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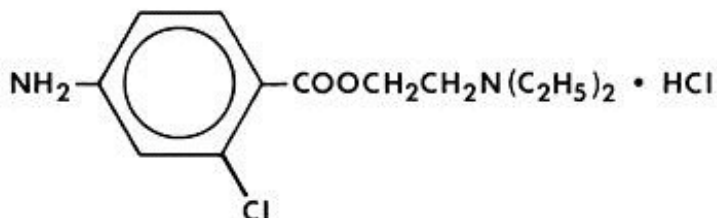
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4 **Nesacaine[®]** (chloroprocaine HCl Injection, USP)

5 **Nesacaine[®]-MPF** (chloroprocaine HCl Injection, USP)

6 *For Infiltration and Nerve Block*

7
8 **DESCRIPTION:**

9 Nesacaine and Nesacaine-MPF Injections are sterile non pyrogenic local anesthetics. The
10 active ingredient in Nesacaine and Nesacaine-MPF Injections is chloroprocaine HCl
11 (benzoic acid, 4-amino-2-chloro-2-(diethylamino) ethyl ester, monohydrochloride),
12 which is represented by the following structural formula:



13
14 Table 1: Composition of Available Injections

Formula (mg/mL)				
Product Identification	Chloro-procaine HCl	Sodium Chloride	Disodium EDTA dihydrate	Methylparaben
Nesacaine 1%	10	6.7	0.111	1
Nesacaine 2%	20	4.7	0.111	1
Nesacaine-MPF 2%	20	4.7	-	-
Nesacaine-MPF 3%	30	3.3	-	-

15
16 The solutions are adjusted to pH 2.7–4.0 by means of sodium hydroxide and/or

1 hydrochloric acid. Filled under nitrogen. Nesacaine and Nesacaine-MPF Injections
2 should not be resterilized by autoclaving.

3 **CLINICAL PHARMACOLOGY:**

4 Chloroprocaine, like other local anesthetics, blocks the generation and the conduction of
5 nerve impulses, presumably by increasing the threshold for electrical excitation in the
6 nerve, by slowing the propagation of the nerve impulse and by reducing the rate of rise of
7 the action potential. In general, the progression of anesthesia is related to the diameter,
8 myelination and conduction velocity of affected nerve fibers. Clinically, the order of loss
9 of nerve function is as follows: (1) pain, (2) temperature, (3) touch, (4) proprioception,
10 and (5) skeletal muscle tone.

11 Systemic absorption of local anesthetics produces effects on the cardiovascular
12 and central nervous systems. At blood concentrations achieved with normal therapeutic
13 doses, changes in cardiac conduction, excitability, refractoriness, contractility, and
14 peripheral vascular resistance are minimal. However, toxic blood concentrations depress
15 cardiac conduction and excitability, which may lead to atrioventricular block and
16 ultimately to cardiac arrest. In addition, with toxic blood concentrations myocardial
17 contractility may be depressed and peripheral vasodilation may occur, leading to
18 decreased cardiac output and arterial blood pressure.

19 Following systemic absorption, toxic blood concentrations of local anesthetics can
20 produce central nervous system stimulation, depression, or both. Apparent central
21 stimulation may be manifested as restlessness, tremors and shivering, which may
22 progress to convulsions. Depression and coma may occur, possibly progressing
23 ultimately to respiratory arrest.

1 However, the local anesthetics have a primary depressant effect on the medulla
2 and on higher centers. The depressed stage may occur without a prior stage of central
3 nervous system stimulation.

4 **PHARMACOKINETICS:**

5 The rate of systemic absorption of local anesthetic drugs is dependent upon the total dose
6 and concentration of drug administered, the route of administration, the vascularity of the
7 administration site, and the presence or absence of epinephrine in the anesthetic injection.
8 Epinephrine usually reduces the rate of absorption and plasma concentration of local
9 anesthetics and is sometimes added to local anesthetic injections in order to prolong the
10 duration of action.

11 The onset of action with chlorprocaine is rapid (usually within 6 to 12 minutes),
12 and the duration of anesthesia, depending upon the amount used and the route of
13 administration, may be up to 60 minutes.

14 Local anesthetics appear to cross the placenta by passive diffusion. However, the
15 rate and degree of diffusion varies considerably among the different drugs as governed
16 by: (1) the degree of plasma protein binding, (2) the degree of ionization, and (3) the
17 degree of lipid solubility. Fetal/maternal ratios of local anesthetics appear to be inversely
18 related to the degree of plasma protein binding, since only the free, unbound drug is
19 available for placental transfer. Thus, drugs with the highest protein binding capacity
20 may have the lowest fetal/maternal ratios. The extent of placental transfer is also
21 determined by the degree of ionization and lipid solubility of the drug. Lipid soluble,
22 nonionized drugs readily enter the fetal blood from the maternal circulation.

1 Depending upon the route of administration, local anesthetics are distributed to
2 some extent to all body tissues, with high concentrations found in highly perfused organs
3 such as the liver, lungs, heart, and brain.

4 Various pharmacokinetic parameters of the local anesthetics can be significantly
5 altered by the presence of hepatic or renal disease, addition of epinephrine, factors
6 affecting urinary pH, renal blood flow, the route of administration, and the age of the
7 patient. The *in vitro* plasma half-life of chloroprocaine in adults is 21 ± 2 seconds for
8 males and 25 ± 1 seconds for females. The *in vitro* plasma half-life in neonates is 43 ± 2
9 seconds.

10 Chloroprocaine is rapidly metabolized in plasma by hydrolysis of the ester
11 linkage by pseudocholinesterase. The hydrolysis of chloroprocaine results in the
12 production of β -diethylaminoethanol and 2-chloro-4-aminobenzoic acid, which inhibits
13 the action of the sulfonamides (see **PRECAUTIONS**).

14 The kidney is the main excretory organ for most local anesthetics and their
15 metabolites. Urinary excretion is affected by urinary perfusion and factors affecting
16 urinary pH.

17 **INDICATIONS AND USAGE:**

18 Nesacaine 1% and 2% Injections, in multidose vials with methylparaben as preservative,
19 are indicated for the production of local anesthesia by infiltration and peripheral nerve
20 block. They are not to be used for lumbar or caudal epidural anesthesia.

21 Nesacaine-MPF 2% and 3% Injections, in single dose vials without preservative
22 and without EDTA, are indicated for the production of local anesthesia by infiltration,
23 peripheral and central nerve block, including lumbar and caudal epidural blocks.

1 Nesacaine and Nesacaine-MPF Injections are not to be used for subarachnoid
2 administration.

3 **CONTRAINDICATIONS:**

4 Nesacaine and Nesacaine-MPF Injections are contraindicated in patients hypersensitive
5 (allergic) to drugs of the PABA ester group.

6 Lumbar and caudal epidural anesthesia should be used with extreme caution in
7 persons with the following conditions: existing neurological disease, spinal deformities,
8 septicemia, and severe hypertension.

9 **WARNINGS:**

10 LOCAL ANESTHETICS SHOULD ONLY BE EMPLOYED BY CLINICIANS WHO
11 ARE WELL VERSED IN DIAGNOSIS AND MANAGEMENT OF DOSE RELATED
12 TOXICITY AND OTHER ACUTE EMERGENCIES WHICH MIGHT ARISE FROM
13 THE BLOCK TO BE EMPLOYED, AND THEN ONLY AFTER ENSURING THE
14 IMMEDIATE AVAILABILITY OF OXYGEN, OTHER RESUSCITATIVE DRUGS,
15 CARDIOPULMONARY RESUSCITATIVE EQUIPMENT, AND THE PERSONNEL
16 RESOURCES NEEDED FOR PROPER MANAGEMENT OF TOXIC REACTIONS
17 AND RELATED EMERGENCIES (see also **ADVERSE REACTIONS** and
18 **PRECAUTIONS**). DELAY IN PROPER MANAGEMENT OF DOSE RELATED
19 TOXICITY, UNDERVENTILATION FROM ANY CAUSE AND/OR ALTERED
20 SENSITIVITY MAY LEAD TO THE DEVELOPMENT OF ACIDOSIS, CARDIAC
21 ARREST AND, POSSIBLY, DEATH. NESACAINE (chloroprocaine HCl Injection,
22 USP) contains methylparaben and should not be used for lumbar or caudal epidural
23 anesthesia because safety of this antimicrobial preservative has not been established with

1 regard to intrathecal injection, either intentional or unintentional. NESACAINE-MPF
2 Injection contains no preservative; discard unused injection remaining in vial after initial
3 use.

4 Intra-articular infusions of local anesthetics following arthroscopic and other
5 surgical procedures is an unapproved use, and there have been post-marketing reports of
6 chondrolysis in patients receiving such infusions. The majority of reported cases of
7 chondrolysis have involved the shoulder joint; cases of gleno-humeral chondrolysis have
8 been described in pediatric and adult patients following intra-articular infusions of local
9 anesthetics with and without epinephrine for periods of 48 to 72 hours. There is
10 insufficient information to determine whether shorter infusion periods are not associated
11 with these findings. The time of onset of symptoms, such as joint pain, stiffness and loss
12 of motion can be variable, but may begin as early as the 2nd month after surgery.
13 Currently, there is no effective treatment for chondrolysis; patients who experienced
14 chondrolysis have required additional diagnostic and therapeutic procedures and some
15 required arthroplasty or shoulder replacement.

16 Vasopressors should not be used in the presence of ergot-type oxytocic drugs,
17 since a severe persistent hypertension may occur.

18 To avoid intravascular injection, aspiration should be performed before the
19 anesthetic solution is injected. The needle must be repositioned until no blood return can
20 be elicited. However, the absence of blood in the syringe does not guarantee that
21 intravascular injection has been avoided.

22 Mixtures of local anesthetics are sometimes employed to compensate for the
23 slower onset of one drug and the shorter duration of action of the second drug.

1 Experiments in primates suggest that toxicity is probably additive when mixtures of local
2 anesthetics are employed, but some experiments in rodents suggest synergism. Caution
3 regarding toxic equivalence should be exercised when mixtures of local anesthetics are
4 employed.

5 **PRECAUTIONS:**

6 *General*

7 The safety and effective use of chlorprocaine depend on proper dosage, correct
8 technique, adequate precautions and readiness for emergencies. Resuscitative equipment,
9 oxygen and other resuscitative drugs should be available for immediate use (see
10 **WARNINGS** and **ADVERSE REACTIONS**). The lowest dosage that results in
11 effective anesthesia should be used to avoid high plasma levels and serious adverse
12 effects. Injections should be made slowly, with frequent aspirations before and during
13 the injection to avoid intravascular injection. Syringe aspirations should also be
14 performed before and during each supplemental injection in continuous (intermittent)
15 catheter techniques. During the administration of epidural anesthesia, it is recommended
16 that a test dose be administered (3 mL of 3% or 5 mL of 2% Nesacaine-MPF Injection)
17 initially and that the patient be monitored for central nervous system toxicity and
18 cardiovascular toxicity, as well as for signs of unintended intrathecal administration,
19 before proceeding. When clinical conditions permit, consideration should be given to
20 employing a chlorprocaine solution that contains epinephrine for the test dose because
21 circulatory changes characteristic of epinephrine may also serve as a warning sign of
22 unintended intravascular injection. An intravascular injection is still possible even if
23 aspirations for blood are negative. With the use of continuous catheter techniques, it is

1 recommended that a fraction of each supplemental dose be administered as a test dose in
2 order to verify proper location of the catheter.

3 Injection of repeated doses of local anesthetics may cause significant increases in
4 plasma levels with each repeated dose due to slow accumulation of the drug or its
5 metabolites. Tolerance to elevated blood levels varies with the physical condition of the
6 patient. Debilitated, elderly patients, acutely ill patients, and children should be given
7 reduced doses commensurate with their age and physical status. Local anesthetics should
8 also be used with caution in patients with hypotension or heart block.

9 Careful and constant monitoring of cardiovascular and respiratory (adequacy of
10 ventilation) vital signs and the patient's state of consciousness should be accomplished
11 after each local anesthetic injection. It should be kept in mind at such times that
12 restlessness, anxiety, tinnitus, dizziness, blurred vision, tremors, depression or drowsiness
13 may be early warning signs of central nervous system toxicity.

14 Local anesthetic injections containing a vasoconstrictor should be used cautiously
15 and in carefully circumscribed quantities in areas of the body supplied by end arteries or
16 having otherwise compromised blood supply. Patients with peripheral vascular disease
17 and those with hypertensive vascular disease may exhibit exaggerated vasoconstrictor
18 response. Ischemic injury or necrosis may result.

19 Since ester-type local anesthetics are hydrolyzed by plasma cholinesterase
20 produced by the liver, chlorprocaine should be used cautiously in patients with hepatic
21 disease. Local anesthetics should also be used with caution in patients with impaired
22 cardiovascular function since they may be less able to compensate for functional changes
23 associated with the prolongation of A-V conduction produced by these drugs.

1 **Use in Ophthalmic Surgery:** When local anesthetic injections are employed for
2 retrobulbar block, lack of corneal sensation should not be relied upon to determine
3 whether or not the patient is ready for surgery. This is because complete lack of corneal
4 sensation usually precedes clinically acceptable external ocular muscle akinesia.

5 ***Information for Patients***

6 When appropriate, patients should be informed in advance that they may experience
7 temporary loss of sensation and motor activity, usually in the lower half of the body,
8 following proper administration of epidural anesthesia.

9 ***Clinically Significant Drug Interactions***

10 The administration of local anesthetic solutions containing epinephrine or norepinephrine
11 to patients receiving monoamine oxidase inhibitors, tricyclic antidepressants or
12 phenothiazines may produce severe, prolonged hypotension or hypertension. Concurrent
13 use of these agents should generally be avoided. In situations when concurrent therapy is
14 necessary, careful patient monitoring is essential.

15 Concurrent administration of vasopressor drugs (for the treatment of hypotension
16 related to obstetric blocks) and ergot-type oxytocic drugs may cause severe, persistent
17 hypertension or cerebrovascular accidents.

18 The para-aminobenzoic acid metabolite of chlorprocaine inhibits the action of
19 sulfonamides. Therefore, chlorprocaine should not be used in any condition in which a
20 sulfonamide drug is being employed.

21 ***Carcinogenesis, Mutagenesis, and Impairment of Fertility***

22 Long-term studies in animals to evaluate carcinogenic potential and reproduction studies
23 to evaluate mutagenesis or impairment of fertility have not been conducted with

1 chloroprocaine.

2 ***Pregnancy: Category C***

3 Animal reproduction studies have not been conducted with chloroprocaine. It is also not
4 known whether chloroprocaine can cause fetal harm when administered to a pregnant
5 woman or can affect reproduction capacity. Chloroprocaine should be given to a
6 pregnant woman only if clearly needed. This does not preclude the use of chloroprocaine
7 at term for the production of obstetrical anesthesia.

8 ***Labor and Delivery***

9 Local anesthetics rapidly cross the placenta, and when used for epidural, paracervical,
10 pudendal or caudal block anesthesia, can cause varying degrees of maternal, fetal and
11 neonatal toxicity (see **CLINICAL PHARMACOLOGY** and
12 **PHARMACOKINETICS**).

13 The incidence and degree of toxicity depend upon the procedure performed, the
14 type and amount of drug used, and the technique of drug administration. Adverse
15 reactions in the parturient, fetus and neonate involve alterations of the central nervous
16 system, peripheral vascular tone and cardiac function.

17 Maternal hypotension has resulted from regional anesthesia. Local anesthetics
18 produce vasodilation by blocking sympathetic nerves. Elevating the patient's legs and
19 positioning her on her left side will help prevent decreases in blood pressure. The fetal
20 heart rate also should be monitored continuously, and electronic fetal monitoring is
21 highly advisable.

22 Epidural, paracervical, or pudendal anesthesia may alter the forces of parturition
23 through changes in uterine contractility or maternal expulsive efforts. In one study,

1 paracervical block anesthesia was associated with a decrease in the mean duration of first
2 stage labor and facilitation of cervical dilation. However, epidural anesthesia has also
3 been reported to prolong the second stage of labor by removing the parturient's reflex
4 urge to bear down or by interfering with motor function. The use of obstetrical
5 anesthesia may increase the need for forceps assistance.

6 The use of some local anesthetic drug products during labor and delivery may be
7 followed by diminished muscle strength and tone for the first day or two of life. The
8 long-term significance of these observations is unknown.

9 Careful adherence to recommended dosage is of the utmost importance in
10 obstetrical paracervical block. Failure to achieve adequate analgesia with recommended
11 doses should arouse suspicion of intravascular or fetal intracranial injection. Cases
12 compatible with unintended fetal intracranial injection of local anesthetic injection have
13 been reported following intended paracervical or pudendal block or both. Babies so
14 affected present with unexplained neonatal depression at birth which correlates with high
15 local anesthetic serum levels and usually manifest seizures within six hours. Prompt use
16 of supportive measures combined with forced urinary excretion of the local anesthetic has
17 been used successfully to manage this complication.

18 Case reports of maternal convulsions and cardiovascular collapse following use of
19 some local anesthetics for paracervical block in early pregnancy (as anesthesia for
20 elective abortion) suggest that systemic absorption under these circumstances may be
21 rapid. The recommended maximum dose of each drug should not be exceeded. Injection
22 should be made slowly and with frequent aspiration. Allow a 5-minute interval between
23 sides.

1 There are no data concerning use of chloroprocaine for obstetrical paracervical
2 block when toxemia of pregnancy is present or when fetal distress or prematurity is
3 anticipated in advance of the block; such use is, therefore, not recommended.

4 The following information should be considered by clinicians who select
5 chloroprocaine for obstetrical paracervical block anesthesia:

- 6 1. Fetal bradycardia (generally a heart rate of less than 120 per minute for more than 2
7 minutes) has been noted by electronic monitoring in about 5 to 10 percent of the cases
8 (various studies) where initial total doses of 120 mg to 400 mg of chloroprocaine were
9 employed. The incidence of bradycardia, within this dose range, might not be dose
10 related.
- 11 2. Fetal acidosis has not been demonstrated by blood gas monitoring around the time of
12 bradycardia or afterwards. These data are limited and generally restricted to non-
13 toxemic cases where fetal distress or prematurity was not anticipated in advance of the
14 block.
- 15 3. No intact chloroprocaine and only trace quantities of a hydrolysis product, 2-chloro-4-
16 aminobenzoic acid, have been demonstrated in umbilical cord arterial or venous
17 plasma following properly administered paracervical block with chloroprocaine.
- 18 4. The role of drug factors and non-drug factors associated with fetal bradycardia
19 following paracervical block are unexplained at this time.

20 ***Nursing Mothers***

21 It is not known whether this drug is excreted in human milk. Because many drugs are
22 excreted in human milk, caution should be exercised when chloroprocaine is
23 administered to a nursing woman.

1 *Pediatric Use*

2 Guidelines for the administration of Nesacaine and Nesacaine-MPF Injections to children
3 are presented in **DOSAGE AND ADMINISTRATION**.

4 *Geriatric Use*

5 Clinical studies of Nesacaine and Nesacaine-MPF did not include sufficient numbers of
6 subjects 65 and over to determine whether they respond differently from younger
7 subjects. Other reported clinical experience has not identified differences in responses
8 between the elderly and younger patients. In general, dose selection for an elderly patient
9 should be cautious usually starting at the low end of the dosing range, reflecting the
10 greater frequency of decreased hepatic, renal, or cardiac function, and of concomitant
11 disease or other drug therapy.

12 This drug and its metabolites are known to be substantially excreted by the
13 kidney, and the risk of toxic reactions to this drug may be greater in patients with
14 impaired renal function. Because elderly patients are more likely to have decreased renal
15 function, care should be taken in dose selection and it may be useful to monitor renal
16 function.

17 **ADVERSE REACTIONS:**

18 **Systemic:** The most commonly encountered acute adverse experiences that demand
19 immediate countermeasures are related to the central nervous system and the
20 cardiovascular system. These adverse experiences are generally dose related and may
21 result from rapid absorption from the injection site, diminished tolerance, or from
22 unintentional intravascular injection of the local anesthetic solution. In addition to
23 systemic dose-related toxicity, unintentional subarachnoid injection of drug during the

1 intended performance of caudal or lumbar epidural block or nerve blocks near the
2 vertebral column (especially in the head and neck region) may result in underventilation
3 or apnea (“Total Spinal”). Factors influencing plasma protein binding, such as acidosis,
4 systemic diseases that alter protein production, or competition of other drugs for protein
5 binding sites, may diminish individual tolerance. Plasma cholinesterase deficiency may
6 also account for diminished tolerance to ester-type local anesthetics.

7 **Central Nervous System Reactions:** These are characterized by excitation and/or
8 depression. Restlessness, anxiety, dizziness, tinnitus, blurred vision or tremors may
9 occur, possibly proceeding to convulsions. However, excitement may be transient or
10 absent, with depression being the first manifestation of an adverse reaction. This may
11 quickly be followed by drowsiness merging into unconsciousness and respiratory arrest.

12 The incidence of convulsions associated with the use of local anesthetics varies
13 with the procedure used and the total dose administered. In a survey of studies of
14 epidural anesthesia, overt toxicity progressing to convulsions occurred in approximately
15 0.1 percent of local anesthetic administrations.

16 **Cardiovascular System Reactions:** High doses, or unintended intravascular injection,
17 may lead to high plasma levels and related depression of the myocardium, hypotension,
18 bradycardia, ventricular arrhythmias, and, possibly, cardiac arrest.

19 **Allergic:** Allergic type reactions are rare and may occur as a result of sensitivity to the
20 local anesthetic or to other formulation ingredients, such as the antimicrobial preservative
21 methylparaben, contained in multiple dose vials. These reactions are characterized by
22 signs such as urticaria, pruritus, erythema, angioneurotic edema (including laryngeal
23 edema), tachycardia, sneezing, nausea, vomiting, dizziness, syncope, excessive sweating,

1 elevated temperature, and possibly, anaphylactoid type symptomatology (including
2 severe hypotension). Cross sensitivity among members of the ester-type local anesthetic
3 group has been reported. The usefulness of screening for sensitivity has not been
4 definitely established.

5 **Neurologic:** In the practice of caudal or lumbar epidural block, occasional unintentional
6 penetration of the subarachnoid space by the catheter may occur (see **PRECAUTIONS**).
7 Subsequent adverse observations may depend partially on the amount of drug
8 administered intrathecally. These observations may include spinal block of varying
9 magnitude (including total spinal block), hypotension secondary to spinal block, loss of
10 bladder and bowel control, and loss of perineal sensation and sexual function.
11 Arachnoiditis, persistent motor, sensory and/or autonomic (sphincter control) deficit of
12 some lower spinal segments with slow recovery (several months) or incomplete recovery
13 have been reported in rare instances (see **DOSAGE AND ADMINISTRATION**
14 discussion of Caudal and Lumbar Epidural Block). Backache and headache have also
15 been noted following lumbar epidural or caudal block.

16 **OVERDOSAGE:**

17 Acute emergencies from local anesthetics are generally related to high plasma levels
18 encountered during therapeutic use of local anesthetics or to unintended subarachnoid
19 injection of local anesthetic solution (see **ADVERSE REACTIONS, WARNINGS** and
20 **PRECAUTIONS**).

21 In mice, the intravenous LD₅₀ of chlorprocaine HCl is 97 mg/kg and the
22 subcutaneous LD₅₀ of chlorprocaine HCl is 950 mg/kg.

23 **Management of Local Anesthetic Emergencies:** The first consideration is prevention,

1 best accomplished by careful and constant monitoring of cardiovascular and respiratory
2 vital signs and the patient's state of consciousness after each local anesthetic injection.
3 At the first sign of change, oxygen should be administered.

4 The first step in the management of convulsions, as well as underventilation or
5 apnea due to unintentional subarachnoid injection of drug solution, consists of immediate
6 attention to the maintenance of a patent airway and assisted or controlled ventilation with
7 oxygen and a delivery system capable of permitting immediate positive airway pressure
8 by mask. Immediately after the institution of these ventilatory measures, the adequacy of
9 the circulation should be evaluated, keeping in mind that drugs used to treat convulsions
10 sometimes depress the circulation when administered intravenously. Should convulsions
11 persist despite adequate respiratory support, and if the status of the circulation permits,
12 small increments of an ultra-short acting barbiturate (such as thiopental or thiamylal) or a
13 benzodiazepine (such as diazepam) may be administered intravenously; the clinician
14 should be familiar, prior to the use of anesthetics, with these anticonvulsant drugs.
15 Supportive treatment of circulatory depression may require administration of intravenous
16 fluids and, when appropriate, a vasopressor dictated by the clinical situation (such as
17 ephedrine to enhance myocardial contractile force).

18 If not treated immediately, both convulsions and cardiovascular depression can
19 result in hypoxia, acidosis, bradycardia, arrhythmias and cardiac arrest. Underventilation
20 or apnea due to unintentional subarachnoid injection of local anesthetic solution may
21 produce these same signs and also lead to cardiac arrest if ventilatory support is not
22 instituted. If cardiac arrest should occur, standard cardiopulmonary resuscitative
23 measures should be instituted. Recovery has been reported after prolonged resuscitative

1 efforts.

2 Endotracheal intubation, employing drugs and techniques familiar to the clinician,
3 may be indicated, after initial administration of oxygen by mask, if difficulty is
4 encountered in the maintenance of a patent airway or if prolonged ventilatory support
5 (assisted or controlled) is indicated.

6 **DOSAGE AND ADMINISTRATION:**

7 Chloroprocaine may be administered as a single injection or continuously through an
8 indwelling catheter. As with all local anesthetics, the dose administered varies with the
9 anesthetic procedure, the vascularity of the tissues, the depth of anesthesia and degree of
10 muscle relaxation required, the duration of anesthesia desired, and the physical condition
11 of the patient. The smallest dose and concentration required to produce the desired result
12 should be used. Dosage should be reduced for children, elderly and debilitated patients
13 and patients with cardiac and/or liver disease. The maximum single recommended doses
14 of chloroprocaine in adults are: without epinephrine, 11 mg/kg, not to exceed a maximum
15 total dose of 800 mg; with epinephrine (1:200,000), 14 mg/kg, not to exceed a maximum
16 total dose of 1000 mg. For specific techniques and procedures, refer to standard
17 textbooks.

18 There have been adverse event reports of chondrolysis in patients receiving intra-
19 articular infusions of local anesthetics following arthroscopic and other surgical
20 procedures. Nesacaine is not approved for this use (see **WARNINGS** and **DOSAGE**
21 **and ADMINISTRATION**).

22 **Caudal and Lumbar Epidural Block:** In order to guard against adverse experiences
23 sometimes noted following unintended penetration of the subarachnoid space, the

- 1 following procedure modifications are recommended:
- 2 1. Use an adequate test dose (3 mL of Nesacaine-MPF 3% Injection or 5 mL of
- 3 Nesacaine-MPF 2% Injection) prior to induction of complete block. This test dose
- 4 should be repeated if the patient is moved in such a fashion as to have displaced the
- 5 epidural catheter. Allow adequate time for onset of anesthesia following
- 6 administration of each test dose.
- 7 2. Avoid the rapid injection of a large volume of local anesthetic injection through the
- 8 catheter. Consider fractional doses, when feasible.
- 9 3. In the event of the known injection of a large volume of local anesthetic injection into
- 10 the subarachnoid space, after suitable resuscitation and if the catheter is in place,
- 11 consider attempting the recovery of drug by draining a moderate amount of
- 12 cerebrospinal fluid (such as 10 mL) through the epidural catheter.

13 As a guide for some routine procedures, suggested doses are given below:

- 14 1. Infiltration and Peripheral Nerve Block: NESACAINE or NESACAINE-MPF
- 15 (chlorprocaine HCl Injection, USP)
- 16

Anesthetic Procedure	Solution Concentration %	Volume (mL)	Total Dose (mg)
Mandibular	2	2 – 3	40 – 60
Infraorbital	2	0.5 – 1	10 – 20
Brachial plexus	2	30 – 40	600 – 800
Digital (without epinephrine)	1	3 – 4	30 – 40
Pudendal	2	10 on each	400
Paracervical (see also PRECAUTIONS)	1	side 3 per each of 4 sites	up to 120

1 2. Caudal and Lumbar Epidural Block: NESACAINE-MPF INJECTION.

2 For caudal anesthesia, the initial dose is 15 to 25 mL of a 2% or 3% solution.

3 Repeated doses may be given at 40 to 60 minute intervals.

4 For lumbar epidural anesthesia, 2 to 2.5 mL per segment of a 2% or 3% solution can
5 be used. The usual total volume of Nesacaine-MPF Injection is from 15 to 25 mL.

6 Repeated doses 2 to 6 mL less than the original dose may be given at 40 to 50 minute
7 intervals.

8 The above dosages are recommended as a guide for use in the average adult.

9 Maximum dosages of all local anesthetics must be individualized after evaluating the size
10 and physical condition of the patient and the rate of systemic absorption from a particular
11 injection site.

12 **Pediatric Dosage:** It is difficult to recommend a maximum dose of any drug for children,
13 since this varies as a function of age and weight. For children over 3 years of age who
14 have a normal lean body mass and normal body development, the maximum dose is
15 determined by the child's age and weight and should not exceed 11 mg/kg (5 mg/lb). For
16 example, in a child of 5 years weighing 50 lbs (23 kg), the dose of chlorprocaine HCl
17 without epinephrine would be 250 mg. Concentrations of 0.5–1% are suggested for
18 infiltration and 1–1.5% for nerve block. In order to guard against systemic toxicity, the
19 lowest effective concentration and lowest effective dose should be used at all times.

20 Some of the lower concentrations for use in infants and smaller children are not available
21 in prepackaged containers; it will be necessary to dilute available concentrations with the
22 amount of 0.9% sodium chloride injection necessary to obtain the required final
23 concentration of chlorprocaine injection.

1 **Preparation of Epinephrine Injections**—To prepare a 1:200,000 epinephrine-
2 chloroprocaine HCl injection, add 0.1 mL of a 1 to 1000 Epinephrine Injection USP to 20
3 mL of Nesacaine-MPF Injection.

4 Chloroprocaine is incompatible with caustic alkalis and their carbonates, soaps,
5 silver salts, iodine and iodides.

6 Parenteral drug products should be inspected visually for particulate matter and
7 discoloration prior to administration, whenever injection and container permit. As with
8 other anesthetics having a free aromatic amino group, Nesacaine and Nesacaine-MPF
9 Injections are slightly photosensitive and may become discolored after prolonged
10 exposure to light. It is recommended that these vials be stored in the original outer
11 containers, protected from direct sunlight. Discolored injection should not be
12 administered. If exposed to low temperatures, Nesacaine and Nesacaine-MPF Injections
13 may deposit crystals of chloroprocaine HCl which will redissolve with shaking when
14 returned to room temperature. The product should not be used if it contains undissolved
15 (eg, particulate) material.

1 **HOW SUPPLIED:**

2 NESACAINE (chloroprocaine HCl Injection, USP) with preservatives is supplied as follows:

Product No.	NDC No.	Strength	Vial size
470530	63323-475-30	1% (10 mg/mL)	30 mL multiple dose vial packaged individually.
470630	63323-476-30	2% (20 mg/mL)	30 mL multiple dose vial packaged individually.

3

4 NESACAINE-MPF (chloroprocaine HCl Injection, USP) without preservatives and without

5 EDTA is supplied as follows:

Product No.	NDC No.	Strength	Vial size
470720	63323-477-20	2% (20 mg/mL)	20 mL single dose vial packaged individually.
470820	63323-478-20	3% (30 mg/mL)	20 mL single dose vial packaged individually.

6

7 Keep from freezing. Protect from light. Store at 20° to 25°C (68° to 77°F) [see USP Controlled
8 Room Temperature].

9 All trademarks are the property of APP Pharmaceuticals, LLC.

10 Manufactured for:



11

12 451109C/Revised: February 2010