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Package Insert (for combined 250mL and 500 mL sizes)

BSS PLUS® STERILE INTRAOCULAR IRRIGATING SOLUTION
(balanced salt solution enriched with bicarbonate, dextrose, and glutathione)

DESCRIPTION: BSS PLUS ® is a sterile intraocular irrigating solution for use during all intraocular surgical procedures, including those requiring a relatively long intraocular perfusion time (e.g., pars plana vitrectomy, phacoemulsification, extracapsular cataract extraction/lens aspiration, anterior segment reconstruction, etc.). The solution does not contain a preservative and should be prepared just prior to use in surgery.

Part I: Part I is a sterile 240 mL or 480 mL solution in a 250 mL or 500 mL single-dose bottle to which the Part II concentrate is added. Each mL of Part I contains: Sodium Chloride 7.44 mg, Potassium Chloride 0.395 mg, Dibasic Sodium Phosphate 0.433 mg, Sodium Bicarbonate 2.19 mg, Hydrochloric Acid and/or Sodium Hydroxide (to adjust pH), in Water for Injection.

Part II: Part II is a sterile concentrate in a 10 mL or 20 mL single-dose vial for addition to Part I. Each mL of Part II contains: Calcium Chloride Dihydrate 3.85 mg, Magnesium Chloride Hexahydrate 5 mg, Dextrose 23 mg, Glutathione Disulfide (Oxidized Glutathione) 4.6 mg, in Water for Injection.

After addition of BSS PLUS Part II to the Part I bottle, each mL of the reconstituted product contains: sodium chloride 7.14 mg, potassium chloride 0.38 mg, calcium chloride dihydrate 0.154 mg, magnesium chloride hexahydrate 0.2 mg, dibasic sodium phosphate 0.42 mg, sodium bicarbonate 2.1 mg, dextrose 0.92 mg, glutathione disulfide (oxidized glutathione) 0.184 mg, hydrochloric acid and/or sodium hydroxide (to adjust pH), in water for injection.

The reconstituted product has a pH of approximately 7.4. Osmolality is approximately 305 mOsm.

CLINICAL PHARMACOLOGY: None of the components of BSS PLUS are foreign to the eye, and BSS PLUS has no pharmacological action. Human perfused cornea studies have shown BSS PLUS to be an effective irrigation solution for providing corneal detumescence and maintaining corneal endothelial integrity during intraocular perfusion. An *in vivo* study in rabbits has shown that BSS PLUS is more suitable than normal saline or Balanced Salt Solution for intravitreal irrigation because BSS PLUS contains the appropriate bicarbonate, pH, and ionic composition necessary for the maintenance of normal retinal electrical activity. Human *in vivo* studies have demonstrated BSS PLUS to be safe and effective when used during surgical procedures such as pars plana vitrectomy, phacoemulsification, cataract extraction/lens aspiration and anterior segment reconstruction. No differences have been observed between adults and pediatric patients following use of this drug product.

INDICATIONS AND USAGE: BSS PLUS is indicated for use as an intraocular irrigating solution during intraocular surgical procedures involving perfusion of the eye.

CONTRAINDICATIONS: There are no specific contraindications to the use of BSS PLUS; however, contraindications for the surgical procedure during which BSS PLUS is to be used should be strictly adhered to.

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WARNINGS: For IRRIGATION during ophthalmic surgery only. Not for injection or intravenous infusion. Do not use unless product is clear, seal is intact, vacuum is present and container is undamaged. Do not use if product is discolored or contains a precipitate.

PRECAUTIONS: **DO NOT USE BSS PLUS UNTIL PART I IS FULLY RECONSTITUTED WITH PART II.** Discard unused contents. BSS PLUS does not contain a preservative; therefore, do not use this container for more than one patient. **Do not use additives other than BSS PLUS Concentrate Part II (10 mL or 20 mL) with this product.** Tissue damage could result if other drugs are added to product. **DISCARD ANY UNUSED PORTION SIX HOURS AFTER PREPARATION.** Studies suggest that intraocular irrigating solutions which are iso-osmotic with normal aqueous fluids should be used with caution in diabetic patients undergoing vitrectomy since intraoperative lens changes have been observed.

There have been reports of corneal clouding or edema following ocular surgery in which BSS PLUS was used as an irrigating solution. As in all surgical procedures appropriate measures should be taken to minimize trauma to the cornea and other ocular tissues.

Preparation: Reconstitute **BSS PLUS®** Intraocular Irrigating Solution just prior to use in surgery. Follow the same strict aseptic procedures in the reconstitution of **BSS PLUS** as is used for intravenous additives. Remove the blue flip-off seal from the **BSS PLUS Part I** (240 mL or 480 mL) bottle. Remove the BSS PLUS Part II overcap and invert the assembly. Insert the transfer spike by holding the vial/collar (to prevent premature activation) into the outer target area of the rubber stopper on the BSS PLUS Part I bottle. Activate fluid transfer by pushing down on the bottom of the Part II vial to actuate the transfer assembly. An excess volume of Part II is provided in each vial. After transfer, remove the Part II vial system from the Part I bottle. Gently agitate the contents to mix the solution. Place a sterile cap over the rubber stopper if the solution is not going to be used immediately. Remove the tear-off portion of the label. Remove the tear-off portion of the label. Record the time and date of reconstitution and the patient's name on the bottle label.

Geriatric Use: No overall differences in safety or effectiveness have been observed between elderly and younger patients.

ADVERSE REACTIONS: Postoperative inflammatory reactions as well as incidents of corneal edema and corneal decompensation have been reported. Their relationship to the use of BSS PLUS has not been established.

OVERDOSAGE: The solution has no pharmacological action and thus no potential for overdose. However, as with any intraocular surgical procedure, the duration of intraocular manipulation should be kept to a minimum.

DOSAGE AND ADMINISTRATION: The solution should be used according to the standard technique employed by the operating surgeon. Use an administration set with an air-inlet in the plastic spike since the bottle does not contain a separate airway tube. Follow the directions for the particular administration set to be used. Insert the spike aseptically into the bottle through the center target area of the rubber stopper. Allow the fluid to flow to remove air from the tubing before intraocular irrigation begins. If a second bottle is necessary to complete the surgical procedure, insure that the vacuum is vented from the second bottle BEFORE attachment to the administration set.

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HOW SUPPLIED: BSS PLUS Solution is supplied in two sizes, 250 mL and 500 mL. BSS PLUS 250 mL is supplied in two packages for reconstitution prior to use: a 250 mL glass bottle, grey butyl stopper and aluminum seal with blue polypropylene flip-off cap containing 240 mL (Part I) and a 24 mL glass vial, grey butyl stopper and plastic transfer spike/overcap assembly containing 10 mL (Part II).

BSS PLUS 500 mL is supplied in two packages for reconstitution prior to use: a 500 mL glass bottle, grey butyl stopper and aluminum seal with blue polypropylene flip-off cap containing 480 mL (Part I) and a 24 mL glass vial, grey butyl stopper and plastic transfer spike/overcap assembly containing 20 mL (Part II).

See the **PRECAUTIONS** section regarding reconstitution of the solution.

250 mL – NDC 0065-0800-25

500 mL - NDC 0065-0800-50

Storage: Store Part I and Part II at 2° - 25°C (36° - 77°F). DO NOT FREEZE. Discard prepared solution after six hours.

Rx Only

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Reconstitution instructions for 250 mL and 500 mL sizes:

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RECONSTITUTION INSTRUCTIONS

BSS PLUS®

STERILE INTRAOCULAR IRRIGATING SOLUTION

(balanced salt solution enriched with bicarbonate, dextrose, and glutathione)

RECONSTITUTION INSTRUCTIONS

DIRECTIONS: Use Aseptic Technique

1. Remove the blue flip-off seal from the BSS PLUS Part I (240 mL or 480 mL) bottle. Prepare the stoppers on the bottle by using a sterile alcohol wipe.
2. Open the BSS PLUS Part II vial/transfer assembly by twisting the overcap at the indicated “Rotate at Arrows”. Place thumb and forefinger above the “Rotate at Arrows” instruction. Squeeze the overcap and lift to remove.
3. Do not touch the transfer spike to avoid contaminating it.
4. Invert the BSS PLUS Part II (10 mL or 20 mL) vial. While holding the vial/collar (to prevent premature activation), insert the transfer spike into the outer target of the rubber stopper of the BSS PLUS Part I (240 mL or 480 mL) bottle. Do not twist the Part II vial while inserting it into the Part I bottle. (See Illustration.)
5. Push down on the BSS PLUS Part II vial and fluid will automatically transfer from the vial into the large Part I bottle. If product flow does not start within a few seconds, grasp the Part I bottle and rotate it in a circular motion. NOTE: An excess amount of BSS PLUS Part II is provided in each vial. A non-transferred solution residual of approximately 0.1 to 0.3 mL can be expected to remain in the vial.
6. Immediately remove transfer assembly from the BSS PLUS Part I container and discard it after solution transfer has been completed.
7. Place a sterile safety cap over the rubber stopper of Part I if the solution is not going to be used immediately. Mix the solution gently until uniform. Peel off the right-hand side of Part I bottle label (fully reconstituted BSS PLUS Solution). Record the patient’s name and the date and time of reconstitution. BSS PLUS Solution is now ready for use.

CAUTION: Reconstituted BSS PLUS Solution must be used within six hours of mixing. Discard any solution which has aged beyond that time. Never use the same bottle of BSS PLUS Solution on more than one patient.

[diagram]