

DESCRIPTION: L-Methyl-MC is an orally administered prescription dietary supplement specifically formulated for the dietary management of patients with unique nutritional needs requiring increased folate levels. L-Methyl-MC should be administered under the supervision of a licensed medical practitioner.

Each round, coated, blue tablet contains the following dietary ingredients:

Supplement Facts	Serving Size 1 Tablet	Servings Per Container 90
	Amount Per Serving	% Daily Value
L-Methylfolate Calcium*, or 4(S)-5-MTHF-Ca	6 mg	**
Riboflavin	5 mg	**
Pyridoxine (as Pyridoxine HCl)	50 mg	**
Cyanocobalamin	1 mg	**

*contains less than 1% d-methylfolate

Other ingredients: See insert for more information.

**Daily Values not established for patients with unique nutritional needs who are in need of supplementation as directed by a licensed medical practitioner.

DOSAGE AND ADMINISTRATION: Usual adult dose is one to two tablets daily or as prescribed by a licensed medical practitioner.

L-Methyl-MC is not recommended for use in children under the age of twelve.



NDC† 76439-207-90

L-METHYL-MC

TABLETS Prescription Dietary Supplement

Rx
90 Tablets

Made in the USA

Store at controlled room temperature 15°-30°C (59°-86°F). [See USP]. Protect from light and moisture. Dispense in a light, light-resistant container.

If you are pregnant or nursing a baby, please ask a health professional.

KEEP THIS OUT OF REACH OF CHILDREN.

All prescriptions using this product shall be pursuant to state statutes as applicable. This is not an Orange Book product. Call your medical practitioner about side effects. You may report side effects by calling 813-283-1344.

NDC† 76439-207-90

† See insert for more information about National Drug Codes (NDCs).

Manufactured for: Virtus Pharmaceuticals, LLC
Tampa, FL 33619

Rev. 1/2012



Lot No.
Exp. Date:

L-METHYL-MC

Tablets

Rx

Prescription Dietary Supplement

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Other Ingredients: Dicalcium Phosphate, Microcrystalline Cellulose, FD&C Blue #1, Titanium Dioxide, Modified Cellulose, Croscarmellose Sodium, Stearic Acid, Magnesium Stearate, Silicon Dioxide, Polyethylene Glycol and Food Glaze.

FOLATE REGULATION: The term "folate" are B vitamins that include folic acid and any forms of active pteroylglutamates regardless of the reduction state of the molecule. Foliates, or vitamin B₉, are primarily hydrolyzed in the intestinal jejunum and the liver to the active circulating form of folate, L-methylfolate, with an intermediate stable form, 5,10-methylenetetrahydrofolate. Individuals with genetic polymorphisms for the genes coding methylenetetrahydrofolate reductase (MTHFR) may not be capable of utilizing or metabolizing folic acid adequately for the vitamin B₁₂ dependent methylation cycle.

Folic acid, including reduced forms* such as folic acid, may obscure pernicious anemia above 0.1 mg doses, and must be administered under the supervision of a licensed medical practitioner. The 1971, 1972, 1973, 1980, 1984, 2000, and 2010 Federal Register Notices addressed this concern while establishing that increased folate was proper therapy in megaloblastic anemias - specifically where homocysteine levels were elevated or risk of neural tube defects (NTDs) was at issue. The Federal Register Notice of August 2, 1973 (38 FR 20750) specifically states that: Dietary supplement preparations are available without a prescription (21 CFR 121.1134). Levels higher than dietary supplement amounts are available only with a prescription.

† It is not known whether or not L-methylfolate can obscure pernicious anemia above 0.1 mg doses, so caution is advised also with this form of folate.

Folic acid - including reduced forms, may be added to medical foods undefined in section 5(b)(3) of the Orphan Drug Act (21 USC 360ee(b)(3)), or to food (21 CFR 172.345).

INDICATIONS AND USAGE:

L-Methyl-MC is indicated for the distinct nutritional requirements of patients in need of dietary supplementation as determined by a licensed medical practitioner.

L-Methyl-MC should be administered under the supervision of a licensed medical practitioner.

CONTRAINDICATIONS: This product is contraindicated in patients with a known hypersensitivity to any of the ingredients.

WARNINGS: Caution is recommended in patients with a history of bipolar illness.

PRECAUTIONS: General: Folate, when administered as a single agent in doses above 0.1 mg daily, may obscure the detection of vitamin B₁₂ deficiency (specifically, the administration of folic acid may reverse the hematological manifestations of B₁₂ deficiency, including pernicious anemia, while not addressing the neurological manifestations). Folate therapy alone is inadequate for treatment of a vitamin B₁₂ deficiency.

PATIENT INFORMATION: L-Methyl-MC is a prescription dietary supplement to be used only under licensed medical supervision.

DRUG INTERACTIONS: Drugs which may interact with folate include:

- Antiepileptic drugs (AED): The AED class including, but not limited to, phenytoin, carbamazepine, primidone, valproic acid, fosphenytoin, valproate, phenobarbital and lamotrigine have been shown to impair folate absorption and increase the metabolism of circulating folate.
- Additionally, concurrent use of folic acid has been associated with enhanced phenytoin metabolism, lowering the level of the AED in the blood and allowing breakthrough seizures to occur. Caution should be used when prescribing this product among patients who are receiving treatment with phenytoin and other anticonvulsants.
- Capecitabine: Folic acid (5-formyltetrahydrofolate) may increase the toxicity of Capecitabine.
- Cholestyramine: Reduces folic acid absorption and reduces serum folate levels.
- Colestipol: Reduces folic acid absorption and reduces serum folate levels.
- Cycloserine: Reduces folic acid absorption and reduces serum folate levels.
- Dihydrofolate Reductase Inhibitors (DHFRi): DHFRi block the conversion of folic acid to its active forms, and lower plasma and red blood cell folate levels. DHFRi include aminopterin, methotrexate, pyrimethamine, triamterene, and trimethoprim.
- Fluoxetine: Fluoxetine exerts a noncompetitive inhibition of the 5-methyltetrahydrofolate active transport in the intestine.

- Isotretinoin: Reduced folate levels have occurred in some patients taking isotretinoin.
- L-dopa, triamterene, colchicine, and trimethoprim may decrease plasma folate levels.
- Nonsteroidal Anti-inflammatory Drugs (NSAIDs): NSAIDs have been shown to inhibit some folate dependent enzymes in laboratory experiments. NSAIDs include ibuprofen, naproxen, indomethacin and sulindac.
- Oral Contraceptives: Serum folate levels may be depressed by oral contraceptive therapy.
- Methylprednisolone: Reduced serum folate levels have been noted after treatment with methylprednisolone.
- Pancreatic Enzymes: Reduced serum folate levels have occurred in some patients taking pancreatic extracts, such as pancreatin and pancrelipase.
- Pentamidine: Reduced folate levels have been seen with prolonged intravenous pentamidine.
- Pyrimethamine: High levels of folic acid may result in decreased serum levels of pyrimethamine.
- Smoking and Alcohol: Reduced serum folate levels have been noted.
- Sulfasalazine: Inhibits the absorption and metabolism of folic acid.
- Metformin treatment in patients with type 2 diabetes decreases serum folate.
- Warfarin can produce significant impairment in folate status after a 6-month therapy.
- Folinic acid may enhance the toxicity of fluorouracil.
- Concurrent administration of chloramphenicol and folic acid in folate-deficient patients may result in antagonism of the haematopoietic response to folate.
- Caution should be exercised with the concomitant use of folic acid and trimethoprim-sulfamethoxazole for the acute treatment of *Pneumocystis carinii* pneumonia in patients with HIV infection as it is associated with increased rates of treatment failure and mortality in a placebo controlled study.

Drugs which interact with vitamin B₆:

- Vitamin B₆ should not be given to patients receiving the drug levodopa because the action of levodopa is antagonized by vitamin B₆. However, vitamin B₆ may be used concurrently in patients receiving a preparation containing both carbidopa and levodopa.

Drugs which may interact with vitamin B₁₂:

- Antibiotics, cholestyramine, colchicines, colestipol, metformin, para-aminosalicylic, and potassium chloride may decrease the absorption of vitamin B₁₂.
- Nitrous oxide can produce a functional vitamin B₁₂ deficiency.

PREGNANCY AND NURSING MOTHERS

L-Methyl-MC is not intended for use as a prenatal/postnatal multivitamin for lactating and non-lactating mothers. This product contains B vitamins. Talk with your medical practitioner before using L-Methyl-MC if pregnant or lactating.

ADVERSE REACTIONS: Allergic sensitization has been reported following both oral and parental administration of folic acid, and may possibly occur with other forms of folate.

Paresthesia, somnolence, nausea, and headaches have been reported with vitamin B₆.

Mild transient diarrhea, polycythemia vera, itching, transitory exanthema and the feeling of swelling of the entire body have been associated with vitamin B₁₂.

If headaches occur with the use of this product, consult your medical practitioner.

DOSAGE AND ADMINISTRATION: Usual adult dose is one to two tablets daily or as prescribed by a licensed medical practitioner. L-Methyl-MC is not recommended for use in children under the age of twelve.

HOW SUPPLIED: L-Methyl-MC is supplied as round, coated, blue tablets debossed "BP" on top and "850" on bottom, dispensed in bottles of 90 tablets.

NDC†: 76439-207-90

† This product is a dietary supplement that due to increased folate levels (AUG 2, 1973 38 FR 20750), requires an Rx on the label because of increased risk associated with masking of B₁₂ deficiency. As such, this product requires licensed medical supervision, an Rx status, and a National Drug Code (NDC) as required by pedigree reporting requirements.

STORAGE: Store at Controlled Room Temperature 15°-30°C (59°-86°F). [See USP]. Protect from light and moisture. Dispense in a light, light-resistant container.

Call your medical practitioner about side effects. You may report side effects by calling (813)-283-1344.

KEEP THIS OUT OF THE REACH OF CHILDREN.

Rx

Prescription Dietary Supplement

All prescriptions using this product shall be pursuant to state statutes as applicable. This is not an Orange Book product.

Manufactured for:
Virtus Pharmaceuticals, LLC
Tampa, FL 33619
www.virtusrx.com
MADE IN USA



REV 12/2012