

TISSEEL FIBRIN SEALANT- fibrinogen human, human thrombin
TISSEEL FIBRIN SEALANT- fibrinogen human, human thrombin solution
Baxter Healthcare Corporation

HIGHLIGHTS OF PRESCRIBING INFORMATION

These highlights do not include all the information needed to use TISSEEL safely and effectively. See full prescribing information for TISSEEL.

TISSEEL [Fibrin Sealant] Frozen solution and lyophilized powder for solution for topical use only

Initial U.S. Approval: 1998

RECENT MAJOR CHANGES

Indication and Usage (1) 09/2025

Dosage and Administration (2.1, 2.2, 2.3) 09/2025

INDICATIONS AND USAGE

Hemostasis: TISSEEL is a fibrin sealant indicated for use as an adjunct to hemostasis in adult and pediatric patients undergoing surgery when control of bleeding by conventional surgical techniques (such as suture, ligature, and cautery) is ineffective or impractical. TISSEEL is effective in heparinized patients. (1)

Sealing: TISSEEL is a fibrin sealant indicated in adult and pediatric patients as an adjunct to standard surgical techniques (such as suture and ligature) to prevent leakage from colonic anastomoses following the reversal of temporary colostomies. (1)

DOSAGE AND ADMINISTRATION

For Topical Use Only. (2.1)

- The recommended dosage of TISSEEL is based on the surface area of the wound to be covered to stop bleeding. (2.1)
- Apply TISSEEL as a thin layer by dripping or spraying using cannula or spray set. (2.1, 2.4)
- Ensure that the amount of TISSEEL to be applied is sufficient to entirely cover the intended application area. (2.1)

DOSAGE FORMS AND STRENGTHS

TISSEEL Kit (Freeze-Dried) is supplied as 2 mL, 4 mL and 10 mL (total volume) pack sizes with and without the DUPLOJECT Fibrin Sealant Preparation and Application System (3). TISSEEL Pre-filled Syringe (Frozen) is supplied as 2 mL, 4 mL and 10 mL (total volume) pack sizes with DUPLOJECT COMBI. (3)

CONTRAINDICATIONS

- Do not inject directly into the circulatory system or into highly vascularized tissue, such as nasal mucosa. (4)
- Do not use in individuals with a known hypersensitivity to aprotinin. (4, 5.1, 6.2)
- Do not use for the treatment of severe or brisk arterial or venous bleeding. (4)
- Do not spray where the minimum recommended distance from the applicator tip to the target site cannot be assured. (4)

WARNINGS AND PRECAUTIONS

- Hypersensitivity Reactions: TISSEEL contains aprotinin, a protein known to be associated with anaphylactic reactions. (5.1, 6.2)
- Air or Gas Embolism: To reduce the risk of potential life-threatening gas embolism, spray using only the appropriate pressurized gas at the recommended pressure and distance. For Open Surgical procedures, use the EASYSpray device connected to CO₂, Medical Air or Nitrogen. For Minimally Invasive Surgery procedures use the DUPLOSPRAY MIS device connected only to CO₂. (5.2)
- TISSEEL may be denatured when exposed to solutions containing alcohol, iodine or heavy metals. (5.2)
- Use in Surgery: When using TISSEEL in surgery do not inject intravascularly. (4, 5.3, 6.2)
- Neurosurgical Procedures: Safety has not been evaluated in neurosurgical procedures. (5.4)
- Transmission of Infectious Agents: TISSEEL is made from pooled human plasma and may carry a risk of transmitting infectious agents. (5.4)

ADVERSE REACTIONS

Most common adverse reaction include hypersensitivity including anaphylactic reaction and anaphylactic shock. (6)

To report SUSPECTED ADVERSE REACTIONS, contact Baxter Healthcare Corporation at 1-888-229-0001 or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

----- DRUG INTERACTIONS -----

Oxidized cellulose-containing preparations can reduce the efficacy of TISSEEL and should not be used as carrier materials. (7)

See 17 for PATIENT COUNSELING INFORMATION.

Revised: 9/2025

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FULL PRESCRIBING INFORMATION

1 INDICATIONS AND USAGE

Hemostasis: TISSEEL is a fibrin sealant indicated for use as an adjunct to hemostasis in adult and pediatric patients undergoing surgery when control of bleeding by conventional surgical techniques (such as suture, ligature, and cautery) is ineffective or impractical. TISSEEL is effective in heparinized patients.

Sealing: TISSEEL is a fibrin sealant indicated in adult and pediatric patients as an adjunct to standard surgical techniques (such as suture and ligature) to prevent leakage from colonic anastomoses following the reversal of temporary colostomies.

2 DOSAGE AND ADMINISTRATION

2.1 Recommended Dosage

FOR TOPICAL USE ONLY

The recommended dosage of TISSEEL is based on the surface area of the wound to be covered to stop bleeding.

The approximate surface areas covered by each package size of TISSEEL are listed in Table 1:

Table 1: Surface Area Coverage

Required package size of TISSEEL	Maximum coverage using spray	Maximum coverage using cannula
2 mL	100 cm ²	8 cm ²
4 mL	200 cm ²	16 cm ²
10 mL	500 cm ²	40 cm ²

- Ensure that the amount applied is sufficient to entirely cover the intended application area. Avoid application beyond the intended area.
- Allow at least 2 minutes after application to achieve sufficient polymerization.
- If repeat application is needed, dry the site as much as possible before reapplying. Reapply after removing residues from the prior application or before polymerization takes place since TISSEEL may not adhere firmly to a polymerized layer.
- Only a thin layer of TISSEEL should be applied. Excessive thickness of the fibrin layer may interfere with the wound healing process.

2.2 Preparation and Reconstitution of TISSEEL Kit (Freeze-Dried)

The procedures described below are provided as general guidance.

Do not expose to temperatures above 37°C.

Do not microwave.

After reconstitution, the product must be used within 4 hours and must not be refrigerated or frozen.

Use aseptic technique when preparing and reconstituting the fibrin sealant components. Prior to reconstitution of the fibrin sealant components, clean the rubber stoppers of all vials with a suitable disinfectant, taking care to avoid direct contact with the product.

The Sealer Protein Concentrate is dissolved with the Fibrinolysis Inhibitor Solution (Aprotinin) to form the Sealer Protein Solution. Air-bubbles may make the reconstituted Sealer Protein Solution appear turbid, but such turbidity does not compromise the efficacy or usability of the product. Thrombin Powder is dissolved in the Calcium Chloride Solution to form the Thrombin Solution. Always use separate syringes and cannulas for reconstituting the two solutions to prevent premature clotting. Both solutions should be clear or slightly opalescent. Check the reconstituted solutions visually prior to administration. Do not use solutions which are cloudy, discolored, have deposits, or other changes in their appearance.

1. Reconstitution using the FIBRINOTHERM device

1a. Pre-warming of TISSEEL Lyo (Kit)

Place all four vials from the TISSEEL Kit into the pre-warmed wells of the FIBRINOTHERM device, using the appropriately sized adapter ring(s), and allow the vials to warm for approximately 3 minutes. The FIBRINOTHERM device maintains a constant temperature of 37°C and enables dissolution of the lyophilized Sealer Protein Concentrate by stirring.

1b. Preparation of Sealer Protein Solution (component 1) (FIBRINOTHERM)

- Remove the caps from the Sealer Protein Concentrate and the Fibrinolysis Inhibitor Solution vials.
- Transfer the Fibrinolysis Inhibitor Solution into the vial containing the Sealer Protein Concentrate using one cannula and the blue-scaled syringe provided in the sterile kit for reconstitution.
- Gently swirl the vial to ensure that the product is completely soaked.
- Place the vial into the largest opening of the FIBRINOTHERM device (use an appropriate adaptor, if necessary).
- Switch on the magnetic stirrer and stir until the Sealer Protein Concentrate is dissolved.
- Reconstitution is complete when no particles are visible.
- If particles are present, dissolution can be facilitated by horizontal agitation of the vial (avoid shaking vertically) and keep on stirring the solution at 37°C until the Sealer Protein Concentrate is dissolved completely.
- Turn off the magnetic stirrer when dissolution is complete.

Notes:

- If the Sealer Protein Concentrate has not fully dissolved within 20 minutes discard the vial and prepare a fresh kit.
- Keep the Sealer Protein Solution at 37°C or at room temperature without stirring if it is not used immediately. To ensure homogeneity, briefly swirl the Sealer Protein Solution before drawing it up into the blue-scaled syringe, provided in the sterile kit for application.

1c. Preparation of Thrombin Solution (component 2) (FIBRINOTHERM)

- Remove the caps from the Thrombin Powder and the Calcium Chloride Solution vials.
- Transfer the Calcium Chloride Solution into the Thrombin vial. Use the second cannula and the black-scaled syringe provided in the sterile kit for reconstitution.
- Swirl briefly to dissolve the lyophilized powder and place the vial into the appropriate opening of the FIBRINOTHERM device.
- Keep the Thrombin Solution at 37°C or at room temperature if it is not used immediately. To ensure homogeneity, briefly swirl the Thrombin solution before drawing it up into the black-scaled syringe, provided in the sterile kit for application.

Transferring to the Sterile Field

For transfer of the Sealer Protein and Thrombin Solutions to the sterile field, the circulating nurse should disinfect the tops of the vials with a germicidal solution and allow to dry. The scrub nurse should withdraw the sterile solutions while the circulating nurse holds the non-sterile vials. Slowly withdraw the solution, by firm constant aspiration, to reduce the risk of large air bubbles.

See FIBRINOTHERM device manual for complete operating instructions. If a FIBRINOTHERM device is not available, contact Baxter (1-800-229-0001) for assistance.

2.3 Preparation of TISSEEL Pre-Filled Syringe (Frozen)

Do not expose to temperatures above 37°C. Do not microwave.

Do not refrigerate or re-freeze after thawing.

Do not use TISSEEL (frozen) until it is completely thawed and warmed to 33 - 37°C. To facilitate removal of the tip cap from the syringe, rock the tip cap by moving it backward and forward, then pulling the protective cap off the syringe.

Sterile Water Bath: Transfer inner pouch to the sterile field, remove pre-filled syringe from inner pouch and place directly into sterile water bath ensuring the syringe is completely immersed in the water. Maintain the product at 33-37°C until use. Once the package is opened or the product is warmed to 33-37°C, it must be used within 4 hours.

Non-Sterile Water Bath: Maintain the pre-filled syringe in pouches and place into a water bath outside the sterile field ensuring the pouches remain submerged. Remove from the water bath after thawing and warming, dry the external pouch and transfer inner pouch with pre-filled syringe onto the sterile field. Maintain the product at 33-37°C until use. Once the package is opened or the product is warmed to 33-37°C, it must be used within 4 hours.

Incubator: Maintain the pre-filled syringe in pouches and place into an incubator. Remove from the incubator after thawing and warming. Transfer inner pouch with pre-filled syringe onto the sterile field. Maintain the product at 33-37°C until use. Once the package is opened or the product is warmed to 33-37°C, it must be used within 4 hours.

Table 2: Approximate Water Bath or Incubator Thawing and Warming Times

Pack Size	Sterile Water Bath (Pouches Removed) 33 - 37°C	Non-Sterile Water Bath (In Pouches) 33 - 37°C	Incubator (In Pouches) 33 - 37°C

2 mL	5 minutes	15 minutes	40 minutes
4 mL	5 minutes	20 minutes	50 minutes
10 mL	10 minutes	35 minutes	90 minutes

Room Temperature Thawing: Unopened pouches can be stored for up to 48 hours at room temperature (15-25°C). Before use, warm the product to 33-37°C and apply immediately. The total thawing and warming time cannot exceed 48 hours.

Table 3: Approximate Room Temperature Thawing Times

Pack Size	Room Temperature (In Pouches) 15-25°C
2 mL	80 minutes
4 mL	90 minutes
10 mL	160 minutes

Table 4: Approximate Water Bath or Incubator Warming Times for Thawed Product

Pack Size	Sterile Water Bath (Pouches Removed) 33 - 37°C	Non-Sterile Water Bath (In Pouches) 33 - 37°C	Incubator (In Pouches) 33 - 37°C
2 mL	2 minutes	5 minutes	16 minutes
4 mL	2 minutes	8 minutes	21 minutes
10 mL	4 minutes	12 minutes	35 minutes

The Sealer Protein and the Thrombin Solutions should be clear or slightly opalescent. Do not use solutions that are cloudy, discolored, have deposits, or other changes in their appearance. If one of the above occurs, dispose of the solutions.

The thawed Sealer Protein Solution should be liquid but slightly viscous. If the solution has the consistency of a solidified gel, do NOT use TISSEEL.

2.4 Method of Application

TISSEEL Kit (Freeze-Dried)

Apply TISSEEL using the DUPLOJECT Fibrin Sealant Preparation and Application System or an equivalent delivery system (including open and minimally invasive spray devices) cleared by FDA for use with TISSEEL. Specific instructions for the use of TISSEEL in conjunction with each cleared delivery device are provided with the device.

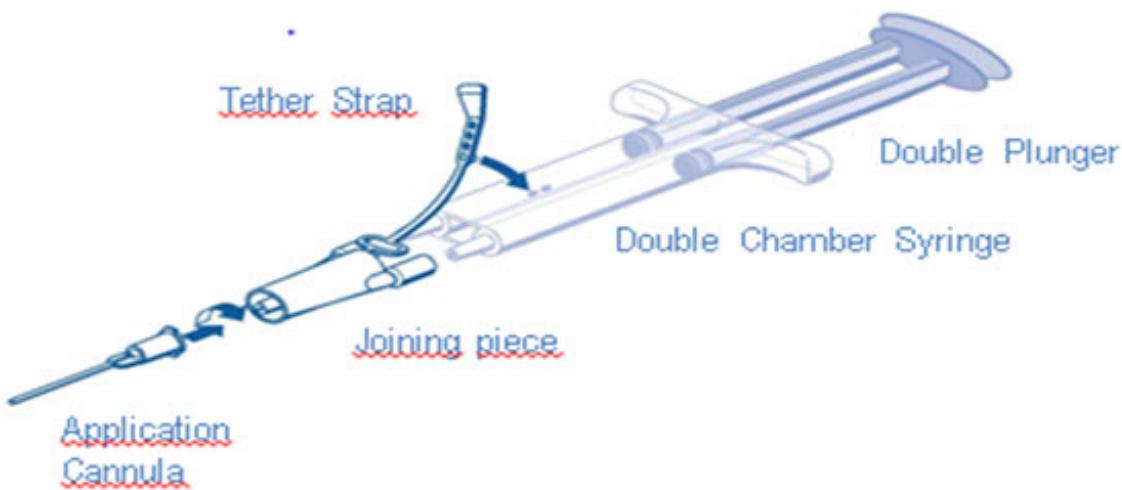
TISSEEL Pre-filled Syringe (Frozen)

Apply pre-filled TISSEEL using the joining piece and application cannula accessory devices provided with the product or an equivalent delivery device (including open and minimally invasive spray devices) cleared by the FDA for use with TISSEEL.

DUPLOJECT COMBI Instructions (see Figure 1)

1. Expel all air from the syringe prior to attaching any application device.
 2. Align the joining piece and tether to the side of the syringe with the tether strap hole.
 3. Connect the nozzles of the double chamber ready-to-use syringe to the joining piece, ensuring that they are firmly attached.
 4. Secure the joining piece by fastening the tether strap to the double chamber ready-to-use syringe.
 5. If the tether strap tears, use the spare joining piece provided in the kit. If a spare joining piece is not available, the system can still be used if care is taken to ensure that the connection is secure and leak-proof.
 6. Do NOT expel the air remaining inside the joining piece.
 7. Attach an application cannula on to the joining piece.
 8. Do NOT expel the air remaining inside the joining piece and inside the application cannula until the start of the actual application because this may clog the application cannula.
- Note: Interruption of TISSEEL application causes clogging in the cannula. Replace the cannula immediately prior to resuming application. If the opening of the joining piece (Y-connector) facing the cannula is clogged, use the spare joining piece provided in the package.

Figure 1
DUPLOJECT COMBI



TISSEEL must be sprayed only onto application sites that are visible. Dry the site of application as much as possible. The surface area of the wound needs to be dried using standard techniques (e.g. intermittent application of compresses, swabs, use of suction devices). Do not use pressurized air or gas to dry the site.

When applying TISSEEL using a spray device, utilize the recommended gas, pressure and distance from tissue within the ranges recommended by the manufacturer as follows:

Table 5: Recommended Application Equipment, Gas and Parameters

Surgery	Spray set / Applicator tips to use	Pressure regulator to use	Gas	Distance	Spray Pressure
Open surgery	TISSEEL /ARTISS Spray Set	EASY SPRAY Pressure Regulator	Medical grade CO ₂ [*] , Compressed Air or Nitrogen	10-15 cm	1.5-2.0 bar (21.8-29.0 psi)
Laparoscopic/ minimally invasive procedures	DUPLOSPRAY MIS Applicator 20 cm DUPLOSPRAY MIS Applicator 30 cm DUPLOSPRAY MIS Applicator 40 cm 360° Flexible Applicator 40 cm Replaceable tip	DUPLOSPRAY MIS Regulator	CO ₂ Only	Range 2-5 cm 3 cm recommended	1.18-1.50 bar (17-22 psi)

* Medical grade CO₂ is the preferred gas for application, however compressed Air or Nitrogen are acceptable gasses for administration of TISSEEL in open surgery.

Apply TISSEEL as a thin layer by dripping or spraying using a cannula or spray set approved for use with TISSEEL. To reduce the risk of potentially life-threatening gas embolism, spray TISSEEL using only the appropriate pressurized gas within the pressure range and distance recommended in the device Instructions For Use (see *Warnings and Precautions (5.2)*).

In cases where very small volumes (1-2 drops) are required, expel and discard the first several drops from the application cannula immediately before application to ensure administration of adequately mixed TISSEEL.

Vials and pre-filled syringes are for single-patient use only. Discard any unused product.

It is strongly recommended that every time TISSEEL is applied to a patient, the name and batch number of the product are recorded in order to maintain a link between the patient and the batch of the product.

3 DOSAGE FORMS AND STRENGTHS

TISSEEL Kit (Freeze-Dried) is supplied as 2 mL, 4 mL and 10 mL (total volume) pack sizes with and without the DUPLOJECT Fibrin Sealant Preparation and Application System.

TISSEEL Pre-Filled Syringe (Frozen) is supplied as 2 mL, 4 mL and 10 mL (total volume) pack sizes with the DUPLOJECT COMBI.

4 CONTRAINDICATIONS

TISSEEL is contraindicated for the following.

- Intravascular Application: Do not inject TISSEEL directly into the circulatory system or into highly vascularized tissue, such as nasal mucosa. Intravascular application of TISSEEL can lead to intravascular coagulation, can result in life-threatening thromboembolic events (see *Warnings and Precautions (5.3) and Adverse Reactions (6.2)*). TISSEEL must be applied with caution to minimize any risk of intravascular application, for example in coronary bypass surgery.
- Aprotinin Hypersensitivity: Do not use TISSEEL in individuals with a known hypersensitivity to aprotinin (see *Warnings and Precautions (5.1) and Adverse Reactions (6)*).
- Severe or Brisk Bleeding: Do not use TISSEEL for treatment of severe or brisk arterial or venous bleeding. In these situations, TISSEEL will be washed away in the flow of blood before hemostasis can be attained.
- Application below minimum recommended distance from target site: Do not spray TISSEEL where the minimum recommended distance from the applicator tip to the target site cannot be assured.

5 WARNINGS AND PRECAUTIONS

5.1 Hypersensitivity Reactions

Hypersensitivity reactions including life-threatening anaphylactic reactions have occurred with the use of TISSEEL [see *Adverse Reactions (6.2)*]. Aprotinin included in TISSEEL for its antifibrinolytic properties is associated with hypersensitivity including anaphylactic reactions. Patients who receive repeated applications of TISSEEL over time or in the same setting, or who previously received systemic aprotinin are at a higher risk for hypersensitivity reactions. Hypersensitivity reactions may occur with initial or subsequent applications of TISSEEL.

Document the use of TISSEEL in the patient's records, noting that TISSEEL contains aprotinin, as required for all aprotinin-containing products. Monitor patients for signs and symptoms of hypersensitivity reactions which include bradycardia, tachycardia, hypotension, flushing, bronchospasm, wheezing, dyspnea, nausea, urticaria, angioedema, pruritus, erythema and paresthesia. Discontinue administration of TISSEEL in the event of hypersensitivity reactions and manage according to clinical practice. Remove remaining product from the application site.

5.2 Air or Gas Embolism

Life threatening or fatal air or gas embolism, tissue rupture or gas entrapment with compression, have occurred when Fibrin Sealants were administered using pressurized gas with open regulator spray devices [see *Adverse Reactions (6)*]. This can occur if a spray device is used at higher than recommended pressures and in closer than recommended proximity to the tissue surface. The solubility of compressed CO₂ is greater than either compressed N₂ or Air thereby reducing the potential effect of embolization.

Regardless of the type of gas used, to reduce the incidence of embolization, spray TISSEEL using only the recommended regulator, set within the recommended pressure range, with the appropriate applicator positioned at the recommended distance in Table 5.

Monitor changes in blood pressure, pulse, oxygen saturation and end tidal CO₂ due to the possibility of air or gas embolism.

Use only spray catheters or applicators approved for use with TISSEEL.

Do not spray TISSEEL in enclosed body areas using the EASYSPRAY device. Spray only on the visible application sites.

For Open Surgical Procedures, use the EASYSPRAY Pressure Regulator connected to medical grade CO₂, compressed Air or a Nitrogen compressed gas source along with the TISSEEL/ARTISS spray set (see *Method of Application* (2.4)).

For Minimally Invasive Surgery Procedures in enclosed body areas use of the DUPLOSPRAY MIS device connected only to compressed CO₂, along with DUPLOSPRAY applicator is recommended. The DUPLOSPRAY MIS device is specifically designed to prevent over pressurization of the body cavity through a dedicated ventline to reduce the risk of gas embolization, (see *Method of Application* (2.4)).

The Sealer Protein and Thrombin solutions may be denatured by alcohol, iodine or heavy metal ions. If any of these substances have been used to clean the wound area, the area must be thoroughly rinsed before the application of TISSEEL.

5.3 Use in Surgery

When using TISSEEL in surgery, do not inject intravascularly (see *Contraindications* (4) and *Adverse Reactions* (6.2)).

5.4 Use in Neurosurgical Procedures

The safety and effectiveness of TISSEEL used alone or in combination with biocompatible carriers in neurosurgical procedures or other surgeries involving confined spaces have not been evaluated, and its use in this setting is not approved by FDA (see *Adverse Reactions* (6.2) and *Drug Interactions* (7)).

5.5 Transmission of Infectious Agents

TISSEEL is made from human blood, therefore it may carry a risk of transmitting infectious agents, e.g., viruses, the variant Creutzfeldt-Jakob disease (vCJD) agent and, theoretically the Creutzfeldt-Jakob disease (CJD) agent. All infections thought possibly to have been transmitted by this product should be reported by the physician or other healthcare provider to Baxter Healthcare Corporation at 1-888-229-0001.

6 ADVERSE REACTIONS

6.1 Clinical Trials Experience

Because clinical trials are conducted under widely varying conditions, adverse reaction rates observed in the clinical trials of a drug cannot be directly compared to rates in the clinical trials of another drug and may not reflect the rates observed in practice.

The safety data described in this section reflects exposure of TISSEEL in five clinical trials, Study 1, Study 2, Study 3, Study 4, and Study 5. A total of 452 patients received TISSEEL as an adjunct to hemostasis undergoing surgery when control of bleeding by conventional surgical techniques is ineffective or impractical in Study 1, Study 2, Study

3, and Study 4, or for sealing as an adjunct to standard surgical technique to prevent leakage from colonic anastomoses following the reversal of temporary colostomies in Study 5. The mean volume of TISSEEL applied was 5.3 mL (range 0.2 – 20.0 mL). The follow up duration for Study 1 was 30 days. The follow up duration for Study 2 and Study 3 was 6 months. The follow up duration for Study 4 was a maximum of 167 days. The follow up duration for Study 5 was a maximum of 69 days (see *Clinical Studies* (14)).

Increased D-Dimer levels have been observed during a clinical study in cardiovascular surgery (see *Clinical Studies* (14)), but did not exceed values reported in the literature occurring after this type of surgery. Postoperatively increased D-Dimers can result at least partly from the degradation of Fibrin Sealant.

There were no reports of serious, associated adverse reactions reported above 1% in clinical studies.

6.2 Post-Marketing Experience

Because adverse reactions are reported voluntarily and the population is of uncertain size, it is not always possible to reliably estimate the frequency of these reactions.

The following adverse reactions have been reported in the post-marketing experience.

Immune System Disorders: Hypersensitivity, including anaphylactic reaction and anaphylactic shock. Anaphylactic reactions and anaphylactic shock have included fatal outcomes.

Vascular Disorders: Hypotension, flushing, embolism, including cerebral artery embolism, cerebral infarction*, air embolism**

Skin and subcutaneous Tissue Disorders: Angioedema, erythema, impaired healing, pruritus, urticaria

Cardiac Disorders: Bradycardia, tachycardia

Respiratory Disorders: Bronchospasm, dyspnea, wheezing

Gastrointestinal Disorders: Nausea

Nervous System Disorders: Paresthesia

* As a result of intravascular application into the superior petrosal sinus

** As with other fibrin sealants life-threatening/fatal air or gas embolism when using devices with pressurized air or gas occurred; this event appears to be related to an inappropriate use of the spray device (e.g. at higher than recommended pressure and in close proximity to the tissue surface),

Class effect: Manifestations of hypersensitivity or allergic reactions associated with the class of fibrin sealant/hemostatic products include: application site irritation, chest discomfort, chills, headache, lethargy, restlessness and vomiting.

There have been reports of serious adverse events such as paralysis and other compressive complications possibly related to the use of fibrin sealants in combination with resorbable hemostatic agents. There have also been reports of fatalities following the misadministration of topical thrombin (see *Warnings and Precautions* (5.4)).

7 DRUG INTERACTIONS

Oxidized cellulose-containing preparations can reduce the efficacy of TISSEEL and should not be used as carrier materials. No interaction studies have been performed.

8 USE IN SPECIFIC POPULATIONS

8.1 Pregnancy

Risk Summary

There are no direct or controlled studies of TISSEEL in pregnant women. No animal reproductive and developmental toxicity studies have been conducted with TISSEEL. It is also not known whether TISSEEL can cause fetal harm when administered to a pregnant woman or can affect reproduction capacity. Some viruses, such as parvovirus B19, are particularly difficult to remove or inactivate. Parvovirus B19 most seriously affects pregnant women (fetal infection). In the United States general population, the estimated background risk of major birth defects and miscarriage in clinically recognized pregnancies is 2% to 4% and 15% to 20%, respectively.

8.2 Lactation

Risk Summary

There is no information regarding the presence of TISSEEL in human milk, the effects on the breastfed infant, or the effects on milk production. The developmental and health benefits of breastfeeding should be considered along with the mother's clinical need for TISSEEL and any potential adverse effects on the breastfed infant from TISSEEL or from underlying maternal condition.

8.4 Pediatric Use

The safety and efficacy of TISSEEL have been established in pediatric patients. The use of TISSEEL as an adjunct to hemostasis was supported by evidence from one clinical study which included 27 pediatric patients aged 1 month to 16 years. The use of TISSEEL as an adjunct to sealing was supported by extrapolation of the data from one clinical study in adult patients [see *Adverse Reactions (6) and Clinical Studies (14)*].

8.5 Geriatric Use

Clinical studies included 218 patients aged 65 years of age or older treated with TISSEEL (159 undergoing cardiac surgery and 59 undergoing vascular surgery) (see *Clinical Studies (14)*). No overall differences in safety or effectiveness were observed between these patients and younger patients.

11 DESCRIPTION

TISSEEL [Fibrin Sealant] is a two-component fibrin sealant made from pooled human plasma. When combined, the two components, Sealer Protein and Thrombin mimic the final stage of the blood coagulation cascade.

Sealer Protein (Human)

Sealer Protein (Human) is a sterile, non-pyrogenic, vapor heated and solvent/detergent treated preparation made from pooled human plasma. Sealer Protein (Human) is provided either as a freeze-dried powder for reconstitution with Aprotinin (Synthetic) solution or as a finished frozen solution pre-filled into one side of a dual-chambered syringe. The active ingredient in Sealer Protein (Human) is fibrinogen. Sealer Protein

(Human) Solution contains fibrinolysis inhibitor, synthetic Aprotinin, that delays fibrinolysis. Aprotinin (Synthetic) is manufactured by solid phase synthesis from materials completely of non-human/non-animal origin.

Thrombin (Human)

Thrombin (Human) is a sterile, non-pyrogenic, vapor-heated and solvent/detergent treated preparation made from pooled human plasma. Thrombin (Human) is also provided either as a freeze-dried powder for reconstitution with Calcium Chloride Solution or as a finished frozen solution pre-filled into one side of a dual-chambered syringe.

The reconstituted solution or pre-filled syringe contains:

Sealer Protein Solution

Total protein: 96 – 125 mg/mL

Fibrinogen: 67 – 106 mg/mL

Aprotinin (Synthetic): 2250 – 3750 KIU/mL

Other ingredients include: human albumin, tri-sodium citrate, histidine, niacinamide, polysorbate 80 and water for injection.

Thrombin Solution

Thrombin (Human): 400 – 625 units/mL*

Calcium Chloride: 36 – 44 µmol/mL

Other ingredients include: human albumin, sodium chloride and water for injection.

* The potency expressed in units is determined with a clotting assay using an in-house internal standard that has been calibrated against the World Health Organization (WHO) Second International Standard for Thrombin, 01/580.

Therefore, a unit (U) is equivalent to an International Unit (IU).

Viral Clearance

TISSEEL is made from pooled human plasma collected at US licensed collection centers. The vapor heat and solvent/detergent treatment steps used in the manufacturing process have been shown to be capable of significant viral reduction. No procedure, however, has been shown to be completely effective in removing viral infectivity from derivatives of human plasma (see *Warnings and Precautions (5.4)*). Validation studies were conducted using samples drawn from manufacturing intermediates for each of the two human plasma derived components. These samples were spiked with stock virus suspensions of known titers followed by further processing under conditions representative of respective manufacturing steps.

The virus reduction factors (expressed as \log_{10}) of manufacturing steps for each of the viruses tested are shown in Table 6.

Table 6: Mean Reduction Factors [\log_{10}] for Virus Removal and/or Inactivation

Sealer Protein Component					
Manufacturing Step	HIV-1 ¹	BVDV ¹	PRV ¹	HAV ²	MMV ²
Early Manufacturing Steps	n.d.	n.d.	n.d.	n.d.	2.7
Solvent/Detergent Treatment	≥4.7	≥5.3	≥5.5	n.d.	n.d.
Vapor Heat Treatment	>5.9	>6.5	>6.8	>6.2	1.4
Overall Reduction Factor	>10.6	>11.8	≥12.3	>6.2	4.1

Thrombin Component					
Manufacturing Step	HIV-1 ¹	BVDV ¹	PRV ¹	HAV ²	MMV ²
Thrombin Precursor Mass Capture	3.2	1.8	2.5	1.5	1.2
Vapor Heat Treatment	>5.3	>5.9	>6.8	>4.6	1.0
Solvent/Detergent Treatment	≥5.3	≥5.3	≥6.2	n.d.	n.d.
Ion Exchange Chromatography	n.d.	n.d.	n.d.	n.d.	3.6
Overall Reduction Factor (ORF)	>13.8	>13.0	>15.5	>6.1	5.8

¹ Enveloped

² Non-enveloped

n.d. = not determined

HIV-1: Human Immunodeficiency Virus 1; **HAV:** Hepatitis A Virus; **BVDV:** Bovine Viral Diarrhea Virus, a model for Hepatitis C Virus; **PRV:** Pseudorabies Virus, a model for lipid enveloped DNA viruses, among those is Hepatitis B Virus; **MMV:** Mouse Minute Virus, a model for B19V.

In addition, Human Parvovirus B19 (B19V) was used to investigate the upstream Thrombin precursor mass capture step, the Sealer Protein early manufacturing steps and the Thrombin and Sealer Protein vapor heating steps. Using quantitative PCR assays, the estimated B19V log reduction factors were: (a) 1.7 for the Thrombin precursor mass capture step, (b) 3.4 for Sealer Protein early manufacturing steps, (c) >4 for Thrombin vapor heat treatment and (d) 1.0 for Sealer Protein vapor heat treatment.

12 CLINICAL PHARMACOLOGY

12.1 Mechanism of Action

Upon mixing Sealer Protein (Human) and Thrombin (Human), soluble fibrinogen is transformed into fibrin, forming a rubber-like mass that adheres to the wound surface and achieves hemostasis and sealing or gluing of tissues. TISSEEL mimics the final coagulation cascade step as it has all relevant components to form a clot. TISSEEL is effective in heparinized patients and in patients medicated with anti-platelet drugs.

12.2 Pharmacodynamics

Thrombin is a highly specific protease that transforms the fibrinogen contained in Sealer Protein (Human) into fibrin. Fibrinolysis inhibitor, Aprotinin (Synthetic), is a polyvalent protease inhibitor that prevents premature degradation of fibrin. Preclinical studies with different fibrin sealant preparations simulating the fibrinolytic activity generated by extracorporeal circulation in patients during cardiovascular surgery have shown that incorporation of aprotinin in the product formulation increases resistance of the fibrin sealant clot to degradation in a fibrinolytic environment.

12.3 Pharmacokinetics

Unincorporated Aprotinin and its metabolites have a half-life of 30 to 60 minutes and are eliminated by the kidney. Pharmacokinetic studies were not conducted. TISSEEL is expected to be completely resorbed in 10 to 14 days.

Because TISSEEL is applied only topically, systemic exposure or distribution to other organs or tissues is not expected.

13 NONCLINICAL TOXICOLOGY

13.1 Carcinogenesis, Mutagenesis, Impairment of Fertility

Long-term animal studies to evaluate the carcinogenic potential of TISSEEL or studies to determine the effect of TISSEEL on fertility have not been performed.

14 CLINICAL STUDIES

The efficacy of TISSEEL was evaluated in five clinical studies, Study 1 (NCT00892957), Study 2 (NCT00161733), Study 3, Study 4, and Study 5 as described below.

Study 1 (Vascular Surgery)

TISSEEL was evaluated in a prospective, controlled, randomized, single-blind, multicenter clinical study against manual compression with gauze pads in 140 patients 33 to 90 years of age undergoing vascular surgery with expanded polytetrafluoroethylene (ePTFE) graft placement (arterio-arterial bypasses and AV shunts for dialysis access in the upper and lower extremity). Patients received standardized dosages of heparin. Protamine was administered after the primary endpoint had been assessed. Long-term antiplatelet treatments were continued perioperatively at the surgeon's discretion.

Patients were randomly assigned to TISSEEL or control when persistent bleeding at the study suture line was present after surgical hemostasis, i.e., sutures. Eligible bleedings before clamping and treatment application were defined as a minimum of 25% of the suture line bleeds or at least 5 suture line bleedings or any pulsatile or spurting needle hole bleeding. The primary efficacy endpoint was hemostasis achieved at the study suture line at 4 minutes and maintained until surgical closure.

The efficacy results are summarized in Table 7 below.

Table 7: Efficacy Results for Study 1 (Vascular Surgery)

Endpoint	TISSEEL n = 70	Manual Compression n = 70	p-value*
Hemostasis at the study suture line within 4 minutes and maintained until surgical closure	44 (63%)	22 (31%)	<0.0001

* likelihood ratio chi-square test; 2.5% one sided

Study 2 (Cardiac Surgery)

TISSEEL was evaluated in a prospective, parallel design, randomized (1:1), double-blind, multicenter clinical study against an earlier formulation of the product, TISSEEL VH, in 317 patients 19 to 94 years of age undergoing cardiac surgery requiring cardiopulmonary bypass (CPB) and median sternotomy. Patients were treated with TISSEEL or the control product only when hemostasis was not achieved by conventional surgical methods. For the endpoint, hemostasis achieved at the primary treatment site within 5 minutes of treatment and maintained until closure of the surgical wound, TISSEEL was non-inferior to the earlier formulation of the product using a one-sided 97.5% confidence interval on the difference in the proportion of patients successfully treated.

Table 8: Efficacy Result for Study 2 (Cardiac Surgery)

Endpoint	TISSEEL n = 144	TISSEEL VH n = 144	One-Sided Confidence Interval
Hemostasis within 5 minutes and maintained until surgical closure	127 (88%)	129 (90%)	97.5%

Study 3 (Cardiac Reoperations)

An earlier formulation of TISSEEL was evaluated in an open-label crossover study against control topical hemostatic agents in 489 patients <10 to 80 years of age undergoing cardiovascular reoperation or resternotomy at 11 institutions. Patients were randomized to TISSEEL or control hemostatic agents when a topical hemostatic was needed at the conclusion of surgery and after all attempts at surgical hemostasis. Patients were crossed to the alternative therapy if bleeding continued after the 5 minute endpoint. At 10 centers, TISSEEL was used after administration of protamine sulfate. At one site, TISSEEL could be used before administration of protamine sulfate. 365 of the 489 patients developed bleeding episodes requiring treatment. For the endpoint (successful hemostasis at 5 minutes), TISSEEL was statistically significantly superior to control topical hemostatic agents in these patients. Similarly, absolute time to cessation of bleeding was statistically significantly shorter for TISSEEL than for control topical hemostatic agents ($p<0.0001$, Gehan-Wilcoxon test, two sided).

Table 9: Efficacy Result for Study 3 (Cardiac Reoperations)

Endpoint	TISSEEL n = 193	Control Topical Hemostatic Agent n = 172	p-value*
Hemostasis within 5 minutes	159 (82%)	76 (45%)	<0.0001

* Pearson χ^2 two sided; $p <0.0001$; intent-to-treat analysis

1. Pearson χ^2 two sided; $p <0.0001$; intent-to-treat analysis

Study 4 (Splenectomy)

In a single center, open label trial, an earlier formulation of TISSEEL was compared to

historical controls in patients undergoing laparotomy for blunt or penetrating traumatic injury to the spleen and/or liver. Use of TISSEEL resulted in the need for statistically significantly fewer splenectomies than control hemostatic maneuvers. TISSEEL did not result in significantly reduced mortality in patients with blunt or penetrating trauma to the liver alone or to the liver and spleen ($p=0.067$, c^2 , one sided).

The population demographics include: 15 - 72 years.

Table 10: Efficacy Result for Study 4 (Splenectomy)

Injury to:	TISSEEL	Historic Controls	p-value*
Spleen	0/19	14/22	$p <0.001$
Spleen and Liver	1/26	19/34	$p <0.001$

Study 5 (Colostomy Closure)

In a single center, prospective open label study of 118 patients 18 to 73 year of age randomized to standard of care (58 patients) or standard of care plus fibrin sealant (60 patients) for elective colostomy closure after temporary colostomy placement for treatment of traumatic injury to the colon, the earlier version of TISSEEL plus standard of care was also shown to be significantly superior to standard of care alone ($p=0.0406$, Jonckheere-Terpstra test for ordinal data, two sided) with regard to anastomotic complications (leakage, intra-abdominal abscess formation, re-operation, septic shock, and death).

16 HOW SUPPLIED/STORAGE AND HANDLING

How Supplied

TISSEEL is supplied in the following pack sizes and presentations:

Table 11: NDC Numbers

Pack Size	TISSEEL Kit (Freeze-Dried)	TISSEEL Kit (Freeze-Dried) with DUPLOJECT System	TISSEEL Pre-Filled Syringe (Frozen) with DUPLOJECT COMBI
2 mL	0338-4210-02	0338-4301-02	0338-9560-01
4 mL	0338-4211-04	0338-4302-04	0338-9564-01
10 mL	0338-4212-10	0338-4303-10	0338-9568-01

TISSEEL Kit contains one vial each of:

1. Sealer Protein Concentrate (Human), Vapor Heated, Solvent/Detergent Treated, Freeze-Dried, Sterile

2. Fibrinolysis Inhibitor Solution, Aprotinin (Synthetic) Liquid, Sterile
3. Thrombin (Human), Vapor Heated, Solvent/Detergent Treated, Freeze-Dried, Sterile
4. Calcium Chloride Solution, Liquid, Sterile
5. With and without DUPLOJECT Fibrin Sealant Preparation and Application System

TISSEEL Pre-Filled Dual-Chambered Syringe contains:

1. Sealer Protein Solution, Vapor Heated, Solvent/Detergent Treated, Frozen Solution, Sterile
2. Thrombin Solution, Vapor Heated, Solvent/Detergent Treated, Frozen Solution, Sterile
3. Sterile accessory devices (DUPLOJECT COMBI)

Storage and Handling

TISSEEL Kit (Freeze-Dried)

Store at 2-25°C. Avoid freezing. Do not freeze or refrigerate reconstituted solutions.

TISSEEL Pre-filled Syringe (Frozen)

Store at ≤ -20°C. Do not refrigerate or re-freeze after thawing. Once removed from the freezer, TISSEEL must be used within 48 hours. Prior to application, TISSEEL must be warmed to 33 - 37°C.

Once the pouches are opened or warmed to 33-37°C, they must be used within 4 hours.

Do not use after the expiration date. Discard if packaging of any components is damaged.

17 PATIENT COUNSELING INFORMATION

Discuss following with the patient and/or caregivers.

- Hypersensitivity Reactions: Inform the patient and/or caregivers that hypersensitivity reactions may occur with TISSEEL and advise to seek immediate medical evaluation if any signs and symptoms of hypersensitivity occur. These may include bradycardia, tachycardia, hypotension, flushing, bronchospasm, wheezing, dyspnea, nausea, urticaria, angioedema, pruritus, erythema and paresthesia [see *Warnings and Precautions (5.1)*].
- Transmission of Infectious agents: Inform the patient and/or caregivers that TISSEEL is made from human plasma and may carry a risk of transmitting infectious agents (e.g., viruses, the vCJD agent and, theoretically, the CJD agent). Instruct patients to report any symptoms that concern them and might be caused by infections [see *Warnings and Precautions (5.5)*].

Instruct patients to consult their physician if symptoms of B19 virus infection appear (fever, drowsiness, chills and runny nose) followed about two weeks later by a rash and joint pain (see *Use in Specific Populations (8.1)*).

Manufactured For Baxter Healthcare Corporation

Deerfield, IL 60015 USA

US License No. 140

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PACKAGE/LABEL PRINCIPAL DISPLAY PANEL





Tissel Lyo 2 mL Vial Carton Label

NDC 0338-4210-02

**Fibrin Sealant
TISSEEL 2 mL
Vapor Heated, Solvent/Detergent Treated, Kit**

GTIN (01)
LOT (10)
EXPIRY (17)
SERIAL (21)

Baxter Logo

NDC 0338-4210-02

Barcode

**Fibrin Sealant
TISSEEL 2 mL
Vapor Heated, Solvent/Detergent Treated, Kit**

Contents:

Sealer Protein Concentrate (Human), Vapor Heated, Solvent/Detergent Treated, freeze dried, sterile, for 1 mL of Sealer Protein Solution 86.5 mg/mL

Fibrinolysis Inhibitor Solution (Synthetic), sterile, 3000 KIU of Aprotinin /mL

Thrombin (Human), Vapor Heated, Solvent/Detergent Treated, freeze dried, sterile, for 1 mL of Thrombin

Solution 500 units/mL

Calcium Chloride Solution, sterile, 40 µmol /mL

The risks and benefits of this product should be discussed with the patient.

Read enclosed directions for reconstitution and application before use.

Use within four hours of reconstitution.

NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only

Manufactured for Baxter Healthcare Corporation

Deerfield, IL 60015 USA

U.S. License No. 140

Made in Austria

U.S. Pat. No.: 5,962,405

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0752668

Fibrin Sealant

TISSEEL 2 mL

**Vapor Heated, Solvent/
Detergent Treated, Kit**

2 mL

Store at 2°C to 25°C (36°F to 77°F).

Avoid freezing.

U.S. Pat. No.: 5,962,405

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Barcode Number: 150451A

Made in Austria

U.S. License No. 140

Deerfield, IL 60015 USA

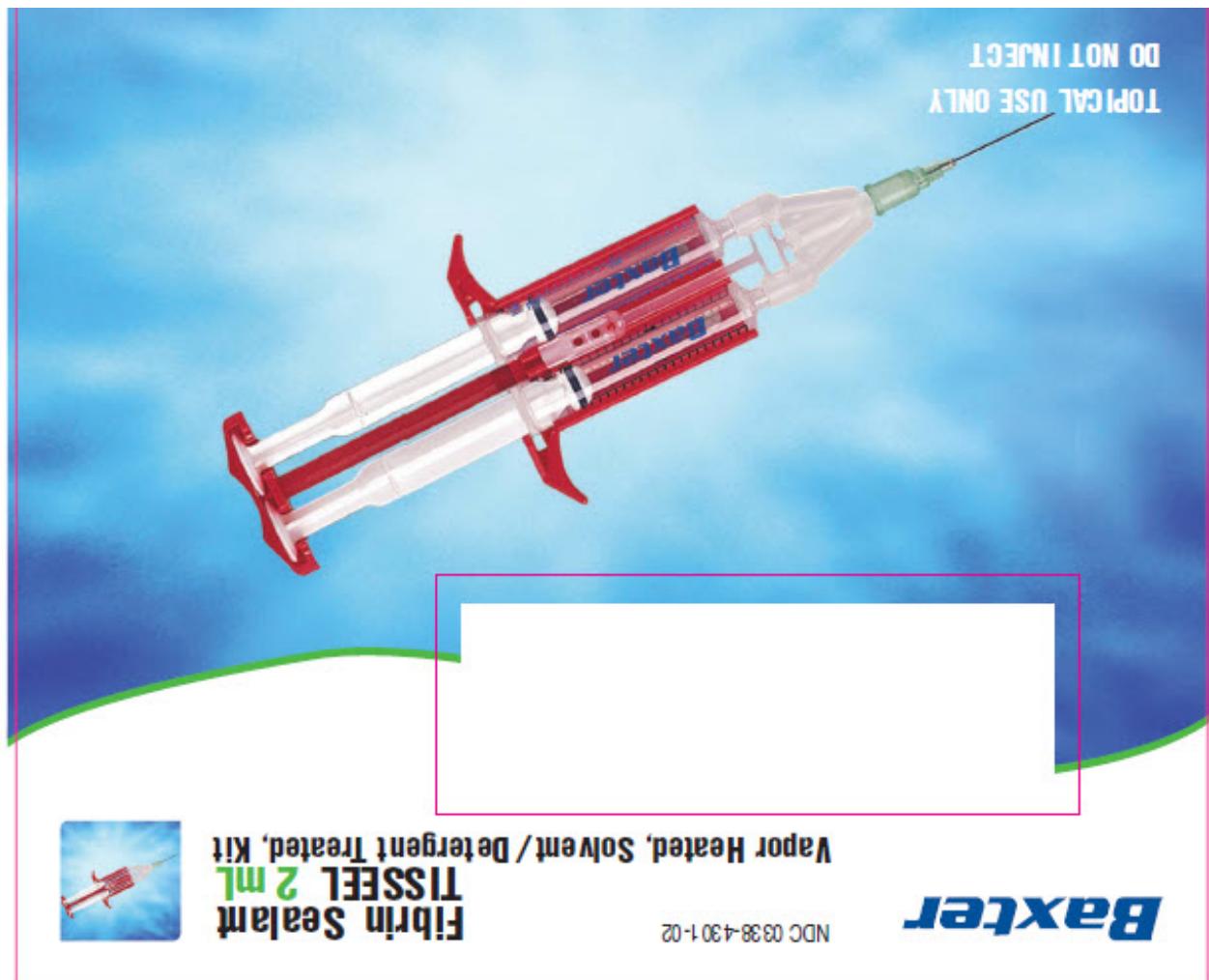
Manufactured for Baxter Healthcare Corporation



Vapor Heated, Solvent/Detergent Treated, Kit
TISSEEL 2 ML
Fibrin Sealant

NDC 0338-4301-02

Baxter



Baxter

NDC 0338-4301-02



**Fibrin Sealant
TISSEEL 2 mL**

Vapor Heated, Solvent / Detergent Treated, Kit



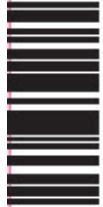
Contents:

Sealer Protein Concentrate (Human), Vapor Heated, Solvent/Detergent Treated, freeze dried, sterile, for 1 mL of Sealer Protein Solution 86.5 mg/mL.
Fibrinolysis Inhibitor Solution (Synthetic), sterile, 3000 KIU of Aprotinin/mL.
Thrombin (Human), Vapor Heated, Solvent/Detergent Treated, freeze dried, sterile, for 1 mL of Thrombin Solution 500 units/mL.
Calcium Chloride Solution, sterile, 40 µmol/mL.

The risks and benefits of this product should be discussed with the patient. Read enclosed directions for reconstitution and application before use. Use within four hours of reconstitution.

NOT FOR INJECTION. For single use only.
Not made with natural rubber latex
Rx only

Also Includes: DUPLOJECT Fibrin Sealant Preparation and Application System 2 mL / 4 mL



Tisseel Lyo 2mL Sleeve Carton Label

Baxter Logo

NDC 0338-4301-02

Fibrin Sealant

TISSEEL 2 mL

Vapor Heated, Solvent/Detergent Treated, Kit

Manufactured for Baxter Healthcare Corporation

Deerfield, IL 60015 USA

U.S. License No. 140

Made in Austria

Reorder Number: 1504514

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U.S. Pat. No.: 5,962,405

Baxter Logo

NDC 0338-4301-02

Fibrin Sealant

TISSEEL 2 mL

Vapor Heated, Solvent/Detergent Treated, Kit

TOPICAL USE ONLY

DO NOT INJECT

Baxter Logo

NDC 0338-4301-02

Barcode

Fibrin Sealant

TISSEEL 2 mL

Vapor Heated, Solvent/Detergent Treated, Kit

Contents:

Sealer Protein Concentrate (Human), Vapor Heated, Solvent/Detergent Treated, freeze dried, sterile, for 1 mL of Sealer Protein Solution 86.5 mg/mL

Fibrinolysis Inhibitor Solution (Synthetic), sterile, 3000 KIU of Aprotinin /mL

Thrombin (Human), Vapor Heated, Solvent /Detergent Treated, freeze dried, sterile, for 1 mL of Thrombin Solution 500 units/mL

Calcium Chloride Solution, sterile, 40 μmol /mL

The risks and benefits of this product should be discussed with the patient. Read enclosed directions for reconstitution and application before use. Use within four hours of reconstitution.

NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only

Also includes: DUPLOJECT Fibrin Sealant Preparation and Application System 2 mL / 4 mL

0752670

Barcode

Barcode

N3 0338-4301-02 3



DUPLOJECT 2 mL / 4 mL
Fibrin Sealant Preparation and Application System

Baxter

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DUPLOJECT 2 mL / 4 mL
Fibrin Sealant Preparation and Application System



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Manufactured for:
Baxter Healthcare Corporation
One Baxter Parkway
Deerfield, IL 60015 USA
1-888-229-0001

DUPLOJECT 2 mL / 4 mL

Fibrin Sealant Preparation and Application System

For Fibrin Sealant
2 mL or 4 mL Kit



Indications for Use:

For the preparation and application of TISSEEL
Fibrin Sealant 2 mL / 4 mL Kit.

STERILE – SINGLE PATIENT USE ONLY – DO NOT RESTERILIZE

Use separate devices for reconstitution of Sealer Protein Concentrate and Thrombin.

Rx Only.

Made in Belgium



BE8001137

DUPLOJET 2 mL / 4 mL

Fibrin Sealant Preparation and
Application System

For Fibrin Sealant
2 mL or 4 mL Kit



Tisseel Lyo 2 mL - 4 mL DUPLOJET Label

Baxter Logo

DUPLOJET 2 mL / 4 mL

**Fibrin Sealant Preparation and
Application System**

Baxter Logo

DUPLOJET 2 mL / 4 mL

**Fibrin Sealant Preparation and
Application System**

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Baxter International Inc., registered in the U.S. Patent and
Trademark office.

Manufactured for:

Baxter Healthcare Corporation
One Baxter Parkway
Deerfield, IL 60015 USA

DUPLOJET 2 mL / 4 mL

**Fibrin Sealant Preparation and
Application System**

**For Fibrin Sealant
2 mL or 4 mL Kit**

2 mL

4 mL

Indications for Use:

For the preparation and application of TISSEEL

[Fibrin Sealant] 2 mL / 4 mL Kit.

STERILE – SINGLE PATIENT USE ONLY – DO NOT RESTERILIZE

Use separate devices for reconstruction of Sealer Protein Concentrate and Thrombin.

Rx only

Made in Belgium

Barcode

BE8001137

BE-80-01-137

DUPLOJET 2 mL / 4 mL

**Fibrin Sealant Preparation and
Application System**

For Fibrin Sealant

2 mL or 4 mL Kit

2 mL

4 mL

DUPLOJECT

Fibrin Sealant Preparation and Application System

Indications for Use:

For the preparation and application of TISSEEL Fibrin Sealant kit.

Note: See TISSEEL Fibrin Sealant package insert for additional preparation and application instructions.

1.0 Reconstitution Instructions for the Circulating Nurse (Using the FIBRINOTHERM Heating and Stirring Device)

- 1.1 Turn on the warming unit of the FIBRINOTHERM device (amber switch).
- 1.2 Open TISSEEL Fibrin Sealant kit and place all vials into the appropriately sized heating wells of the FIBRINOTHERM device.
- 1.3 Place the Sealer Protein vial into the large magnetic stirring well fitted with the appropriately sized adapter ring (if necessary). Do not activate the stirring mechanism at this time. The indicator light will remain lit until a temperature of 37°C (98.6°F) is reached. Allow several minutes for proper warming.
- 1.4 Open Pack A of the DUPLOJECT Preparation and Application System and assemble the blue-scaled and black-scaled syringes with the needles provided.
- 1.5 Remove the flip-off caps from the blue-capped Sealer Protein Concentrate and Fibrinolysis Inhibitor Solution vials and wipe with a non-iodine based disinfectant.
- 1.6 Using the blue-scaled syringe, withdraw the Fibrinolysis Inhibitor Solution from the vial, tilting the vial slightly to facilitate removal of all solution. Inject the Fibrinolysis Inhibitor Solution into the Sealer Protein Concentrate vial. Gently swirl the vial to ensure that the freeze-dried material is fully soaked. Note: Do not invert or inject air into vials.
- 1.7 Place the Sealer Protein Concentrate vial in the stirring well and activate the stirring mechanism of the FIBRINOTHERM device (green switch). Reconstitution

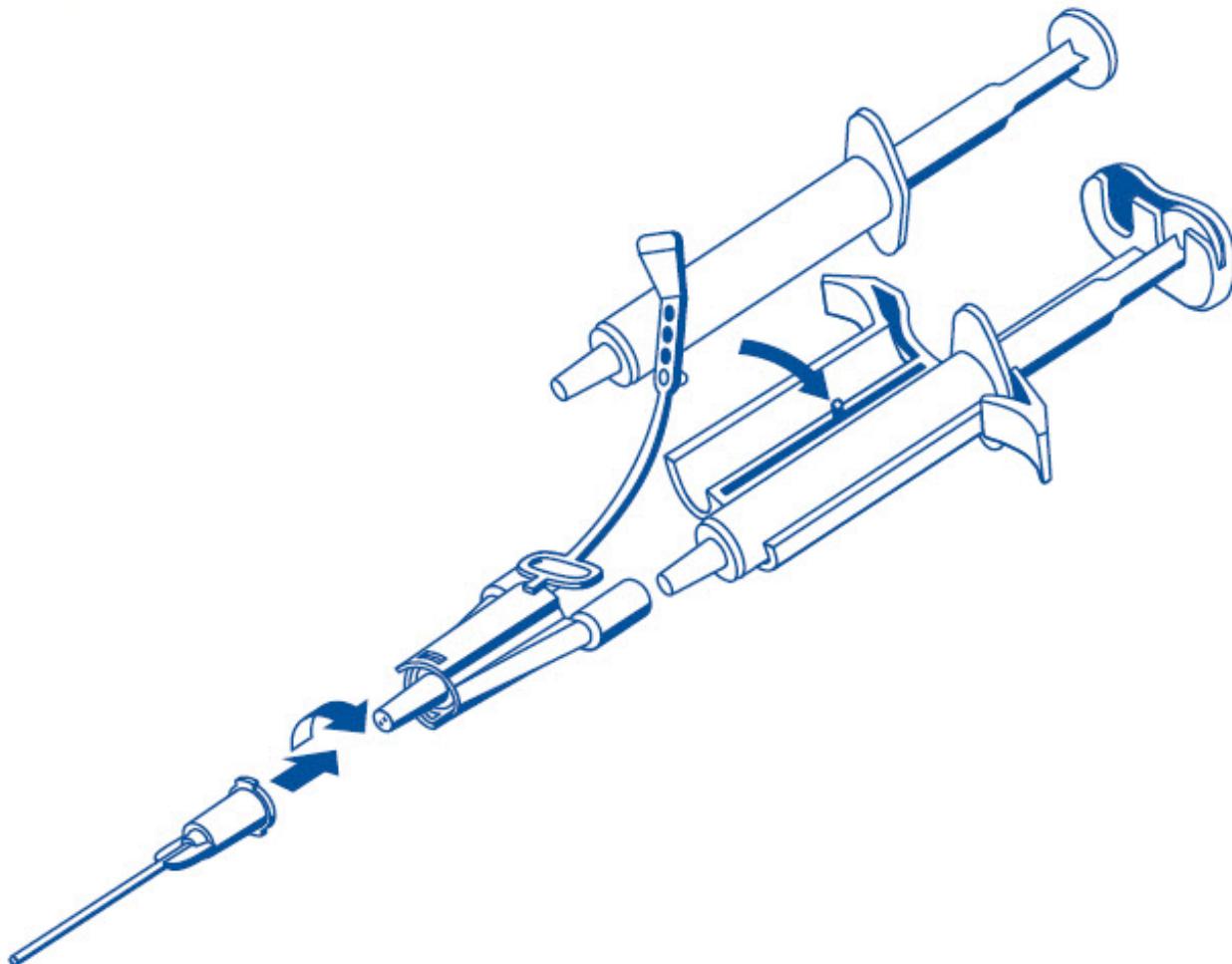
Stirring mechanism of the FIBRINOTHERM device (green switch). Reconstruction is complete when no undissolved particles are visible. If Sealer Protein is not fully dissolved after 20 minutes, discard and prepare a fresh kit. Continue with steps 8 through 11.

- 1.8 Remove the flip-off caps from the black-capped Thrombin and Calcium Chloride Solution vials and wipe with a non-iodine based disinfectant.
- 1.9 Using the black-scaled syringe, withdraw the Calcium Chloride Solution from the vial, tilting the vial slightly to facilitate removal of all solution. Inject the Calcium Chloride Solution into the Thrombin vial. Note: Do not invert or inject air into vials.
- 1.10 Swirl contents of the Thrombin vial briefly and return it to an appropriately sized heating well in the FIBRINOTHERM device.
- 1.11 Leave the Sealer Protein and Thrombin vials in the FIBRINOTHERM device until the solutions are ready to be passed into the sterile field.

2.0 Preparation Instructions for the Scrub Nurse

- 2.1 Open Pack B of the DUPLOJECT Preparation and Application System into the sterile field.
- 2.2 Open Packs 1, 2, and 3.
- 2.3 Assemble the blue-scaled and black-scaled syringes with the needles provided. Note: For 10 mL DUPLOJECT systems, attach the larger needle to the blue-scaled syringe.
- 2.4 While the Circulating Nurse holds the vial slightly tilted, insert needle into vial (bevel side down) and withdraw all of the Sealer Protein Solution into the blue-scaled syringe using slow, constant aspiration. Discard needle in sharps container.
- 2.5 While the Circulating Nurse holds the vial slightly tilted, insert needle into vial (bevel side down) and withdraw all of the Thrombin Solution into the black-scaled syringe using slow, constant aspiration. Discard needle in sharps container.
- 2.6 Remove any air bubbles from the syringes and ensure both syringes contain the same volume.

- 2.7 Snap the filled syringes into the two-syringe clip with the flanges in an up/down position, as illustrated below.



- 2.8 Attach the joining piece to the syringe nozzles, ensuring that both are firmly seated. Secure the joining piece by fastening the retaining strap to the double syringe clip. Note: For 10 ml DUPLOJECT systems, align the syringe nozzles toward the middle for proper attachment.
- 2.9 Fit one of the application cannulas onto the joining piece. A second application cannula is provided in Pack 3 as a spare.
- 2.10 DUPLOJECT Applicator is ready for use.
- 2.11 If application of TISSEEL Fibrin Sealant is interrupted, replace the cannula immediately before application is resumed. Note: If the apertures of the joining piece become clogged, Pack 4 contains one spare joining piece and two additional application cannulas.

Rx Only.

DO NOT REUSE OR RESTERILIZE.

NOT MADE WITH NATURAL RUBBER LATEX.

STERILE AND NON-PYROGENIC IN UNOPENED AND UNDAMAGED PACKAGE.

DISPOSE OF CONTAMINATED AND SHARP COMPONENTS PROPERLY.

Baxter, Duploject, Fibrinotherm and Tisseel are registered trademarks of Baxter International Inc.

Manufactured for:

Baxter Healthcare Corporation

One Baxter Parkway

Deerfield, IL 60015 USA

1-888-229-0001

Made in Belgium

Rev. Date: 2017-05-17

DUPLOJET IFU Label

DUPLOJECT
Fibrin Sealant Preparation and Application System

Indications for Use:

For the preparation and application of TISSEEL [Fibrin Sealant] kit.

Note: See TISSEEL Fibrin Sealant package insert for additional preparation and application instructions.

1.0 Reconstitution Instructions for the Circulating Nurse (Using the FIBRINOTHERM Heating and Stirring Device)

- 1.1 Turn on the warming unit of the FIBRINOTHERM device (amber switch).
- 1.2 Open TISSEEL [Fibrin Sealant] kit and place all vials into the appropriately sized heating wells of the FIBRINOTHERM device.
- 1.3 Place the Sealer Protein vial into the large magnetic stirring well fitted with the appropriately sized adapter ring (if necessary). Do not activate the stirring mechanism at this time. The indicator light will remain lit until a temperature of 37°C (98.6°F) is reached. Allow several minutes for proper warming.
- 1.4 Open Pack A of the DUPLOJECT Preparation and Application System and assemble the blue-scaled and black-scaled syringes with the needles provided.
- 1.5 Remove the flip-off caps from the blue-capped Sealer Protein Concentrate and Fibrinolysis Inhibitor Solution vials and wipe with a non-iodine based disinfectant.
- 1.6 Using the blue-scaled syringe, withdraw the Fibrinolysis Inhibitor Solution from the vial, tilting the vial slightly to facilitate removal of all solution. Inject the Fibrinolysis Inhibitor Solution into the Sealer Protein Concentrate vial. Gently swirl the vial to ensure that the freeze-dried material is fully soaked. Note: Do not invert or inject air into vials.
- 1.7 Place the Sealer Protein Concentrate vial in the stirring well and activate the stirring mechanism of the FIBRINOTHERM device (green switch). Reconstitution is complete when no undissolved particles are visible. If Sealer Protein is not fully dissolved after 20 minutes, discard and prepare a fresh kit. Continue with steps 8 through 11.
- 1.8 Remove the flip-off caps from the black-capped Thrombin and Calcium Chloride Solution vials and wipe with a non-iodine based disinfectant.
- 1.9 Using the black-scaled syringe, withdraw the Calcium Chloride Solution from the vial, tilting the vial slightly to facilitate removal of all solution. Inject the Calcium Chloride Solution into the Thrombin vial. Note: Do not invert or inject air into vials.
- 1.10 Swirl contents of the Thrombin vial briefly and return it to an appropriately sized heating well in the FIBRINOTHERM device.
- 1.11 Leave the Sealer Protein and Thrombin vials in the FIBRINOTHERM device until

the solutions are ready to be passed into the sterile field.

2.0 Preparation Instructions for the Scrub Nurse

2.1 Open Pack B of the DUPLOJECT Preparation and Application System into the sterile field.

2.2 Open Packs 1, 2, and 3.

2.3 Assemble the blue-scaled and black-scaled syringes with the needles provided.

Note: For 5 mL DUPLOJECT systems, attach the larger needle to the blue-scaled syringe.

2.4 While the Circulating Nurse holds the vial slightly tilted, insert needle into vial (bevel side down) and withdraw all of the Sealer Protein Solution into the blue-scaled syringe using slow, constant aspiration. Discard needle in sharps container.

2.5 While the Circulating Nurse holds the vial slightly tilted, insert needle into vial (bevel side down) and withdraw all of the Thrombin Solution into the black-scaled syringe using slow, constant aspiration. Discard needle in sharps container.

2.6 Remove any air bubbles from the syringes and ensure both syringes contain the same volume.

2.7 Snap the filled syringes into the two-syringe clip with the flanges in an up/down position, as illustrated below.

2.8 Attach the joining piece to the syringe nozzles, ensuring that both are firmly seated. Secure the joining piece by fastening the retaining strap to the double syringe clip. Note: Align the syringe nozzles toward the middle for proper attachment.

2.9 Fit one of the application needles onto the joining piece. A second application needle is provided in Pack 3 as a spare.

2.10 DUPLOJECT Applicator is ready for use.

2.11 If application of TISSEEL [Fibrin Sealant] is interrupted, replace the needle immediately before application is resumed. Note: If the apertures of the joining piece become clogged, Pack 4 contains one spare joining piece and two additional application needles.

Rx Only

DO NOT REUSE OR RESTERILIZE.

NOT MADE WITH NATURAL RUBBER LATEX.

STERILE AND NON-PYROGENIC IN UNOPENED AND UNDAMAGED PACKAGE.

DISPOSE OF CONTAMINATED AND SHARP COMPONENTS PROPERLY.

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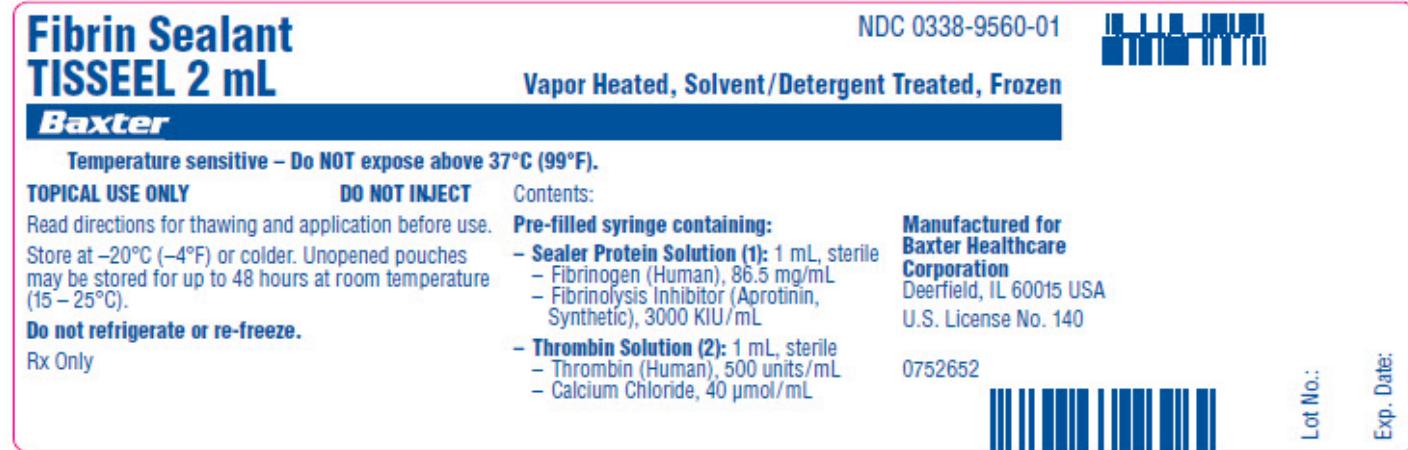
One Baxter Parkway

Deerfield, IL 60015 USA

1-888-229-0001

Made in Belgium

Rev. Date: 2017-05-17



2 mL PRIMA Pouch Label

Fibrin Sealant TISSEEL 2 mL

NDC 0338-9560-01

Vapor Heated, Solvent/Detergent Treated, Frozen

Baxter Logo

Temperature sensitive - Do NOT expose above 37°C (99°F).

TOPICAL USE ONLY DO NOT INJECT

Read directions for thawing and application before use.

Store at -20°C (-4°F) or colder. Unopened pouches may be stored for up to 48 hours at room temperature (15 – 25°C).

Do not refrigerate or re-freeze.

Rx Only

Contents:

Pre-filled syringe containing:

- **Sealer Protein Solution (1):** 1 mL, sterile
- Fibrinogen (Human), 86.5 mg/mL
- Fibrinolysis Inhibitor (Aprotinin, Synthetic), 3000 KIU/mL

- **Thrombin Solution (2):** 1 mL, sterile
- Thrombin (Human), 500 units/mL
- Calcium Chloride, 40 µmol/mL

**Manufactured for
Baxter Healthcare**

Corporation
Deerfield, IL 60015 USA
U.S. License No. 140
0752652

Barcode

Lot No.:
Exp. Date:





2 mL PRIMA Carton Label

0752653

**Fibrin Sealant
TISSEEL
Vapor Heated, Solvent/
Detergent Treated, Frozen
TOPICAL USE ONLY**

NDC 0338-9560-01

Barcode

**Barcode
3 03389 56001 2**

2 mL

**Fibrin Sealant
TISSEEL
Vapor Heated, Solvent/Detergent Treated, Frozen
with Pre-filled PRIMA Syringe**

TOPICAL USE ONLY

Contents:

Pre-filled syringe containing:

Sealer Protein Solution: 1 mL, sterile

- Fibrinogen (Human), 86.5 mg/mL
- Fibrinolysis Inhibitor (Aprotinin, Synthetic),
3000 KIU/mL

Thrombin Solution: 1 mL, sterile

- Thrombin (Human), 500 units/mL
- Calcium Chloride, 40 µmol /mL

The risks and benefits of this product should
be discussed with the patient.

Rx Only

DO NOT INJECT

Store at -20°C (-4°F) or colder.

Read enclosed directions for thawing and
application before use.

Unopened pouches may be stored for up to
48 hours at room temperature (15 – 25°C).

Do not refrigerate or re-freeze.

Not made with natural rubber latex
For Single Patient Use Only.

NDC 0338-9560-01

Fibrin Sealant

TISSEEL

Vapor Heated, Solvent/Detergent Treated, Frozen

TOPICAL USE ONLY

2 mL

Fibrin Sealant

TISSEEL

Vapor Heated, Solvent/

Detergent Treated, Frozen

TOPICAL USE ONLY

2 mL

Fibrin Sealant

TISSEEL

Vapor Heated, Solvent/

Detergent Treated, Frozen

TOPICAL USE ONLY

2 mL

Fibrin Sealant

TISSEEL

Vapor Heated, Solvent/Detergent Treated, Frozen

with Pre-filled PRIMA Syringe

TOPICAL USE ONLY

DO NOT INJECT

U.S. Pat. No.: 5,962,405

Manufactured for
Baxter Healthcare Corporation

Deerfield IL, 60015 USA

1-888-229-0001

U.S. License No. 140

Made in Austria

Reorder Number: 1506078

Baxter and Tisseel are trademarks of
Baxter International Inc.**2 mL****GTIN (01) 00303389560012**

EXPIRY (17)

LOT (10)

SERIAL (21)

Baxter Logo**Fibrin Sealant****TISSEEL****Vapor Heated, Solvent/
Detergent Treated, Frozen
TOPICAL USE ONLY****2 mL****Fibrin Sealant
TISSEEL 4 mL**

NDC 0338-9564-01



Vapor Heated, Solvent/Detergent Treated, Frozen

Baxter

Temperature sensitive – Do NOT expose above 37°C (99°F).

TOPICAL USE ONLY**DO NOT INJECT**

Read directions for thawing and application before use.

Store at -20°C (-4°F) or colder. Unopened pouches may be stored for up to 48 hours at room temperature (15 – 25°C).

Do not refrigerate or re-freeze.

Rx Only

Contents:

Pre-filled syringe containing:

- Sealer Protein Solution (1): 2 mL, sterile
 - Fibrinogen (Human), 86.5 mg/mL
 - Fibrinolysis Inhibitor (Aprotinin, Synthetic), 3000 KIU/mL
- Thrombin Solution (2): 2 mL, sterile
 - Thrombin (Human), 500 units/mL
 - Calcium Chloride, 40 µmol/mL

Manufactured for
Baxter Healthcare
Corporation
Deerfield, IL 60015 USA
U.S. License No. 140

0752656



Lot No.:

Exp. Date:

4 mL PRIMA Pouch Label**Fibrin Sealant
TISSEEL 4 mL**

NDC 0338-9564-01

Barcode

Vapor Heated, Solvent/Detergent Treated, Frozen

Baxter Logo

Temperature sensitive - Do NOT expose above 37°C (99°F).

TOPICAL USE ONLY DO NOT INJECT

Read directions for thawing and application before use.

Store at -20°C (-4°F) or colder. Unopened pouches may be stored for up to 48 hours at room temperature (15 – 25°C).

Do not refrigerate or re-freeze.

Rx Only

Contents:

Pre-filled syringe containing:

- **Sealer Protein Solution (1):** 2 mL, sterile
 - Fibrinogen (Human), 86.5 mg/mL
 - Fibrinolysis Inhibitor (Aprotinin, Synthetic), 3000 KIU/mL
-
- **Thrombin Solution (2):** 2 mL, sterile
 - Thrombin (Human), 500 units/mL
 - Calcium Chloride, 40 µmol/mL

Barcode

Manufactured for

Baxter Healthcare

Corporation

Deerfield, IL 60015 USA

U.S. License No. 140

0752656

Barcode

Lot No.:

Exp. Date:





4 mL PRIMA Carton Label

0752657

Barcode
3 03389 56401 0

**Fibrin Sealant
TISSEEL
Vapor Heated, Solvent/Detergent Treated, Frozen
TOPICAL USE ONLY
with Pre-filled PRIMA Syringe**

NDC 0338-9564-01

TOPICAL USE ONLY

Contents:

Pre-filled syringe containing:

Sealer Protein Solution: 2 mL, sterile

- Fibrinogen (Human), 86.5 mg/mL

- Fibrinolysis Inhibitor (Aprotinin, Synthetic),

3000 KIU/mL

Thrombin Solution: 2 mL, sterile

- Thrombin (Human), 500 units/mL

- Calcium Chloride, 40 µmol /mL

The risks and benefits of this product should be discussed with the patient.

Rx Only

DO NOT INJECT

Store at -20°C (-4°F) or colder.

Read enclosed directions for thawing and application before use.

Unopened pouches may be stored for up to 48 hours at room temperature (15 – 25°C).

Do not refrigerate or re-freeze.

Not made with natural rubber latex
For Single Patient Use Only.

Fibrin Sealant

TISSEEL

**Vapor Heated, Solvent/
Detergent Treated, Frozen
TOPICAL USE ONLY**

4 mL

Fibrin Sealant

TISSEEL

**Vapor Heated, Solvent/
Detergent Treated, Frozen
TOPICAL USE ONLY**

4 mL

Fibrin Sealant

TISSEEL

**Vapor Heated, Solvent/
Detergent Treated, Frozen
TOPICAL USE ONLY**

4 mL

Fibrin Sealant

TISSEEL

**Vapor Heated, Solvent/Detergent Treated, Frozen
with Pre-filled PRIMA Syringe**

TOPICAL USE ONLY

DO NOT INJECT

U.S. Pat. No.: 5,962,405

Manufactured for
Baxter Healthcare Corporation

Deerfield IL, 60015 USA

1-888-229-0001

U.S. License No. 140

Made in Austria

Reorder Number: 1506079

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Baxter International Inc.

4 mL

Baxter Logo

GTIN (01) 00303389564010

LOT (10)

EXPIRY (17)

SERIAL (21)

Fibrin Sealant TISSEEL 10 mL

NDC 0338-9568-01



Vapor Heated, Solvent/Detergent Treated, Frozen

Baxter

Temperature sensitive – Do NOT expose above 37°C (99°F).

TOPICAL USE ONLY

DO NOT INJECT

Read directions for thawing and application before use.

Store at -20°C (-4°F) or colder. Unopened pouches may be stored for up to 48 hours at room temperature (15 – 25°C).

Do not refrigerate or re-freeze.

Rx Only

Contents:

Pre-filled syringe containing:

- **Sealer Protein Solution (1):** 5 mL, sterile
 - Fibrinogen (Human), 86.5 mg/mL
 - Fibrinolysis Inhibitor (Aprotinin, Synthetic), 3000 KIU/mL
- **Thrombin Solution (2):** 5 mL, sterile
 - Thrombin (Human), 500 units/mL
 - Calcium Chloride, 40 µmol/mL

**Manufactured for
Baxter Healthcare
Corporation**

Deerfield, IL 60015 USA

U.S. License No. 140

0752660



Lot No.:

Exp. Date:

10 mL PRIMA Pouch Label

Fibrin Sealant TISSEEL 10 mL

NDC 0338-9568-01

Vapor Heated, Solvent/Detergent Treated, Frozen

Baxter Logo

Temperature sensitive – Do NOT expose above 37°C (99°F).

TOPICAL USE ONLY DO NOT INJECT

Read directions for thawing and application before use.

Store at -20°C (-4°F) or colder. Unopened pouches may be stored for up to 48 hours at room temperature (15 – 25°C).

Do not refrigerate or re-freeze.

Rx Only

Contents:

Pre-filled syringe containing:

- **Sealer Protein Solution (1):** 5 mL, sterile
 - Fibrinogen (Human), 86.5 mg/mL
 - Fibrinolysis Inhibitor (Aprotinin, Synthetic), 3000 KIU/mL
 - **Thrombin Solution (2):** 5 mL, sterile
 - Thrombin (Human), 500 units/mL
 - Calcium Chloride, 40 μmol/mL

Manufactured for Baxter Healthcare Corporation

Deerfield, IL 60015 USA
U.S. License No. 140

0752660

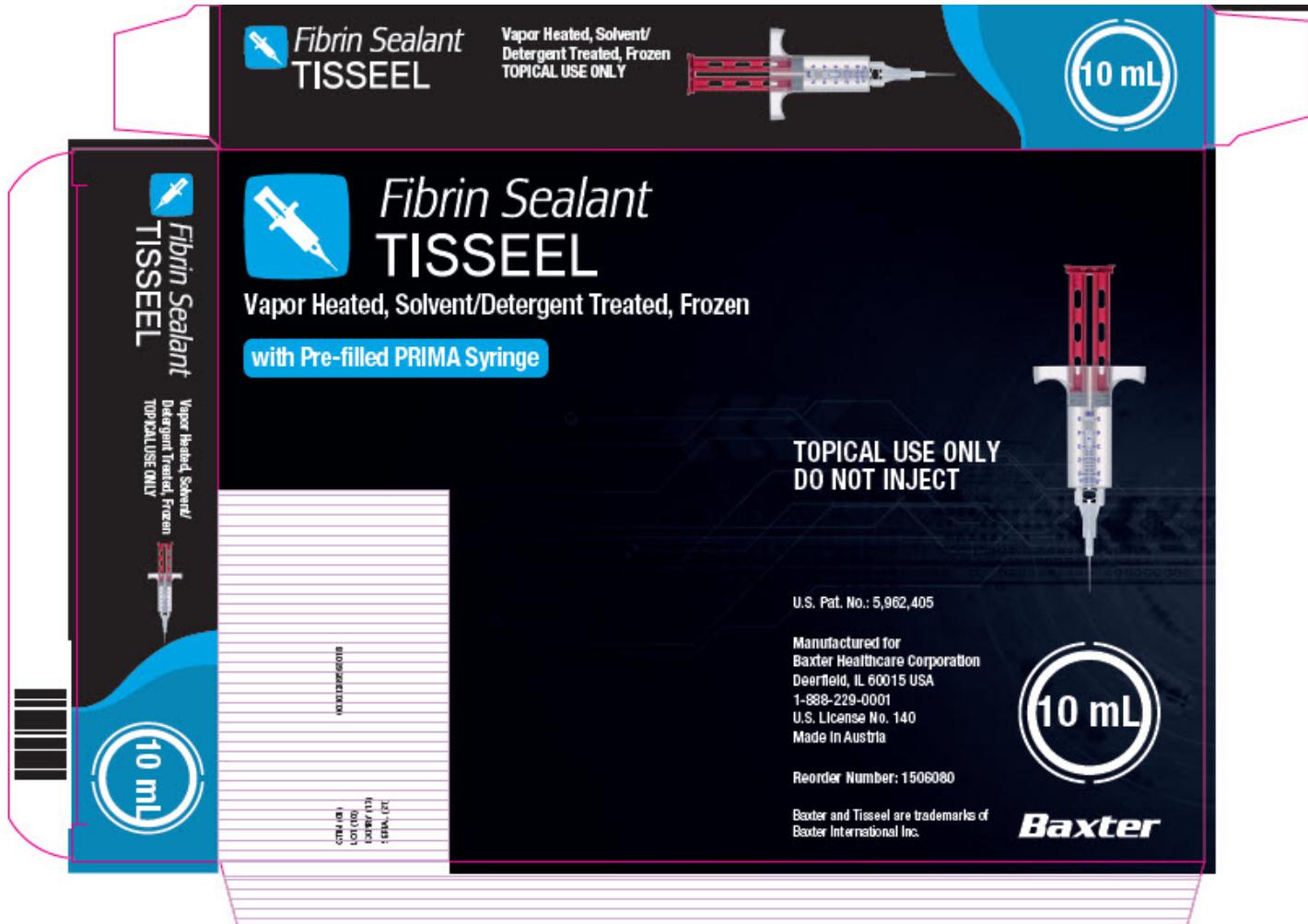
Barcode

Barcode

Lot No.:

Exp. Date:





0752661

Barcode
3 03389 56801 8

**Fibrin Sealant
TISSEEL
Vapor Heated, Solvent/
Detergent Treated, Frozen
TOPICAL USE ONLY**

NDC 0338-9568-01

Barcode

10 mL

**Fibrin Sealant
TISSEEL
Vapor Heated, Solvent/
Detergent Treated, Frozen
TOPICAL USE ONLY**

10 mL

Fibrin Sealant

TISSEEL

**Vapor Heated, Solvent/Detergent Treated, Frozen
with Pre-filled PRIMA Syringe**

TOPICAL USE ONLY

Contents:

Pre-filled syringe containing:

Sealer Protein Solution: 5 mL, sterile

- Fibrinogen (Human), 86.5 mg/mL

- Fibrinolysis Inhibitor (Aprotinin, Synthetic),

3000 KIU/mL

Thrombin Solution: 5 mL, sterile

- Thrombin (Human), 500 units/mL

- Calcium Chloride, 40 µmol /mL

The risks and benefits of this product should be discussed with the patient.

Rx Only

DO NOT INJECT

Store at -20°C (-4°F) or colder.

Read enclosed directions for thawing and application before use.

Unopened pouches may be stored for up to 48 hours at room temperature (15 – 25°C).

Do not refrigerate or re-freeze.

Not made with natural rubber latex

For Single Patient Use Only

Fibrin Sealant**TISSEEL**

**Vapor Heated, Solvent/
Detergent Treated, Frozen**
TOPICAL USE ONLY

10 mL

Fibrin Sealant**TISSEEL**

**Vapor Heated, Solvent/
Detergent Treated, Frozen**
TOPICAL USE ONLY

10 mL

Barcode**Fibrin Sealant****TISSEEL**

Vapor Heated, Solvent/Detergent Treated, Frozen

with Pre-filled PRIMA Syringe

TOPICAL USE ONLY

DO NOT INJECT

U.S. Pat. No.: 5,962,405

Manufactured for
Baxter Healthcare Corporation

Deerfield IL, 60015 USA

1-888-229-0001

U.S. License No. 140

Made in Austria

Reorder Number: 1506080

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Baxter International Inc.

10 mL

Baxter Logo

GTIN (01) 00303389568018

EXPIRY (17)

LOT (10)

SERIAL (21)



Baxter

NDC 0338-4211-04



**Fibrin Sealant
TISSEEL 4 mL**

Vapor Heated, Solvent/Detergent Treated, Kit



Contents:

**Sealer Protein Concentrate (Human), Vapor Heated,
Solvent/Detergent Treated, freeze dried, sterile, for
2 mL of Sealer Protein Solution 86.5 mg/mL**

**Fibrinolysis Inhibitor Solution (Synthetic), sterile,
3000 KIU of Aprotinin/mL**

**Thrombin (Human), Vapor Heated, Solvent/Detergent
Treated, freeze dried, sterile, for 2 mL of Thrombin
Solution 500 units/mL**

Calcium Chloride Solution, sterile, 40 µmol/mL

The risks and benefits of this product should be discussed
with the patient.

Read enclosed directions for reconstitution and application
before use.

Use within four hours of reconstitution.

NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only



GTIN (01)
LOT (10)
EXPIRY (17)
SERIAL (21)
NDC 0338-4211-04

**Fibrin Sealant
TISSEEL 4 mL
Vapor Heated, Solvent/Detergent Treated, Kit**

Baxter logo

NDC 0338-4211-04
barcode

**Fibrin Sealant
TISSEEL 4 mL
Vapor Heated, Solvent/Detergent Treated, Kit**

Contents:

Sealer Protein Concentrate (Human), Vapor Heated, Solvent /Detergent Treated, freeze dried, sterile, for 2 mL of Sealer Protein Solution

Fibrinolysis Inhibitor Solution (Synthetic), sterile, 3000 KIU of Aprotinin /mL

Thrombin (Human), Vapor Heated, Solvent /Detergent Treated, freeze dried, sterile, for 2 mL of Thrombin Solution 500 units/mL

Calcium Chloride Solution, sterile, 40 µmol /mL

The risks and benefits of this product should be discussed with the patient.

Read enclosed directions for reconstitution and application before use.

Use within four hours of reconstitution.

NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only

Manufactured for Baxter Healthcare Corporation

Deerfield, IL 60015 USA

U.S. License No. 140

Made in Austria

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U.S. Pat. No.: 5,962,405

Barcode

0752678

Fibrin Sealant

TISSEEL 4 mL

**Vapor Heated, Solvent/
Detergent Treated, Kit**

4 mL

Store at 2°C to 25°C (36°F to 77°F).

Avoid freezing.



Baxter

NDC 0338-4212-10



**Fibrin Sealant
TISSEEL 10 mL**



Vapor Heated, Solvent / Detergent Treated, Kit

Contents:

**Sealer Protein Concentrate (Human), Vapor Heated,
Solvent/Detergent Treated, freeze dried, sterile, for
5 mL of Sealer Protein Solution 86.5 mg/mL.**

**Fibrinolysis Inhibitor Solution (Synthetic), sterile,
3000 KIU of Aprotinin/ mL**

**Thrombin (Human), Vapor Heated, Solvent/Detergent
Treated, freeze dried, sterile, for 5 mL of Thrombin
Solution 500 units/mL.**

Calcium Chloride Solution, sterile, 40 µmol/ mL.

The risks and benefits of this product should be discussed
with the patient.

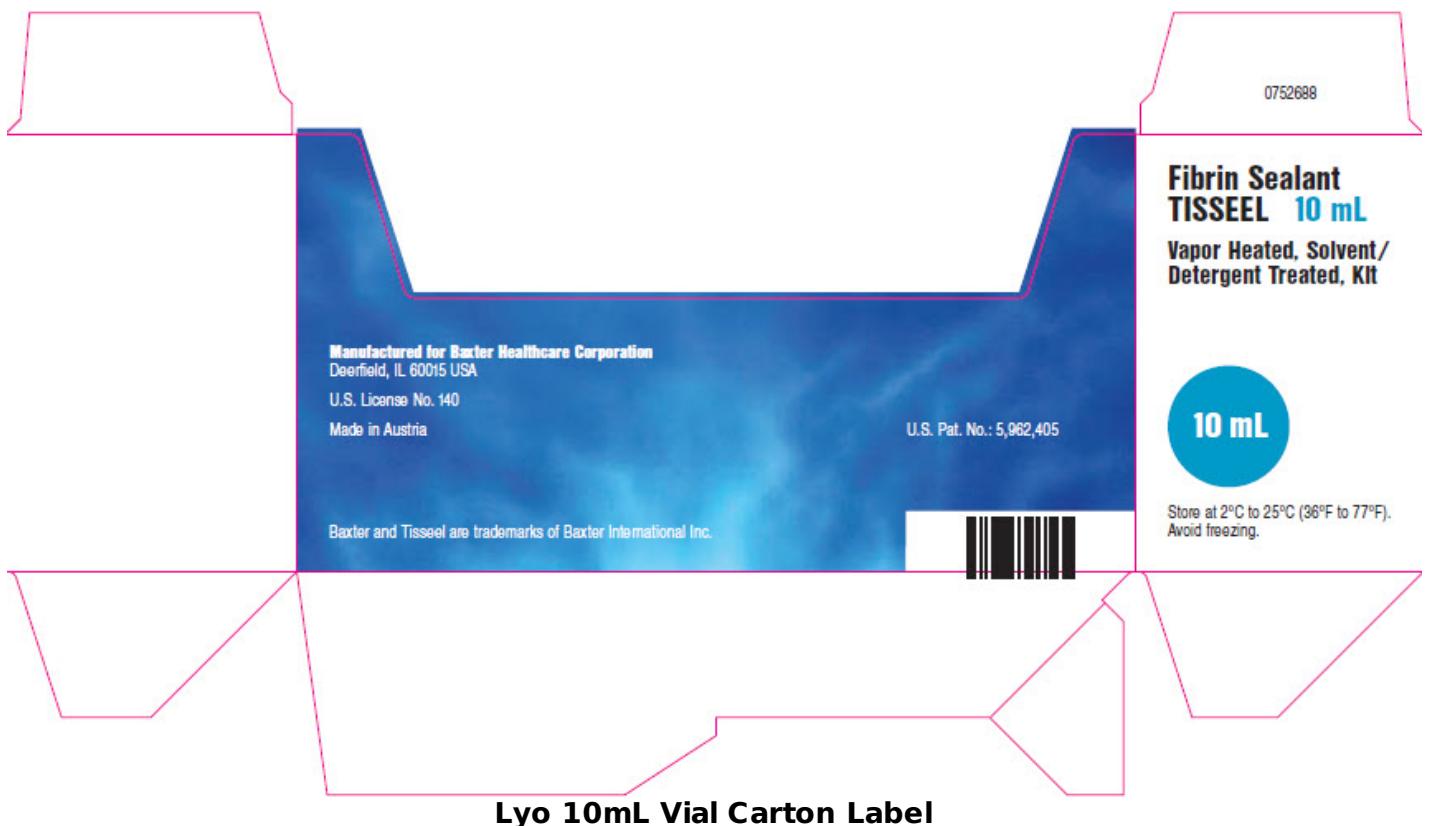
Read enclosed directions for reconstitution and application
before use.

Use within four hours of reconstitution.

NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only



GTIN (01)

LOT (10)

EXPIRY (17)

SERIAL (21)

NDC 0338-4212-10

**Fibrin Sealant
TISSEEL 10 mL
Vapor Heated, Solvent/Detergent Treated, Kit**

Baxter Logo

NDC 0338-4212-10

barcode

**Fibrin Sealant
TISSEEL 10 mL
Vapor Heated, Solvent/Detergent Treated, Kit**

Contents:

Sealer Protein Concentrate (Human), Vapor Heated, Solvent /Detergent Treated, freeze dried, sterile, for 5 mL of Sealer Protein Solution 86.5 mg/mL

Fibrinolysis Inhibitor Solution (Synthetic), sterile, 3000 KIU of Aprotinin /mL

Thrombin (Human), Vapor Heated, Solvent /Detergent Treated, freeze dried, sterile, for 5 mL of Thrombin

Solution 500 units/mL

Calcium Chloride Solution, sterile, 40 µmol /mL

The risks and benefits of this product should be discussed with the patient.

Read enclosed directions for reconstitution and application before use.

Use within four hours of reconstitution.

NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only

Manufactured for Baxter Healthcare Corporation

Deerfield, IL 60015 USA

U.S. License No. 140

Made in Austria

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U.S. Pat. No.: 5,962,405

Barcode

0752688

Fibrin Sealant

TISSEEL 10 mL

**Vapor Heated, Solvent/
Detergent Treated, Kit**

10 mL

Store at 2°C to 25°C (36°F to 77°F).

Avoid freezing.

Baxter

NDC 0338-4302-04

**Fibrin Sealant
TISSEEL 4 mL**

Vapor Heated, Solvent/Detergent Treated, Kit



**TOPICAL USE ONLY
DO NOT INJECT**

Baxter

NDC 0338-4302-04

**Fibrin Sealant
TISSEEL 4 mL**

Vapor Heated, Solvent/Detergent Treated, Kit



Manufactured for Baxter Healthcare Corporation
Deerfield, IL 60015 USA

U.S. License No. 140

Made in Austria

Reorder Number: 1504515

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U.S. Pat. No.: 5,962,405



NDC 0338-4302-04



**Fibrin Sealant
TISSEEL 4 mL**



Vapor Heated, Solvent/Detergent Treated, Kit

Contents:

Sealer Protein Concentrate (Human), Vapor Heated, Solvent/Detergent Treated, freeze dried, sterile, for 2 mL of Sealer Protein Solution 86.5 mg/mL
Fibrinolysis Inhibitor Solution (Synthetic), sterile, 3000 KIU of Aprotinin / mL
Thrombin (Human), Vapor Heated, Solvent/Detergent Treated, freeze dried, sterile, for 2 mL of Thrombin Solution 500 units/ mL
Calcium Chloride Solution, sterile, 40 µmol/mL

Also includes: DUPLOJECT Fibrin Sealant Preparation and Application System 2 mL / 4 mL

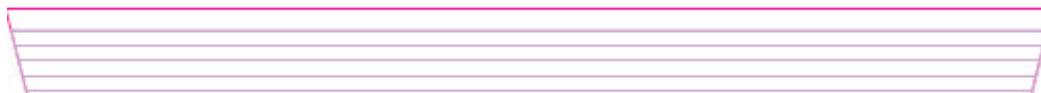
The risks and benefits of this product should be discussed with the patient. Read enclosed directions for reconstitution and application before use. Use within four hours of reconstitution.

NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only

0752680



Lyo 4mL Sleeve Carton Label

Baxter Logo

NDC 0338-4302-04

**Fibrin Sealant
TISSEEL 4 mL
Vapor Heated, Solvent/Detergent Treated, Kit**

**Manufactured for Baxter Healthcare Corporation
Deerfield, IL 60015 USA**

U.S. License No. 140

Made in Austria

Reorder Number: 1504515

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U.S. Pat. No.: 5,962,405

Baxter Logo

NDC 0338-4302-04

Fibrin Sealant

TISSEEL 4 mL

Vapor Heated, Solvent/Detergent Treated, Kit

TOPICAL USE ONLY

DO NOT INJECT

Baxter Logo

NDC 0338-4302-04

barcode

Fibrin Sealant

TISSEEL 4 mL

Vapor Heated, Solvent/Detergent Treated, Kit

Contents:

Sealer Protein Concentrate (Human), Vapor Heated,
Solvent /Detergent Treated, freeze dried, sterile, for
2 mL of Sealer Protein Solution 86.5 mg/mL

Fibrinolysis Inhibitor Solution (Synthetic), sterile,
3000 KIU of Aprotinin /mL

Thrombin (Human), Vapor Heated, Solvent /Detergent
Treated, freeze dried, sterile, for 2 mL of Thrombin Solution
500 units/mL

Calcium Chloride Solution, sterile, 40 μ mol /mL

The risks and benefits of this product should be discussed
with the patient. Read enclosed directions for
reconstitution and application before use. Use within four
hours of reconstitution.

NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only

**Also includes: DUPLOJECT Fibrin Sealant Preparation and Application System 2
mL / 4 mL**

0752680

Barcode

Barcode

N3 0338-4302-04 4



Vapor Heated, Solvent / Detergent Treated, Kit
TISSUEEL 10 mL
Fibrin Sealant

NDC 0338-4303-10

Baxter



Vapor Heated, Solvent / Detergent Treated, Kit
TISSUEEL 10 mL
Fibrin Sealant

NDC 0338-4303-10

Baxter



NDC 0338-4303-10

**Fibrin Sealant
TISSEEL 10 mL****Vapor Heated, Solvent / Detergent Treated, Kit****Contents:****Sealer Protein Concentrate (Human), Vapor Heated, Solvent/Detergent Treated, freeze dried, sterile, for 5 mL of Sealer Protein Solution 86.5 mg/mL.****Fibrinolytic Inhibitor Solution (Synthetic), sterile, 3000 KIU of Aprotinin/mL.****Thrombin (Human), Vapor Heated, Solvent/Detergent Treated, freeze dried, sterile, for 5 mL of Thrombin Solution 500 units/mL.****Calcium Chloride Solution, sterile, 40 µmol/mL.**

The risks and benefits of this product should be discussed with the patient. Read enclosed directions for reconstitution and application before use. Use within four hours of reconstitution.

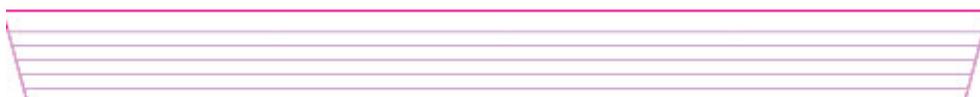
NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only

Also Includes: DUPLOJECT Fibrin Sealant Preparation and Application System 10 mL

0752690

**Lyo 10mL Sleeve Carton Label****Baxter Logo**

NDC 0338-4303-10

Fibrin Sealant**TISSEEL 10 mL****Vapor Heated, Solvent/Detergent Treated, Kit****Manufactured for Baxter Healthcare Corporation**

Deerfield, IL 60015 USA

U.S. License No. 140

Made in Austria

Reorder Number: 1504516

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U.S. Pat. No.: 5,962,405

Baxter Logo

NDC 0338-4303-10

Fibrin Sealant

TISSEEL 10 mL

Vapor Heated, Solvent/Detergent Treated, Kit

TOPICAL USE ONLY

DO NOT INJECT

Baxter Logo

NDC 0338-4303-10

Barcode

Fibrin Sealant

TISSEEL 10 mL

Vapor Heated, Solvent/Detergent Treated, Kit

Contents:

Sealer Protein Concentrate (Human), Vapor Heated,
Solvent /Detergent Treated, freeze dried, sterile, for
5 mL of Sealer Protein Solution 86.5 mg/mL

Fibrinolysis Inhibitor Solution (Synthetic), sterile,
3000 KIU of Aprotinin /mL

Thrombin (Human), Vapor Heated, Solvent /Detergent
Treated, freeze dried, sterile, for 5 mL of Thrombin Solution
500 units/mL

Calcium Chloride Solution, sterile, 40 µmol /mL

The risks and benefits of this product should be discussed
with the patient. Read enclosed directions for
reconstitution and application before use. Use within four
hours of reconstitution.

NOT FOR INJECTION. For single use only.

Not made with natural rubber latex

Rx only

**Also includes: DUPLOJECT Fibrin Sealant Preparation and Application System
10 mL**

0752690

Barcode

Barcode

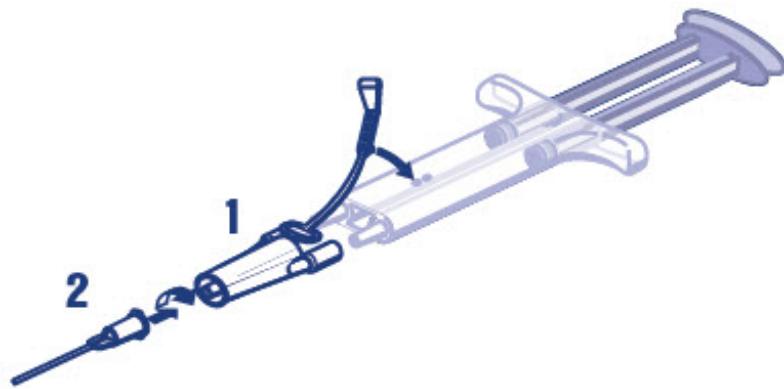
N3 0338-4303-10 2

Baxter

DUPLOJECT COMBI

Indication for Use:

For the application of TISSEEL Fibrin Sealant.



For single use only. Not made with natural rubber latex.
Do not reuse or resterilize. Sterilized using ethylene oxide.
Sterile and non-pyrogenic in unopened and undamaged package. Rx Only.

Manufactured for:
Baxter Healthcare Corporation
One Baxter Parkway
Deerfield, IL 60015 USA
1-888-229-0001

BE-90-01-044

Made in Belgium

Duploject Combi Label

Baxter Logo

Duploject Combi

Indication for Use:

For the application of TISSEEL Fibrin Sealant.

REF

Batch code

Use-by date

For single use only. Not made with natural rubber latex.

Do not reuse or resterilize. Sterilized using ethylene oxide.

Sterile and non-pyrogenic in unopened and undamaged package. Rx Only.

Manufactured for:

Baxter Healthcare Corporation

One Baxter Parkway

Deerfield, IL 60015 USA

1-888-229-0001

Made in Belgium

BE-90-01-044

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin kit

Product Information

Product Type	PLASMA DERIVATIVE	Item Code (Source)	NDC:0338-4210
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Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-4210-02	1 in 1 CARTON; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Quantity of Parts

Part #	Package Quantity	Total Product Quantity
Part 1	1 VIAL, GLASS	1 mL
Part 2	1 VIAL, GLASS	1 mL
Part 3	1 VIAL, GLASS	1 mL
Part 4	1 VIAL, GLASS	1 mL

Part 1 of 4

SEALER PROTEIN CONCENTRATE HUMAN

fibrinogen human powder, for solution

Product Information

Item Code (Source)	NDC:0338-7112
Route of Administration	TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - UNII:N94833051K)	FIBRINOGEN HUMAN	86.5 mg in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	
NIACINAMIDE (UNII: 25X51I8RD4)	
POLYSORBATE 80 (UNII: 6OZP39ZG8H)	
TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7112-01	1 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 2 of 4

HUMAN THROMBIN

human thrombin powder, for solution

Product Information

Item Code (Source)	NDC:0338-7332
Route of Administration	TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [iU] in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
SODIUM CHLORIDE (UNII: 451W47IQ8X)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7332-01	1 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 3 of 4

FIBRINOLYSIS INHIBITOR SOLUTION

aprotinin liquid

Product Information

Item Code (Source)	NDC:0338-7201
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name	Strength
APROTININ (UNII: 04XPW8C0FL)	3000 [iU] in 1 mL
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-	1 mL in 1 VIAL, GLASS; Type 0: Not a Combination		

#	7201-01	Product		
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Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 4 of 4

CALCIUM CHLORIDE SOLUTION

calcium chloride liquid

Product Information

Item Code (Source)	NDC:0338-7401
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name	Strength
CALCIUM CHLORIDE (UNII: M4I0D6VV5M)	40 umol in 1 mL
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7401-01	1 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin kit

Product Information

Product Type

PLASMA DERIVATIVE

Item Code (Source)

NDC:0338-4211

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-4211-04	1 in 1 CARTON; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Quantity of Parts

Part #	Package Quantity	Total Product Quantity
Part 1	1 VIAL, GLASS	2 mL
Part 2	1 VIAL, GLASS	2 mL
Part 3	1 VIAL, GLASS	2 mL
Part 4	1 VIAL, GLASS	2 mL

Part 1 of 4

SEALER PROTEIN CONCENTRATE HUMAN

fibrinogen human powder, for solution

Product Information

Item Code (Source)

NDC:0338-7112

Route of Administration

TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - UNII:N94833051K)	FIBRINOGEN HUMAN	86.5 mg in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	
NIACINAMIDE (UNII: 25X51I8RD4)	
POLYSORBATE 80 (UNII: 6OZP39ZG8H)	
TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7112-02	2 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 2 of 4

HUMAN THROMBIN

human thrombin powder, for solution

Product Information

Item Code (Source)	NDC:0338-7332
Route of Administration	TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [iU] in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
SODIUM CHLORIDE (UNII: 451W47IQ8X)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7332-02	2 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing	Application Number or Monograph	Marketing Start	Marketing End
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Category	Citation	Date	Date
BLA	BLA103980	05/01/1998	

Part 3 of 4

FIBRINOLYSIS INHIBITOR SOLUTION

aprotinin liquid

Product Information

Item Code (Source)	NDC:0338-7201
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name	Strength
APROTININ (UNII: 04XPW8C0FL)	3000 [iU] in 1 mL
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7201-02	2 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 4 of 4

CALCIUM CHLORIDE SOLUTION

calcium chloride liquid

Product Information

Item Code (Source)	NDC:0338-7401
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name		Strength
CALCIUM CHLORIDE (UNII: M4I0D6VV5M)		40 umol in 1 mL
WATER (UNII: 059QF0KO0R)		

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7401-02	2 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin kit

Product Information

Product Type	PLASMA DERIVATIVE	Item Code (Source)	NDC:0338-4212
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Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-4212-10	1 in 1 CARTON; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Quantity of Parts

Part #	Package Quantity	Total Product Quantity
Part 1	1 VIAL, GLASS	5 mL
Part 2	1 VIAL, GLASS	5 mL
Part 3	1 VIAL, GLASS	5 mL
Part 4	1 VIAL, GLASS	5 mL

Part 1 of 4

SEALER PROTEIN CONCENTRATE HUMAN

fibrinogen human powder, for solution

Product Information

Item Code (Source)	NDC:0338-7112
Route of Administration	TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - UNII:N94833051K)	FIBRINOGEN HUMAN	86.5 mg in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	
NIACINAMIDE (UNII: 25X51I8RD4)	
POLYSORBATE 80 (UNII: 6OZP39ZG8H)	
TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7112-05	5 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 2 of 4

HUMAN THROMBIN

human thrombin powder, for solution

Product Information

Item Code (Source)	NDC:0338-7332			
Route of Administration	TOPICAL			
Active Ingredient/Active Moiety				
Ingredient Name	Basis of Strength	Strength		
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [iU] in 1 mL		
Inactive Ingredients				
Ingredient Name	Strength			
ALBUMIN HUMAN (UNII: ZIF514RVZR)				
SODIUM CHLORIDE (UNII: 451W47IQ8X)				
Packaging				
#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7332-05	5 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		
Marketing Information				
Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date	
BLA	BLA103980	05/01/1998		
Part 3 of 4				
FIBRINOLYSIS INHIBITOR SOLUTION				
aprotinin liquid				
Product Information				
Item Code (Source)	NDC:0338-7201			
Route of Administration	TOPICAL			
Inactive Ingredients				
Ingredient Name	Strength			
APROTININ (UNII: 04XPW8C0FL)	3000 [iU] in 1 mL			
WATER (UNII: 059QF0KO0R)				

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7201-05	5 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 4 of 4

CALCIUM CHLORIDE SOLUTION

calcium chloride liquid

Product Information

Item Code (Source)	NDC:0338-7401
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name	Strength
CALCIUM CHLORIDE (UNII: M4I0D6VV5M)	40 umol in 1 mL
WATER (UNII: 059QF0KO0R)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7401-05	5 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date

BLA

BLA103980

05/01/1998

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin kit

Product Information

Product Type	PLASMA DERIVATIVE	Item Code (Source)	NDC:0338-4301
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Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-4301-02	1 in 1 CARTON; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Quantity of Parts

Part #	Package Quantity	Total Product Quantity
Part 1	1 VIAL, GLASS	1 mL
Part 2	1 VIAL, GLASS	1 mL
Part 3	1 VIAL, GLASS	1 mL
Part 4	1 VIAL, GLASS	1 mL

Part 1 of 4

SEALER PROTEIN CONCENTRATE HUMAN

fibrinogen human powder, for solution

Product Information

Item Code (Source)	NDC:0338-7112
Route of Administration	TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - UNII:N94833051K)	FIBRINOGEN HUMAN	86.5 mg in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	

NIACINAMIDE (UNII: 25X51I8RD4)

POLYSORBATE 80 (UNII: 6OZP39ZG8H)

TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7112-01	1 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 2 of 4

HUMAN THROMBIN

human thrombin powder, for solution

Product Information

Item Code (Source)	NDC:0338-7332		
Route of Administration	TOPICAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [iU] in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
SODIUM CHLORIDE (UNII: 451W47IQ8X)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7332-01	1 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 3 of 4

FIBRINOLYSIS INHIBITOR SOLUTION

aprotinin liquid

Product Information

Item Code (Source)	NDC:0338-7201
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name	Strength
APROTININ (UNII: 04XPW8C0FL)	3000 [iU] in 1 mL
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7201-01	1 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 4 of 4

CALCIUM CHLORIDE SOLUTION

calcium chloride liquid

Product Information

Item Code (Source)	NDC:0338-7401
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Inactive Ingredients

Ingredient Name	Strength
CALCIUM CHLORIDE (UNII: M4I0D6VV5M)	40 umol in 1 mL
WATER (UNII: 059QF0KO0R)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7401-01	1 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin kit

Product Information

Product Type	PLASMA DERIVATIVE	Item Code (Source)	NDC:0338-4302
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Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-4302-04	1 in 1 CARTON; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Quantity of Parts

Part #	Package Quantity	Total Product Quantity
Part 1	1 VIAL, GLASS	2 mL
Part 2	1 VIAL, GLASS	2 mL
Part 3	1 VIAL, GLASS	2 mL

Part 1 of 4**SEALER PROTEIN CONCENTRATE HUMAN**

fibrinogen human powder, for solution

Product Information

Item Code (Source)	NDC:0338-7112
Route of Administration	TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - UNII:N94833051K)	FIBRINOGEN HUMAN	86.5 mg in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	
NIACINAMIDE (UNII: 25X5118RD4)	
POLYSORBATE 80 (UNII: 6OZP39ZG8H)	
TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7112-02	2 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 2 of 4**HUMAN THROMBIN**

human thrombin powder, for solution

Product Information

Item Code (Source)	NDC:0338-7332
Route of Administration	TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [iU] in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
SODIUM CHLORIDE (UNII: 451W47IQ8X)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7332-02	2 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 3 of 4

FIBRINOLYSIS INHIBITOR SOLUTION

aprotinin liquid

Product Information

Item Code (Source)	NDC:0338-7201
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name	Strength

APROTININ (UNII: 04XPW8C0FL)	3000 [iU] in 1 mL
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7201-02	2 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 4 of 4

CALCIUM CHLORIDE SOLUTION

calcium chloride liquid

Product Information

Item Code (Source)	NDC:0338-7401
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name	Strength
CALCIUM CHLORIDE (UNII: M4I0D6VV5M)	40 umol in 1 mL
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7401-02	2 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin kit

Product Information

Product Type	PLASMA DERIVATIVE	Item Code (Source)	NDC:0338-4303
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Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-4303-10	1 in 1 CARTON; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Quantity of Parts

Part #	Package Quantity	Total Product Quantity
Part 1	1 VIAL, GLASS	5 mL
Part 2	1 VIAL, GLASS	5 mL
Part 3	1 VIAL, GLASS	5 mL
Part 4	1 VIAL, GLASS	5 mL

Part 1 of 4

SEALER PROTEIN CONCENTRATE HUMAN

fibrinogen human powder, for solution

Product Information

Item Code (Source)	NDC:0338-7112
Route of Administration	TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - UNII:N94833051K)	FIBRINOGEN HUMAN	86.5 mg in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	
NIACINAMIDE (UNII: 25X51I8RD4)	
POLYSORBATE 80 (UNII: 6OZP39ZG8H)	
TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7112-05	5 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 2 of 4

HUMAN THROMBIN

human thrombin powder, for solution

Product Information

Item Code (Source)	NDC:0338-7332
Route of Administration	TOPICAL

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [iU] in 1 mL

Inactive Ingredients

Ingredient Name	Strength
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
SODIUM CHLORIDE (UNII: 451W47IQ8X)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7332-05	5 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 3 of 4

FIBRINOLYSIS INHIBITOR SOLUTION

aprotinin liquid

Product Information

Item Code (Source)	NDC:0338-7201
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name	Strength
APROTININ (UNII: 04XPW8C0FL)	3000 [iU] in 1 mL
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7201-05	5 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Part 4 of 4

CALCIUM CHLORIDE SOLUTION

calcium chloride liquid

Product Information

Item Code (Source)	NDC:0338-7401
Route of Administration	TOPICAL

Inactive Ingredients

Ingredient Name	Strength
CALCIUM CHLORIDE (UNII: M4I0D6VV5M)	40 umol in 1 mL
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-7401-05	5 mL in 1 VIAL, GLASS; Type 0: Not a Combination Product		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin solution

Product Information

Product Type	PLASMA DERIVATIVE	Item Code (Source)	NDC:0338-8402
Route of Administration	TOPICAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - UNII:N94833051K)	FIBRINOGEN HUMAN	90 mg in 1 mL
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [iU] in 1 mL

Inactive Ingredients

Ingredient Name	Strength
APROTININ (UNII: 04XPW8C0FL)	
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	
NIACINAMIDE (UNII: 25X51I8RD4)	
POLYSORBATE 80 (UNII: 6OZP39ZG8H)	
TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)	
CALCIUM CHLORIDE (UNII: M4I0D6VV5M)	
SODIUM CHLORIDE (UNII: 451W47IQ8X)	
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-8402-02	1 in 1 CARTON		
1	NDC:0338-8402-01	2 mL in 1 SYRINGE, PLASTIC; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		
2	NDC:0338-8402-04	1 in 1 CARTON		
2	NDC:0338-8402-03	4 mL in 1 SYRINGE, PLASTIC; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		
3	NDC:0338-8402-10	1 in 1 CARTON		
3	NDC:0338-8402-09	10 mL in 1 SYRINGE, PLASTIC; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	12/31/2020

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin solution

Product Information

Product Type	PLASMA DERIVATIVE	Item Code (Source)	NDC:0338-9560
Route of Administration	TOPICAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - FIBRINOGEN HUMAN)		86.5 mg

UNII:N94833051K	FIBRINOGEN HUMAN	in 1 mL
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [iU] in 1 mL

Inactive Ingredients

Ingredient Name	Strength
APROTININ (UNII: 04XPW8C0FL)	
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	
NIACINAMIDE (UNII: 25X51I8RD4)	
POLYSORBATE 80 (UNII: 6OZP39ZG8H)	
TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)	
CALCIUM CHLORIDE (UNII: M4I0D6VV5M)	
SODIUM CHLORIDE (UNII: 451W47IQ8X)	
WATER (UNII: 059QF0KO0R)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-9560-01	2 mL in 1 SYRINGE, PLASTIC; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin solution

Product Information

Product Type	PLASMA DERIVATIVE	Item Code (Source)	NDC:0338-9564
Route of Administration	TOPICAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - UNII:N94833051K)	FIBRINOGEN HUMAN	86.5 mg in 1 mL
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [iU] in 1 mL

Inactive Ingredients

Ingredient Name	Strength
APROTININ (UNII: 04XPW8C0FL)	
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	
NIACINAMIDE (UNII: 25X5118RD4)	
POLYSORBATE 80 (UNII: 6OZP39ZG8H)	
TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)	
CALCIUM CHLORIDE (UNII: M4I0D6VV5M)	
SODIUM CHLORIDE (UNII: 451W47IQ8X)	
WATER (UNII: 059QF0KOOR)	

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-9564-01	4 mL in 1 SYRINGE, PLASTIC; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

TISSEEL FIBRIN SEALANT

fibrinogen human, human thrombin solution

Product Information

Product Type	PLASMA DERIVATIVE	Item Code (Source)	NDC:0338-9568
Route of Administration	TOPICAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
FIBRINOGEN HUMAN (UNII: N94833051K) (FIBRINOGEN HUMAN - UNII:N94833051K)	FIBRINOGEN HUMAN	86.5 mg in 1 mL
HUMAN THROMBIN (UNII: 6K15ABL77G) (HUMAN THROMBIN - UNII:6K15ABL77G)	HUMAN THROMBIN	500 [IU] in 1 mL

Inactive Ingredients

Ingredient Name	Strength
APROTININ (UNII: 04XPW8C0FL)	
ALBUMIN HUMAN (UNII: ZIF514RVZR)	
HISTIDINE (UNII: 4QD397987E)	

NIACINAMIDE (UNII: 25X51I8RD4)

POLYSORBATE 80 (UNII: 60ZP39ZG8H)

TRISODIUM CITRATE DIHYDRATE (UNII: B22547B95K)

CALCIUM CHLORIDE (UNII: M4I0D6VV5M)

SODIUM CHLORIDE (UNII: 451W47IQ8X)

WATER (UNII: 059QF0KO0R)

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0338-9568-01	10 mL in 1 SYRINGE, PLASTIC; Type 9: Other Type of Part 3 Combination Product (e.g., Drug/Device/Biological Product)		

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
BLA	BLA103980	05/01/1998	

Labeler - Baxter Healthcare Corporation (005083209)

Revised: 9/2025

Baxter Healthcare Corporation