Penicillin allergy skin testing: Essential supplies at a glance

Prep for success: Supplies needed for a single penicillin allergy skin test

Reagents/antigens: Single PRE-PEN[®] ampule [0.2 cc], penicillin G (PenG) dilution (10,000 units/mL) [0.2 cc], histamine (1.0 mg/mL) [0.1 cc], saline [0.2 cc]

Additional supplies: (5) skin testing medical devices,* (4) syringe labels,* (4) alcohol swabs,* (1) reaction guide,* (1) recording form,* (4) syringes (detachable needles), (7) detachable needles total

Optional oral challenge: Although optional, the CDC recommends performing an oral challenge using a 250-mg amoxicillin capsule or another appropriate alternative.

*Conveniently available as part of the ALK Penicillin Skin Testing Supply Kit (see ordering information below).

Penicillin allergy skin testing ordering information

Item	Description	NDC/UPC	ALK	AmerisourceBergen	Cardinal	McKesson	Diagnostic
PRE-PEN® (benzylpenicilloyl polylysine injection USP) Skin Test Antigen (5 ampules)	5 ampules x 0.25 mL each (1 ampule used per test)	49471-001-05	1058481	10002772	4300927	2166528	Skin test antigen
Histatrol® (histamine phosphate) 1.0 mg/mL 5 mL DV	5-mL dropper vial (multi-use)	0268-0247-05	HIST14	10099823	1612761	1985357	Positive control
Penicillin G 10,000 U/mL (obtain from pharmacy or order available brand)	Usually available in 5-MU or 20-MU vials	Recommended strength for skin testing per NIH (or equivalent) is 10,000 units/mL					Skin test antigen
Normal saline (obtain from pharmacy or order available brand)	Saline	N/A					Negative control
Penicillin Skin Testing Supply Kit (5 kits per order)	Contains testing supplies only, no antigens	N/A	1059360	10153136	5324173	3559937	Testing supplies
4 or 7, 25-28G syringes (TB)—for prepping testing solutions/intradermals	Follow hospital protocols to determine the number of syringes needed. The 3 syringes used for intradermals requir						re needles
DuoTip-Test® II Skin Test Device 20 Packs of 20 Duotip	Quantity: 400 tips total (5 tips used per test)	892093002050	1042815	10099767	4511549	3495285	Skin test medical device
Alternative ordering option for DuoTip-Test® II	Description	NDC/UPC	ALK	AmerisourceBergen	Cardinal	McKesson	Diagnostic
DuoTip-Test® II Skin Test Device Box of 50 packs. 5 Duotips/pack	1 pack needed per test	N/A	1058155	Available via drop ship from ALK			Skin test medical device

INDICATION

PRE-PEN is indicated for the assessment of sensitization to penicillin (benzylpenicillin or penicillin G) in patients suspected to have clinical penicillin hypersensitivity. A negative skin test to PRE-PEN is associated with an incidence of immediate allergic reactions of less than 5% after the administration of therapeutic penicillin, whereas the incidence may be more than 50% in a history-positive patient with a positive skin test to PRE-PEN. These allergic reactions are predominantly dermatologic. Whether a negative skin test to PRE-PEN predicts a lower risk of anaphylaxis is not established. Similarly, when deciding the risk of proposed penicillin treatment, there are not enough data at present to permit relative weighing in individual cases of a history of clinical penicillin hypersensitivity as compared to positive skin tests to PRE-PEN and/or minor penicillin determinants.

PRE-PEN[®] (benzylpenicilloyl polylysine injection USP Skin Test Antigen /isit penallergytest.com or contact us at <u>prepen@alk.net</u> to learn how ALK can help you impact antibiotic treatment hrough penicillin allergy skin testing.

PRE-PEN®

benzylpenicilloyl polylysine injection, solution Skin Test Antigen

DESCRIPTION:

PRE-PEN has the following structure:



R = Benzylpenicilloyl or H

CLINICAL PHARMACOLOGY:

PRE-PEN is a skin test antigen reagent that reacts specifically with benzylpenicilloy IgE antibodies initiating the release of chemical mediators which produce an immediate wheal and flare reaction at a skin test site. All individuals exhibiting a positive skin test to PRE-PEN possess IgE against the benzylpenicilloyI structural group which is a hapten. A hapten is a low molecular weight chemical that conjugates with a carrier (e.g. poly-I-lysine) resulting in the formation of an antigen with the hapten's specificity. The benzylpenicilloyI hapten is the major antigenic determinant in penicillin-allergic individuals. However, many individuals reacting positively to PRE-PEN will not develop a systemic allergic reaction on subsequent exposure to therapeutic penicillin, especially among those who have not reacted to penicillins in the past. Thus, the PRE-PEN skin test determines the presence of penicilloyI IgE antibodies which are necessary but not sufficient for acute allergic reactions due to the major penicilloyI determinant. Non-benzylpenicilloyI haptens are designated as minor determinants, since they less frequently elicit an immune response in penicillin treated individuals. The minor determinants may nevertheless be associated with significant clinical hypersensitivity. PRE-PEN does not react with IgE antibodies directed against non-benzylpenicilloyI haptens.

INDICATIONS AND USAGE:

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CONTRAINDICATIONS:

PRE-PEN is contraindicated in those patients who have exhibited either a systemic or marked local reaction to its previous administration. Patients known to be extremely hypersensitive to penicillin should not be skin tested.

WARNINGS:

The risk of sensitization to repeated skin testing with PRE-PEN is not established. Rarely, a systemic allergic reaction including anaphylaxis (see below) may follow a skin test with PRE-PEN. To decrease the risk of a systemic allergic reaction, puncture skin testing should be performed first. Intradermal skin testing should be performed only if the puncture test is entirely negative.

PRECAUTIONS:

General:

No reagent, test, or combination of tests will completely assure that a reaction to penicillin therapy will not occur.

The value of the PRE-PEN skin test alone as a means of assessing the risk of administering therapeutic penicillin (when penicillin is the preferred drug of choice) in the following situations is not established:

- 1. Adult patients who give no history of clinical penicillin hypersensitivity.
- 2. Pediatric patients.

In addition, the clinical value of PRE-PEN where exposure to penicillin is suspected as a cause of a current drug reaction or in patients who are undergoing routine allergy evaluation is not known. Likewise, the clinical value of PRE-PEN skin tests alone in determining the risk of administering semi-synthetic penicillins (phenoxymethyl penicillin, ampicillin, carbenicillin, dicloxacillin, methicillin, natcillin, oxacillin, amoxicillin), cephalosporin-derived antibiotics, and penem antibiotics is not known.

In addition to the results of the PRE-PEN skin test, the decision to administer or not administer penicillin should take into account individual patient factors. Healthcare professionals should keep in mind the following:

- 1. A serious allergic reaction to therapeutic penicillin may occur in a patient with a negative skin test to PRE-PEN.
- It is possible for a patient to have an anaphylactic reaction to therapeutic penicillin in the presence of a negative PRE-PEN skin test and a negative history of clinical penicillin hypersensitivity.

3. If penicillin is the drug of choice for a life-threatening infection, successful desensitization with therapeutic penicillin may be possible irrespective of a positive skin test and/or a positive history of clinical penicillin hypersensitivity.

Pregnancy – Pregnancy Category C:

Animal reproduction studies have not been conducted with PRE-PEN. It is not known whether PRE-PEN can cause fetal harm when administered to a pregnant woman or can affect reproduction capacity. The hazards of skin testing in such patients should be weighed against the hazard of penicillin therapy without skin testing.

ADVERSE REACTIONS:

Occasionally, patients may develop an intense local inflammatory response at the skin test site. Rarely, patients will develop a systemic allergic reaction, manifested by generalized erythema, pruritus, angioedema, urticaria, dyspnea, hypotension, and anaphylaxis. The usual methods of treating a skin test antigen-induced reaction — the applications of a venous occlusion tourniquet proximal to the skin test site and administration of epinephrine are recommended. The patient should be kept under observation for several hours.

DOSAGE AND ADMINISTRATION:

SKIN TESTING DOSAGE AND TECHNIQUE Skin testing responses can be attenuated by interfering drugs (e.g. H1-antihistamines and vasopressors). Skin testing should be delayed until the effects of such drugs have dissipated, or a separate skin test with histamine can be used to evaluate persistent antihistaminic effects in vivo. Due to the risk of potential systemic allergic reactions, skin testing should be performed in an appropriate healthcare setting under direct medical supervision.

Puncture Testing:

Skin testing is usually performed on the inner volar aspect of the forearm. The skin test antigen should always be applied first by the puncture technique. After preparing the skin surface, apply a small drop of PRE-PEN solution using a sterile 22-28 gauge needle. The same needle can then be used to make a single shallow puncture of the epidermis through the drop of PRE-PEN. Very little pressure is required to break the epidermal continuity. Observe for the appearance of a wheal, erythema, and the occurrence of itching at the test site during the succeeding 15 minutes at which time the solution over the puncture site is wiped off. A positive reaction consists of the development within 10 minutes of a pale wheal, sometimes with pseudopods, surrounding the puncture site and varying in diameter from 5 to 15 mm (or more). This wheal may be surrounded by a variable diameter of erythema, and accompanied by a variable degree of itching. The most sensitive individuals develop itching quickly, and the wheal and erythema are prompt in their appearance. As soon as a positive response as defined above is clearly evident, the solution over the scratch should be immediately wiped off. If the puncture test is either negative or equivocally positive (less than 5 mm wheal with little or no erythema and no itching), an intradermal test may

The Intradermal Test:

Using a 0.5 to 1.0 cc syringe with a 3/8" to 5/8"long, 26 to 30 gauge, short bevel needle, withdraw the contents of the ampule. Prepare with an alcohol swab a skin test area on the upper, outer arm, sufficiently below the deltoid muscle to permit proximal application of a tourniquet later, if necessary. Be sure to eject all air from the syringe through the needle, then insert the needle, bevel up immediately below the skin surface. Inject an amount of PRE-PEN sufficient to raise a small intradermal bleb of about 3 mm in diameter, in duplicate at least 2 cm apart. Using a separate syringe and needle, inject a like amount of saline or allergen diluting solution as a control at least 5 cm removed from the antigen test sites. Most skin reactions will develop within 5-15 minutes and response to the skin test is read at 20 minutes as follows:

 $\ensuremath{\textit{Negative response}}\xspace - \ensuremath{\mathsf{no}}\xspace$ in size of original bleb and no greater reaction than the control site.

Ambiguous response — wheal only slightly larger than initial injection bleb, with or without accompanying erythematous flare and slightly larger than the control site; OR discordance between duplicates.

Positive response — itching and significant increase in size of original blebs to at least 5 mm. Wheal may exceed 20 mm in diameter and exhibit pseudopods.

If the control site exhibits a wheal greater than 2-3 mm, repeat the test, and if the same reaction is observed, a physician experienced with allergy skin testing should be consulted.

HOW SUPPLIED: NDC 49471-001-05

PRE-PEN® (benzylpenicilloyl polylysine injection USP) is a clear, colorless, sterile solution supplied in ampules containing 0.25 mL.

Box of 5 single dose ampules. Ampules are opened by snapping the neck of the ampule using two forefingers of each hand. Visually inspect for glass chards before use. Each ampule is for single patient use only. Discard any unused portion.

PRE-PEN is optimally stored under refrigeration (2-8° C). PRE-PEN subjected to ambient temperatures for more than 24 hours should be discarded. As with all parenteral drug products, PRE-PEN should be inspected visually for particulate matter and discoloration prior to administration.

${\bf R}$ only

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Printed in USA

PRPE399999 07/13 18232 ©2013 ALK-Abelló, Inc. and AllerQuest LLC

