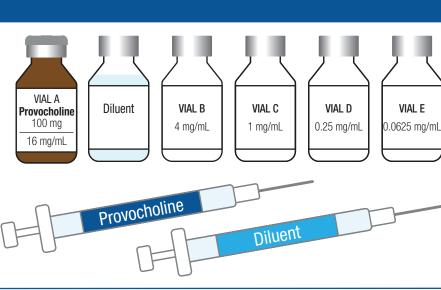
Steps to Diluting Methacholine Chloride **ATS "Short" - Quadrupling Doses**

Dilution Sequence Protocol for Provocholine 100 mg/vial (Canada DIN 02239656)

Getting Started

- 1. Attach labels to sterile empty vials.
- Wipe down the stoppers of the Provocholine vial, diluent vial, and sterile empty vials with alcohol prep pads.
- 3. Label two (2) appropriately sized syringes (one for Provocholine, one for diluent), and attach needles to each.



Preparing Vial A

16 mg/mL Solution

- 1. Using the diluent syringe, draw 6.25 mL of diluent and transfer it to the Provocholine 100 mg vial.
- 2. Shake well.

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Preparing Vial B

- 4 mg/mL Solution
- 1. Using the Provocholine syringe, draw 3 mL of the solution from Vial A and transfer it to Vial B.
- 2. Using the diluent syringe, draw 9 mL of diluent and transfer to Vial B.
- 3. Shake well.
- 3

Preparing Vial C

- 1 mg/mL Solution
- 1. Using the Provocholine needle and syringe, draw 3 mL of solution from Vial B and transfer to Vial C.
- 2. Using the diluent needle and syringe, draw 9 mL of diluent and transfer to Vial C.
- 3. Shake well.

Provocholine[®] (methacholine chloride)

www.provocholine.ca

NOTE

1. Transfer all dilutions (in vials A through E) to nebulizer through the 0.22 μm sterile bacterial retentive filter (Millex GV®).*

Vial C

1 ma/mL

Vial B

4 mg/mL

2. Be sure to follow your institutional policy for needle and sharps safety.

3 ml

VIAL A

Provocholin 100 mg

16 mg/mL

Vial B

4 ma/ml

*Do not attach filter until AFTER solution has been drawn into syringe.

6.25 mL

9 mL

9 mL

Diluent

Diluent

Diluent

VIAL A

100 mg

Vial B

4 ma/mL

Vial C

1 mg/ml

Preparing Vial D

- 0.25 mg/mL Solution
- 1. Using the Provocholine needle and syringe, draw 3 mL of solution from Vial C and transfer to Vial D.
- 2. Using the diluent needle and syringe, draw 9 mL of diluent and transfer to Vial D.
- 3. Shake well.

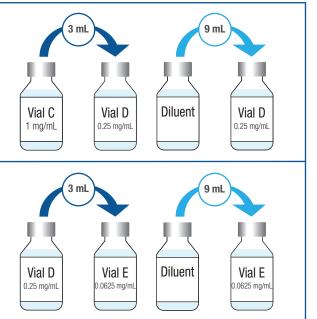
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Preparing Vial E

- 0.0625 mg/mL Solution
- 1. Using the Provocholine needle and syringe, draw 3 mL of solution from Vial D and transfer to Vial E.
- 2. Using the diluent needle and syringe, draw 9 mL of diluent and transfer to Vial E.
- 3. Shake well.

Dilution Check Sheet and Control Record



PROVOCHOLINE DILUTIONS FOR CHALLENGE TESTING

Date	Prepared by	Checked by
Provocholine (see label on vial)	Lot Number	Expiration Date
Diluent	Lot Number	Expiration Date

Take Provocholine	Add diluent (Shake well!)	Obtain dilution	Vial name	Initial / Date
Provocholine 100 mg	6.25 mL	16 mg/mL	Vial A - 16 mg/mL	
3 mL from Vial A	9 mL	4 mg/mL	Vial B - 4 mg/mL	
3 mL from Vial B	9 mL	1 mg/mL	Vial C - 1 mg/mL	
3 mL from Vial C	9 mL	0.25 mg/mL	Vial D - 0.25 mg/mL	
3 mL from Vial D	9 mL	0.0625 mg/mL	Vial E - 0.0625 mg/mL	

Provocholine (methacholine chloride) www.provocholine.ca

NOTE

1. Transfer all dilutions (in vials A through E) to nebulizer through the 0.22 μm sterile bacterial retentive filter (Millex GV®).*

2. Be sure to follow your institutional policy for needle and sharps safety.

*Do not attach filter until AFTER solution has been drawn into syringe.

- Provocholine powder should be stored at 59° to 86°F (15° to 30°C).
- Dilutions A through E (16 mg/mL through 0.0625 mg/mL) should be stored at 36° to 46°F (2° to 8°C) in a refrigerator for no more than 2 weeks.
- Be sure to remove the Provocholine dilutions from the refrigerator 30 minutes prior to testing to allow them to warm to room temperature (59° to 77°F or 15° to 25°C as per World Health Organization).

WARNING: SEVERE BRONCHOCONSTRICTION

Severe bronchoconstriction can result from Provocholine administration (including the lowest dose). The use of Provocholine is contraindicated in pediatric and adult patients with baseline $FEV_1 < 70\%$ predicted or adults with $FEV_1 < 1.5$ L. Because of thepotential for severe bronchoconstriction, the use of Provocholine in patients with clinically apparent asthma or wheezing is not recommended *[see Warnings and Precautions (7) of the Product Monograph].*

Emergency equipment and medication should be immediately available to treat acute respiratory distress. If severe bronchoconstriction occurs, reverse immediately with arapid-acting inhaled bronchodilator agent (β -agonist) [see Warnings and Precautions (7) of the Product Monograph].

If baseline spirometry is not performed or measured inaccurately, the initial FEV_1 may be underestimated. In this situation, decreases in FEV_1 may not be detected after administration of escalating Provocholine doses, which may result in administration of unnecessary higher doses and an increased risk for excessive bronchoconstriction [see Warnings and Precautions (7) of the Product Monograph].

Please see Product Monograph for full prescribing information, which is available on request by calling Methapharm Medical Information at 1-866-701-4636 or by email: medinfo@methapharm.com. You can report any suspected side effects associated with the use of health products to Health Canada by calling toll-free 1-866-234-2345 or by visiting the web page on Adverse Reaction Reporting (<u>http://www.hc-sc.gc.ca/dhp-mps/medeff/report-declaration/index-eng.php</u>) for information on how to report online, by mail or by fax. This information is provided as a professional courtesy, and it is intended to provide data available to us that may assist you in deriving your own conclusions and opinions. This information is not intended to advocate any indications, dosage, or other described in the Product Monograph. For all other inquiries, please call 1-800-287-7686 for Customer Service. This protocol is commonly known as the "ATS Short," and is available for download at <u>https://www.thoracic.org/statements/resources/pfet/methacholine1-21.pdf</u>.

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For technical support, please contact us Toll-Free at 1-800-287-7686

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