub>ClO 156.61

## 5 Structural Formula



## 6 Functional Category

Antimicrobial preservative.

## 7 Applications in Pharmaceutical Formulation or Technology

As a pharmaceutical excipient, chloroxylenol is commonly used in low concentrations, typically at concentrations of 0.1-0.8%, as an