

Andexanet Alfa (Andexxa)

Indicated for the treatment of life-threatening bleeding caused by rivaroxaban or apixaban.

This drug is restricted and **MUST** meet inclusion criteria, meet no exclusion criteria and have appropriate approval before use.

Criteria for use:

- Intracranial hemorrhage or other CNS bleed – evaluation of patient must be completed and use approved by neurosurgery
- Other bleeding site with hemodynamic instability – use must be approved by “hematology staff consult” attending
- Reversal for emergent surgery (non-CNS related) – use must be approved by “hematology staff consult” attending

Exclusion Criteria:

- Anticipated use of heparin or LMWH in the next 12 hours
- Already received PCC, factor VII or other factor-based reversal agent
- Expected survival < 1 months or non-survivable CNS bleed
- Reversal of enoxaparin
- Reversal for planned procedure

Pharmacist Expectations:

- Confirm what anticoagulant the patient was taking AND the time of last dose (if possible)
- Review the patient’s renal function and expected time of drug clearance
- Confirm indication for use and if it meets approved indication
 - This should only be used for life-threatening or debilitating bleeding
 - PCC, PRBC are appropriate alternatives for other types of bleeding
- If CNS bleed – is it approved by Neuro or Spine; If non-CNS bleed was hematology attending involved
 - If for emergent procedure – is there a time for this procedure?
- Confirm that a STAT heparin level by anti-Xa (UH only) was ordered AND a drug specific Xa level was ordered
 - If time allows the STAT heparin level should be resulted prior to administration of andexanet. If this level is undetectable then reversal is not needed.
 - Do NOT wait for the drug-specific level to result
- Cost of andexanet - \$25,000-50,000 per dose

If use is approved recommend:

Dosing Table

	Dose Taken	Timing of last dose	
		< 8 hours or unknown	≥ 8 hours
Apixaban	> 5 mg or unknown	800 mg bolus then 8 mg/min for 120 mins	400 mg bolus then 4 mg/min for 120 minutes
	≤ 5 mg	400 mg bolus then 4 mg/min for 120 mins	
Rivaroxaban	> 10 mg or unknown	800 mg bolus then 8 mg/min for 120 mins	
	≤ 10 mg	400 mg bolus then 4 mg/min for 120 mins	
Edoxaban	Any dose	800 mg bolus then 8 mg/min for 120 mins	

Reconstitution Guide

NOTE: For efficient and timely infusion, it is recommended that the IV bolus dose of ANDEXXA be prepared first and administered to the patient. The continuous IV infusion dose can then be prepared while the bolus dose is being administered to the patient so that it is ready for administration immediately (within 2 minutes) following the bolus dose.

RECONSTITUTION*:

- Remove flip-top from each vial and wipe each exposed top with alcohol swabs



- Withdraw Sterile Water for Injection (SWFI) from vial
- Using syringe with 20-gauge or higher needle, insert the syringe needle through the center of the rubber stopper

- Slowly inject Sterile Water for Injection, directing SWFI onto inside wall of the vial to minimize foaming
- For most efficient reconstitution of needed dose, inject each vial needed with SWFI before proceeding to the next step

Syringe size and SWFI volume for ANDEXXA reconstitution

Vial strength	Syringe size	SWFI volume
100 mg	10 mL	10 mL
200 mg	20 mL	20 mL

- Gently swirl each vial until complete dissolution of cake or powder occurs—avoid prolonged or vigorous agitation



DO NOT SHAKE
the vials as it can lead to foaming

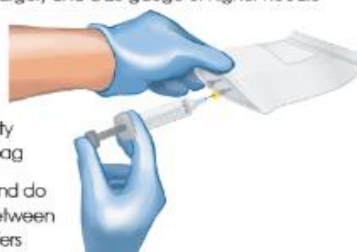
- Dissolution time for each vial is approximately 3-5 minutes

- Inspect reconstituted vials for particulate matter and discoloration prior to administration:

- Solution should be clear colorless to light yellow
- Solution may develop a few translucent particles, as ANDEXXA is a protein
- Do not use if opaque particles, discoloration, or other foreign particles are present

- Once all that are needed are reconstituted, withdraw reconstituted solution from each using an appropriate size syringe (60 mL or larger) and a 20-gauge or higher needle

- Transfer the reconstituted solution from syringe into the appropriate empty 250 mL or less IV bag



- Hold syringe up and do not set it down between multiple vial transfers

- Finally, discard all used syringes, needles, and vials, including any unused portion of reconstituted solution

STORAGE

- Reconstituted ANDEXXA vials can be stored at room temperature for up to 8 hours
- Reconstituted vials may be stored for up to 24 hours at 2-8 degrees C
- Reconstituted ANDEXXA is stable in IV bags for up to 8 hours at room temperature

*Note: ANDEXXA does not need to be brought to room temperature before reconstitution or administration to the patient.