

some countries. Carbonated solutions of prilocaïne have also been tried in some countries in epidural and brachial plexus nerve blocks (see under Administration, p.1852). Prilocaïne is used for surface anaesthesia in a eutectic mixture with lidocaïne. (Local anaesthetic techniques are discussed on p.1853.)

The dosage used in various **local anaesthetic procedures** varies with the site of injection and the procedure used. The recommended **maximum single dose** in adults for prilocaïne hydrochloride is 400 mg if used alone, or 300 mg if used with felypressin. Doses should be reduced in elderly or debilitated patients. The dose for children over 6 months of age is up to 5 mg/kg. For **dental infiltration or dental nerve blocks**, the usual adult dose of prilocaïne hydrochloride without felypressin is 40 to 80 mg (1 to 2 mL) as a 4% solution; children under 10 years generally require about 40 mg (1 mL). Similar doses of the 4% solution with adrenaline (1:200 000) may be used for most routine dental procedures. The usual adult dose of prilocaïne hydrochloride with felypressin 0.03 international units/mL is 30 to 150 mg (1 to 5 mL) as a 3% solution; children under 10 years generally require 30 to 60 mg (1 to 2 mL).

A eutectic mixture of prilocaïne base 2.5% and lidocaïne base 2.5% (see Surface Anaesthesia, under Lidocaïne, p.1866) is applied as a cream under an occlusive dressing to produce **surface anaesthesia** of the skin before procedures requiring needle puncture, surgical treatment of localised lesions, and split skin grafting; it has been used similarly, but without an occlusive dressing, before removal of genital warts.

Action. For a comparison of the vasoactivity of prilocaïne and some other local anaesthetics, see p.1852.

Infiltration anaesthesia. Addition of felypressin at a concentration of 0.03 international units/mL to prilocaïne 3% injection did not reduce plasma concentrations of prilocaïne after infiltration of a 60-mg dose into the upper premolar region.¹

1. Cannell H, Whelpton R. Systemic uptake of prilocaïne after injection of various formulations of the drug. *Br Dent J* 1986; **160**: 47-9.

Preparations

BP 2008: Prilocaïne Injection;

USP 31: Lidocaïne and Prilocaïne Cream; Prilocaïne and Epinephrine Injection; Prilocaïne Hydrochloride Injection.

Proprietary Preparations (details are given in Part 3)

Austral.: Citanest; Citanest Dental; **Belg.:** Citanest; **Braz.:** Citanest; Citocaina; **Canad.:** Citanest; **Denm.:** Citanest Octapressin; **Fin.:** Citanest Octapressin; **Ger.:** Xylonest; **Ital.:** Citanest con Octapressin; **Mex.:** Citanest Octapressin; **Neth.:** Citanest; Citanest Octapressin; **Norw.:** Citanest Octapressin; **NZ:** Citanest; Citanest with Octapressin; **Spain:** Citanest; Citanest Octapressin; **Swed.:** Citanest; Citanest Octapressin; **Switz.:** Citanest Octapressin; Xylonest; **Turk.:** Citanest; Citanest Octapressin; **UK:** Citanest; Citanest with Octapressin; **USA:** Citanest.

Multi-ingredient: **Arg.:** Emla; **Austral.:** Emla; **Austria:** Emla; **Belg.:** Emla; **Braz.:** Emla; **Canad.:** Emla; **Chile:** Eutecaina; **Cz.:** Emla; **Denm.:** Emla; **Oraqix Fin.:** Emla; **Oraqix Fr.:** Emla; Emlapatch; **Oraqix Ger.:** Emla; **Gr.:** Emla; **Pinex Hong Kong:** Emla; **Indon.:** Emla; **Estesia:** Topsy; **Irl.:** Emla; **Israel:** Emla; **Ital.:** Emla; **Malaysia:** Emla; **Mex.:** Emla; **Neth.:** Emla; **Oraqix Norw.:** Emla; **Oraqix NZ:** Emla; **Philipp.:** Emla; **Pol.:** Emla; **Port.:** Emla; **Oraqix Rus.:** Emla (Эмла); **S.Afr.:** Emla; **Topla Singapore:** Emla; **Spain:** Emla; **Swed.:** Emla; **Oraqix Switz.:** Emla; **Thai.:** Emla; **Turk.:** Emla; **UK:** Emla; **Oraqix USA:** Emla.

Procaïne Hydrochloride (BANM, rINN)

Allocaïne; Ethocaïne Hydrochloride; Hidrocloruro de procaína; Novocaïn; Procaïne, chlorhydrate de; Procaïn hydrochloridum; Procainii Chloridum; Procainum Chloride; Prokainihydrokloridi; Prokain Hidroklörür; Prokain-hidroklör; Prokain-hydrochlorid; Prokainhydroklorid; Prokaino hidrokloridas; Prokainy chlorowodorek; Syncaine. 2-Diethylaminoethyl 4-aminobenzoate hydrochloride.

Прокаина Гидрохлорид

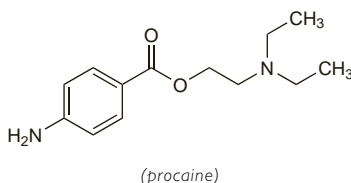
C₁₃H₂₀N₂O₂.HCl = 272.8.

CAS — 59-46-1 (procaïne); 51-05-8 (procaïne hydrochloride).

ATC — C05AD05; N01BA02; S01HA05.

ATC Vet — QC05AD05; QN01BA02; QS01HA05.

The symbol † denotes a preparation no longer actively marketed



Pharmacopoeias. In *Chin., Eur.* (see p.vii), *Int., Jpn, US,* and *Viet.*

Ph. Eur. 6.2 (Procaine Hydrochloride). A white or almost white crystalline powder or colourless crystals. Very soluble in water; soluble in alcohol. A 2% solution in water has a pH of 5.0 to 6.5. Protect from light.

USP 31 (Procaine Hydrochloride). Odourless, small, white crystals or white, crystalline powder. Soluble 1 in 1 of water and 1 in 15 of alcohol; slightly soluble in chloroform; practically insoluble in ether.

Incompatibility. Procaine hydrochloride has been reported to be incompatible with aminophylline, barbiturates, magnesium sulfate, phenytoin sodium, sodium bicarbonate, and amphotericin B.

Stability of solutions. Degradation of procaine in a cardioplegic solution containing magnesium, sodium, potassium, and calcium salts was found to be temperature dependent.¹ At a storage temperature of 6° the shelf-life of the solution was 5 weeks and this was increased to 9 weeks when the storage temperature was -10°. Using carbon dioxide instead of nitrogen in the head space did not affect stability of procaine.

1. Synave R, *et al.* Stability of procaine hydrochloride in a cardioplegic solution containing bicarbonate. *J Clin Hosp Pharm* 1985; **10**: 385-8.

Adverse Effects, Treatment, and Precautions
As for Local Anaesthetics in general, p.1850.

Effects on the cardiovascular system. Severe hypotension leading to cardiac arrest and death developed in a patient following the infusion of 600 mg of procaine for malignant hyperthermia.¹

1. MacLachlan D, Forrest AL. Procaine and malignant hyperthermia. *Lancet* 1974; **i**: 355.

Hypersensitivity. Of 600 persons with dermatitis or eczema submitted to patch testing with 2% aqueous solution of procaine hydrochloride, 4.8% gave a positive reaction.¹

For reports of hypersensitivity including anaphylactic reactions associated with procaine and other local anaesthetics, see under Adverse Effects of Local Anaesthetics, p.1850.

1. Rudzki E, Kleniewska D. The epidemiology of contact dermatitis in Poland. *Br J Dermatol* 1970; **83**: 543-5.

Systemic lupus erythematosus. The limited theoretical risk from using procaine for local anaesthesia in patients who have had procainamide-induced SLE was aired some years ago.¹⁻³

1. Dubois EL. Procaine anesthesia after procainamide-induced systemic erythematosus. *JAMA* 1977; **238**: 2201.
2. Alarcón-Segovia D. Procaine anesthesia after procainamide-induced systemic erythematosus. *JAMA* 1977; **238**: 2201.
3. Lee SL. Procaine anesthesia after procainamide-induced systemic erythematosus. *JAMA* 1977; **238**: 2201.

Interactions

For interactions associated with local anaesthetics, see p.1851.

Diuretics. Use with **acetazolamide** extends the plasma half-life of procaine.¹

1. Calvo R, *et al.* Effects of disease and acetazolamide on procaine hydrolysis by red blood cell enzymes. *Clin Pharmacol Ther* 1980; **27**: 179-83.

Pharmacokinetics

Procaine is poorly absorbed from mucous membranes and is usually given parenterally. It is rapidly hydrolysed by plasma cholinesterase to para-aminobenzoic acid and diethylaminoethanol; some may also be metabolised in the liver. Only about 6% is bound to plasma proteins. About 80% of the para-aminobenzoic acid is excreted unchanged or conjugated in the urine. About 30% of the diethylaminoethanol is excreted in the urine, the remainder being metabolised in the liver.

See also under Local Anaesthetics, p.1852.

Uses and Administration

Procaine hydrochloride, a para-aminobenzoic acid ester, is a local anaesthetic with actions and uses similar to those described on p.1852. Because of its poor penetration of intact mucous membranes, procaine is ineffective for surface application and has been chiefly used by injection, although in general it has been replaced by lidocaïne and other local anaesthetics. It has a slow onset of action and a short duration of action. It has vasodilator activity and therefore a vasoconstrictor may be added to delay absorption and increase the duration of action. Procaine has

mainly been used for infiltration anaesthesia, peripheral nerve blocks, and spinal block. (Local anaesthetic techniques are discussed on p.1853.) It has also been used in cardioplegic solutions to protect the myocardium during cardiac surgery.

For **infiltration anaesthesia** 0.25 or 0.5% solutions of procaine hydrochloride have been used in doses of 350 to 600 mg.

For **peripheral nerve block** a usual dose of 500 mg of procaine hydrochloride has been given as a 0.5% (100 mL), 1% (50 mL), or 2% (25 mL) solution. Doses up to 1 g have been used. For infiltration and peripheral nerve block adrenaline has been added to solutions, in general to give a final concentration of 1 in 200 000 to 1 in 100 000.

Procaine hydrochloride has been used with propoxycaine in dentistry.

Procaine forms poorly soluble salts or conjugates with some drugs, for example penicillin, and is used to prolong their action after injection. It may also reduce the pain of injection.

Procaine-*N*-glucoside hydrochloride has been included in a preparation for gastrointestinal disorders, and procaine ascorbate has been included in a multivitamin preparation.

Action. For a comparison of the vasoactivity of procaine and some other local anaesthetics, see p.1852.

Preparations

USP 31: Procaine and Tetracaine Hydrochlorides and Levonordefrin Injection; Procaine Hydrochloride and Epinephrine Injection; Procaine Hydrochloride Injection; Propoxycaine and Procaine Hydrochlorides and Levonordefrin Injection; Propoxycaine and Procaine Hydrochlorides and Norepinephrine Bitartrate Injection.

Proprietary Preparations (details are given in Part 3)

Arg.: Endocaina†; **Fadacaina;** Procanest; **Austria:** Geroaslan H3; Geroval H3; Novanaest; **Canad.:** Novocain†; **Ger.:** Hewedolor Procaïn; Lophakomp-Procaïn; Novocain†; Pasconeural-Injektions; **Hong Kong:** Geroval H3; **Ital.:** Lenident; **USA:** Novocain; **Venez.:** Artrocel; Bloquet; Genaplex.

Multi-ingredient: **Arg.:** 6 Copin; Dastoni; Gero H3 Aslan†; Gingeron; Muco-Anestyl†; Otalex G; Otonorthia; Sicadental Plus†; **Austral.:** Cardioplegia Concentrate; **Austria:** Aslavit†; Causat; Gerontin; KH3; Regenerin†; **Braz.:** Afine; Algident†; Bismu-Jet; Claudemor; Colutoide; Dentisan; Fongergin; Otobell†; Otoloide; Oturga; Passaja†; Pradente†; Timpanol†; Usedent†; **Chile:** Betonovit†; Diltotal; KH3-Vit†; KH3†; Megavit†; Pantiban; **Cz.:** Solutan†; Solutio Thomas cum Procaino; **Denm.:** Cardioplex; **Ger.:** Bismolan N†; Cardioplegic N†; Gero H3 Aslan; Hewedolor plus Coffein; KH3†; NeyPulpin N (Revitorgan-Dilution N Nr 10)†; Otalgan; Polyamin†; Procaneural†; Revicain comp plus†; Revicain comp†; Revicain†; Veno-Kattwiga N†; **Gr.:** Cardioplegia; **Hong Kong:** Cardioplegia; KH3; **Hung.:** Hemorid; Noditrin†; Trypsin†; **Indon.:** Cardioplegia; **Israel:** Bedodeka Antineuralgic†; **Ital.:** Dentosedina; Ginvapast; Mios; Neo-Ustol; Otalgan; Otomidone; Otopax; Rinantipol†; Ustiosan; **Malaysia:** Cardioplegia; **NZ:** KH3; **Port.:** Claudemor†; Otocalma†; **Rus.:** Solutan (Covyrat); **S.Afr.:** Salusa†; Universal Earache Drops; **Singapore:** Cardioplegia; **Spain:** Anestesia Loc Braun S/A; Co Bucal; Colirioclina Adren Astr; Dentol Topico; Eupnol; Kanafosal; Kanafosal Predni; Neocolan; Nulacin Fermentos; Oftalmol Ocular; Otalgan†; Otosedol; Tangenol†; **Switz.:** Anaestalgin; Ginvapast; Otalgan; Otosan; **Thai.:** Cardioplegia; KH3.

Used as an adjunct in: **Arg.:** Betametasona B12; **Braz.:** Cianotrat-Dexa; Dexa-Neurib†; Dexacab†; Dexador; Dexagil; Dexaneurin; **Ger.:** Eukalisan N; **Ital.:** Neurofal†; **Malaysia:** Alinamin B12†; **Singapore:** Alinamin B12†; **Spain:** Sulmetin Papaverina†; Sulmetin†; **USA:** Hytunic†.

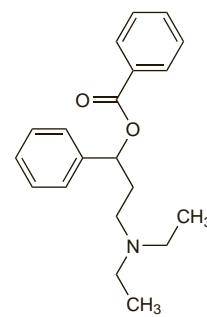
Propanocaine Hydrochloride (rINN)

467D3; Hidrocloruro de propanocaína; Propanocaïne, Chlorhydrate de; Propanocaini Hydrochloridum. 3-Diethylamino-1-phenylpropyl benzoate hydrochloride.

Пропанокaina Гидрохлорид

C₂₀H₂₅NO₂.HCl = 347.9.

CAS — 493-76-5 (propanocaine); 1679-79-4 (propanocaine hydrochloride).



(propanocaine)

Profile

Propanocaine hydrochloride, a benzoic acid ester, is a local anaesthetic (p.1850) that has been used topically for surface anaesthesia.

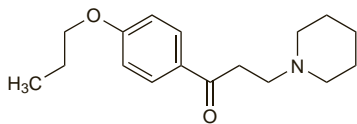
Preparations**Proprietary Preparations** (details are given in Part 3)**Multi-ingredient:** **Spain:** Detraïne.**Propipocaine** (*rINN*)

Propipocaina; Propipocaine; Propipocainum; Propoxypropipocaine. 3-Piperidino-4'-propoxypropiphenone.

Пропипокаин

 $C_{17}H_{25}NO_2 = 275.4$.

CAS — 3670-68-6.

**Profile**

Propipocaine is a local anaesthetic (p.1850) that has been used for surface anaesthesia.

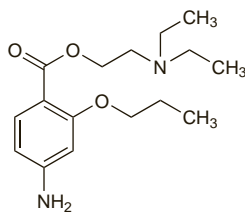
Propoxycaine Hydrochloride (*rINN*)

Hidrocloruro de propoxicaína; Propoxycaine, Chlorhydrate de; Propoxycaini Hydrochloridum; Propoxycainium Chloride. 2-Diethylaminoethyl 4-amino-2-propoxybenzoate hydrochloride.

Пропоксикаина Гидрохлорид

 $C_{16}H_{26}N_2O_3 \cdot HCl = 330.9$.

CAS — 86-43-1 (propoxycaine); 550-83-4 (propoxycaine hydrochloride).



(propoxycaine)

Pharmacopoeias. In *US*.**USP 31** (Propoxycaine Hydrochloride). A white odourless crystalline solid. It discolours on prolonged exposure to light and air. Soluble 1 in 2 of water, 1 in 10 of alcohol, and 1 in 80 of ether; practically insoluble in acetone and in chloroform. A 2% solution in water has a pH of about 5.4. Protect from light.**Profile**

Propoxycaine hydrochloride, a para-aminobenzoic acid ester, is a local anaesthetic (p.1850). It has been used with procaine hydrochloride and a vasoconstrictor for infiltration anaesthesia and nerve block in dental procedures. Propoxycaine has a more rapid onset and a longer duration of action than that of procaine.

Preparations**USP 31:** Propoxycaine and Procaine Hydrochlorides and Levonordefrin Injection; Propoxycaine and Procaine Hydrochlorides and Norepinephrine Bitartrate Injection.**Proxymetacaine Hydrochloride**(BANM, *rINN*)

Hidrocloruro de proximetacaína; Proksimetakain Hidroklorür; Proparacaine Hydrochloride; Proparakain Hidroklorür; Proxymetacaine, Chlorhydrate de; Proxymetacaini Hydrochloridum. 2-Diethylaminoethyl 3-amino-4-propoxybenzoate hydrochloride.

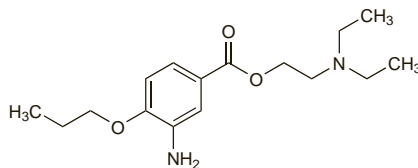
Проксиметакаина Гидрохлорид

 $C_{16}H_{26}N_2O_3 \cdot HCl = 330.9$.

CAS — 499-67-2 (proxymetacaine); 5875-06-9 (proxymetacaine hydrochloride).

ATC — S01HA04.

ATC Vet — QS01HA04.



(proxymetacaine)

NOTE. PROX is a code approved by the BP 2008 for use on single unit doses of eye drops containing proxymetacaine hydrochloride where the individual container may be too small to bear all the appropriate labelling information. PROXFLN is a similar code approved for eye drops containing proxymetacaine hydrochloride and fluorescein sodium.

Pharmacopoeias. In *Br* and *US*.**BP 2008** (Proxymetacaine Hydrochloride). A white or almost white, odourless or almost odourless, crystalline powder. Soluble in water and in chloroform; very soluble in dehydrated alcohol; practically insoluble in ether. A 1% solution in water has a pH of 5.7 to 6.4. Protect from light.**USP 31** (Proparacaine Hydrochloride). A white to off-white, or faintly buff-coloured, odourless, crystalline powder. Soluble in water, in warm alcohol, and in methyl alcohol; insoluble in ether and in benzene.**Adverse Effects, Treatment, and Precautions**

As for Local Anaesthetics in general, p.1850.

A severe immediate-type corneal reaction to proxymetacaine may rarely occur. Allergic contact dermatitis has also been reported.

Effects on the skin. Exacerbation of Stevens-Johnson syndrome has been reported¹ in a woman after ophthalmic anaesthesia with proxymetacaine hydrochloride.1. Ward B, *et al.* Dermatologic reaction in Stevens-Johnson syndrome after ophthalmic anesthesia with proparacaine hydrochloride. *Am J Ophthalmol* 1978; **86**: 133-5.**Interactions**

For interactions associated with local anaesthetics, see p.1851.

Pharmacokinetics

See under Local Anaesthetics, p.1852.

Uses and Administration

Proxymetacaine hydrochloride, a meta-aminobenzoic acid ester, is a local anaesthetic with actions and uses similar to those described on p.1852. It is used for surface anaesthesia (p.1853) in ophthalmology in a concentration of 0.5%. Proxymetacaine is of similar potency to tetracaine in equal concentrations and induces anaesthesia within about 20 seconds. The duration of action may be 15 minutes or longer. Instillation of 1 or 2 drops permits tonometry after 30 seconds. For removal of foreign bodies or sutures from the cornea 1 or 2 drops are instilled every 5 to 10 minutes for up to 3 applications, or 1 or 2 drops are instilled 2 to 3 minutes before the procedure. For deeper anaesthesia such as needed for cataract extraction 1 drop is instilled every 5 to 10 minutes to a total of 5 to 7 applications.

Trigeminal neuralgia. There have been anecdotal reports that proxymetacaine eye drops relieved trigeminal neuralgia (p.9) refractory to carbamazepine.^{1,2} However, a controlled study failed to demonstrate any benefit.³

1. Zvon MR, Fichte CM. Trigeminal neuralgia relieved by ophthalmic anesthetic. *JAMA* 1991; **265**: 2807.
2. Zvon MR, Fichte CM. Trigeminal neuralgia relieved by optical anesthesia. *JAMA* 1991; **266**: 1649.
3. Kondziolka D, *et al.* The effect of single-application topical ophthalmic anesthesia in patients with trigeminal neuralgia: a randomized double-blind placebo-controlled trial. *J Neurosurg* 1994; **80**: 993-7.

Preparations**BP 2008:** Proxymetacaine Eye Drops;**USP 31:** Fluorescein Sodium and Proparacaine Hydrochloride Ophthalmic Solution; Proparacaine Hydrochloride Ophthalmic Solution.**Proprietary Preparations** (details are given in Part 3)**Arg.:** Anestalcon; Poencaina; **Austral.:** Alcaine; Ophthetici; **Belg.:** Alcaine; **Braz.:** Anestalcon; Visonest; **Canad.:** Ak-Taine; Alcaine; Diocaine; Ophthetici; **Chile:** Anestalcon; **Ger.:** Proparakain-POS; **Gr.:** Alcaine; **Hong Kong:** Alcaine; **Malaysia:** Alcaine; **Mex.:** Alcaine; **Norw.:** Alcaine; **NZ:** Ophthetici; **Philipp.:** Alcaine; **Pol.:** Alcaine; **Rus.:** Alcaine (Алкаин); **Singap.****pore:** Alcaine; **Switz.:** Alcaine; **Turk.:** Alcaine; Opticaine; **USA:** Ak-Taine; Alcaine; Ocu-Caine; Ophthetici; Paracaine; **Venez.:** Alcaine; Oftaine; Poencaina.**Multi-ingredient:** **Canad.:** Fluoracaine; **USA:** Fluoracaine; Fluorocaine.**Quinisocaine Hydrochloride** (BANM, *rINN*)Chinisocainum Hydrochloride; Dimethisoquin Hydrochloride (*USAN*); Dimethisoquinium Chloride; Hidrocloruro de quinisocaina; Quinisocaine, Chlorhydrate de; Quinisocaini Hydrochloridum. 2-(3-Butyl-1-isquinolinyloxy)-*NN*-dimethylethylamine hydrochloride.

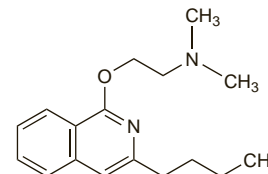
Хинизокаина Гидрохлорид

 $C_{17}H_{24}N_2O \cdot HCl = 308.8$.

CAS — 86-80-6 (quinisocaine); 2773-92-4 (quinisocaine hydrochloride).

ATC — D04AB05.

ATC Vet — QD04AB05.



(quinisocaine)

Profile

Quinisocaine hydrochloride is a local anaesthetic (p.1850) that has been used as a surface anaesthetic in the form of an ointment or cream in a concentration of 0.5% for the relief of pruritus, anogenital or anorectal irritation, and minor skin conditions. It has also been used as suppositories.

Preparations**Proprietary Preparations** (details are given in Part 3)**Fr.:** Quotane; **Ger.:** Haenal; Isochiniol; **Switz.:** Isochiniol.**Multi-ingredient:** **Fr.:** Rectoquotane.**Ropivacaine Hydrochloride**(BANM, *rINN*)

AL-281; Hidrocloruro de ropivacaína; Ropivacaine, chlorhydrate de; Ropivacaini hydrochloridum; Ropivakainihydroklorid; Ropivakain Hidroklorür; Ropivakainhydroklorid. (S)-2',6'-Dimethyl-1-propylpiperidine-2-carboxanilide hydrochloride monohydrate.

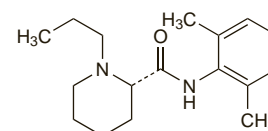
Ропивакаина Гидрохлорид

 $C_{17}H_{26}N_2O \cdot HCl \cdot H_2O = 328.9$.

CAS — 84057-95-4 (ropivacaine); 98717-15-8 (anhydrous ropivacaine hydrochloride); 132112-35-7 (ropivacaine hydrochloride monohydrate).

ATC — N01BB09.

ATC Vet — QN01BB09.



(ropivacaine)

Pharmacopoeias. In *Eur*; (see p.vii) and *US*.**Ph. Eur. 6.2** (Ropivacaine Hydrochloride Monohydrate). A white or almost white, crystalline powder. Soluble in water and in alcohol; slightly soluble in dichloromethane. pH of a 2% solution in water is 4.5 to 6.0.**USP 31** (Ropivacaine Hydrochloride). A white crystalline powder. Soluble in water. A 1% solution in water has a pH of 4.5 to 6.0.**Adverse Effects, Treatment, and Precautions**

As for Local Anaesthetics in general, p.1850.

Ropivacaine is contra-indicated for use in intravenous regional anaesthesia (Bier's block) and for paracervical block in obstetrics.

Effects on the cardiovascular system. Ropivacaine is structurally related to bupivacaine, but data from extensive animal studies suggest that ropivacaine may be less cardiotoxic than bupivacaine.¹ Results from a study² in 12 healthy male volunteers support these data; at doses producing CNS symptoms car-