

POTASSIUM CHLORIDE- potassium chloride solution

Apotex Corp

HIGHLIGHTS OF PRESCRIBING INFORMATION

These highlights do not include all the information needed to use POTASSIUM CHLORIDE ORAL SOLUTION safely and effectively. See full prescribing information for POTASSIUM CHLORIDE ORAL SOLUTION.

POTASSIUM CHLORIDE oral solution
Initial U.S. Approval: 1948

INDICATIONS AND USAGE

Potassium chloride oral solution is indicated for the treatment and prophylaxis of hypokalemia with or without metabolic alkalosis, in patients for whom dietary management with potassium-rich foods or diuretic dose reduction are insufficient. (1)

DOSAGE AND ADMINISTRATION

Dilute prior to administration. (2.1, 5.1)

Monitor serum potassium and adjust dosage accordingly. (2.2, 2.3)

Treatment of hypokalemia:

- Adults: Initial doses range from 40 to 100 mEq/day in 2 to 5 divided doses: limit doses to 40 mEq per dose. Total daily dose should not exceed 200 mEq (2.2)
- Pediatric patients aged birth to 16 years old: 2 to 4 mEq/kg/day in divided doses; not to exceed 1 mEq/kg as a single dose or 40 mEq whichever is lower; if deficits are severe or ongoing losses are great, consider intravenous therapy. Total daily dose should not exceed 100 mEq (2.3)

Maintenance or Prophylaxis of hypokalemia:

- Adults: Typical dose is 20 mEq per day (2.2)
- Pediatric patients aged birth to 16 years old: typical dose is 1 mEq/kg/day. Do not exceed 3 mEq/kg/day (2.3)

DOSAGE FORMS AND STRENGTHS

- Oral Solution: 10%; 1.3 mEq potassium per mL (3)
- Oral Solution: 20%; 2.6 mEq potassium per mL (3)

CONTRAINDICATIONS

- Concomitant use with potassium sparing diuretics. (4)

WARNINGS AND PRECAUTIONS

- Gastrointestinal Irritation: Dilute before use, take with meals (5.1)

ADVERSE REACTIONS

Most common adverse reactions are nausea, vomiting, flatulence, abdominal pain/discomfort, and diarrhea. (6)

To report SUSPECTED ADVERSE REACTIONS, contact Apotex Corp. at 1-800-706-5575 or FDA at 1-800-FDA-1088 or www.fda.gov/medwatch.

DRUG INTERACTIONS

- Potassium sparing diuretics: Avoid concomitant use (7.1)
- Renin-angiotensin-aldosterone inhibitors: Monitor for hyperkalemia (7.2)
- Nonsteroidal Anti-Inflammatory drugs: Monitor for hyperkalemia (7.3)

USE IN SPECIFIC POPULATIONS

Cirrhosis: Initiate therapy at the low end of the dosing range (8.5)

Renal Impairment: Initiate therapy at the low end of the dosing range (8.6)

Revised: 7/2022

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FULL PRESCRIBING INFORMATION

1 INDICATIONS AND USAGE

Potassium chloride oral solution is indicated for the treatment and prophylaxis of hypokalemia with or without metabolic alkalosis, in patients for whom dietary management with potassium-rich foods or diuretic dose reduction are insufficient.

2 DOSAGE AND ADMINISTRATION

2.1 Administration and Monitoring

Monitoring

Monitor serum potassium and adjust dosages accordingly. For treatment of hypokalemia, monitor potassium levels daily or more often depending on the severity of hypokalemia until they return to normal. Monitor potassium levels monthly to biannually for maintenance or prophylaxis.

The treatment of potassium depletion, particularly in the presence of cardiac disease, renal disease, or acidosis requires careful attention to acid-base balance, volume status, electrolytes, including magnesium, sodium, chloride, phosphate, and calcium, electrocardiograms and the clinical status of the patient. Correct volume status, acid-base balance and electrolyte deficits as appropriate.

Administration

Dilute the potassium chloride solution with at least 4 ounces of cold water [see *Warnings and Precautions* (5.1)].

Take with meals or immediately after eating.

If serum potassium concentration is < 2.5 mEq/L, use intravenous potassium instead of oral supplementation.

2.2 Adult Dosing

Treatment of hypokalemia

Daily dose range from 40 to 100 mEq. Give in 2 to 5 divided doses; limit doses to 40 mEq per dose. The total daily dose should not exceed 200 mEq in a 24 hour period.

Maintenance or Prophylaxis

Typical dose is 20 mEq per day. Individualize dose based upon serum potassium levels.

Studies support the use of potassium replacement in digitalis toxicity. When alkalosis is present, normokalemia and hyperkalemia may obscure a total potassium deficit. The advisability of use of potassium replacement in the setting of hyperkalemia is uncertain.

2.3 Pediatric Dosing

Treatment of hypokalemia

Pediatric patients aged birth to 16 years old: The initial dose is 2 to 4 mEq/kg/day in divided doses; do not exceed as a single dose 1 mEq/kg or 40 mEq, whichever is lower; maximum daily doses should not exceed 100 mEq. If deficits are severe or ongoing losses are great, consider intravenous therapy.

Maintenance or Prophylaxis

Pediatric patients aged birth to 16 years old: Typical dose is 1 mEq/kg/day. Do not exceed 3 mEq/kg/day.

3 DOSAGE FORMS AND STRENGTHS

Oral Solution 10%: 1.3 mEq potassium per mL.

Oral Solution 20%: 2.6 mEq potassium per mL.

4 CONTRAINDICATIONS

Potassium chloride is contraindicated in patients on potassium sparing diuretics.

5 WARNINGS AND PRECAUTIONS

5.1 Gastrointestinal Irritation

May cause gastrointestinal irritation if administered undiluted. Increased dilution of the solution and taking with meals may reduce gastrointestinal irritation [*see Dosage and Administration (2.1)*].

6 ADVERSE REACTIONS

The most common adverse reactions to oral potassium salts are nausea, vomiting, flatulence, abdominal pain/discomfort, and diarrhea.

7 DRUG INTERACTIONS

7.1 Potassium-Sparing Diuretics

Use with potassium-sparing diuretics can produce severe hyperkalemia. Avoid concomitant use.

7.2 Renin-Angiotensin-Aldosterone System Inhibitors

Drugs that inhibit the renin-angiotensin-aldosterone system (RAAS) including angiotensin converting enzyme (ACE) inhibitors, angiotensin receptor blockers (ARBs), spironolactone, eplerenone, or aliskiren produce potassium retention by inhibiting aldosterone production. Closely monitor potassium in patients receiving concomitant RAAS therapy.

7.3 Nonsteroidal Anti-Inflammatory Drugs (NSAIDs)

NSAIDs may produce potassium retention by reducing renal synthesis of prostaglandin E and impairing the renin-angiotensin system. Closely monitor potassium in patients on concomitant NSAIDs.

8 USE IN SPECIFIC POPULATIONS

8.1 Pregnancy

There are no human data related to use of Potassium Chloride during pregnancy, and animal studies have not been conducted. Potassium supplementation that does not lead to hyperkalemia is not expected to cause fetal harm.

The background risk for major birth defects and miscarriage in the indicated population is unknown. All pregnancies have a background risk of birth defect, loss, or other adverse outcomes. In the U.S. general population, the estimated background risk of

major birth defects and miscarriage in clinically recognized pregnancies is 2-4% and 15-20%, respectively.

8.2 Lactation

Risk Summary

The normal potassium ion content of human milk is about 13 mEq per liter. Since potassium from oral supplements such as Potassium Chloride becomes part of the body potassium pool, as long as body potassium is not excessive, the contribution of potassium chloride supplementation should have little or no effect on the level in human milk.

8.4 Pediatric Use

The safety and effectiveness of potassium chloride have been demonstrated in children with diarrhea and malnutrition from birth to 16 years.

8.5 Geriatric Use

Clinical studies of potassium chloride did not include sufficient numbers of subjects aged 65 and over to determine whether they respond differently from younger subjects. Other reported clinical experience has not identified differences in responses between the elderly and younger patients. In general, dose selection for an elderly patient should be cautious, usually starting at the low end of the dosing range, reflecting the greater frequency of decreased hepatic, renal, or cardiac function, and of concomitant disease or other drug therapy.

This drug is known to be substantially excreted by the kidney, and the risk of toxic reactions to this drug may be greater in patients with impaired renal function. Because elderly patients are more likely to have decreased renal function, care should be taken in dose selection, and it may be useful to monitor renal function.

8.6 Cirrhotics

Patients with cirrhosis should usually be started at the low end of the dosing range, and the serum potassium level should be monitored frequently. [see *Clinical Pharmacology* (12.3)].

8.7 Renal Impairment

Patients with renal impairment have reduced urinary excretion of potassium and are at substantially increased risk of hyperkalemia. Patients with impaired renal function, particularly if the patient is on ACE inhibitors, ARBs, or nonsteroidal anti-inflammatory drugs should usually be started at the low end of the dosing range because of the potential for development of hyperkalemia. The serum potassium level should be monitored frequently. Renal function should be assessed periodically.

10 OVERDOSAGE

10.1 Symptoms

The administration of oral potassium salts to persons with normal excretory

mechanisms for potassium rarely causes serious hyperkalemia. However, if excretory mechanisms are impaired or if potassium is administered too rapidly potentially fatal hyperkalemia can result.

Hyperkalemia is usually asymptomatic and may be manifested only by an increased serum potassium concentration (6.5 to 8.0 mEq/L) and characteristic electrocardiographic changes (peaking of T-waves, loss of P-waves, depression of S-T segment, and prolongation of the QT-interval). Late manifestations include muscle paralysis and cardiovascular collapse from cardiac arrest (9 to 12 mEq/L).

10.2 Treatment

Treatment measures for hyperkalemia include the following:

1. Monitor closely for arrhythmias and electrolyte changes.
2. Eliminate foods and medications containing potassium and of any agents with potassium-sparing properties such as potassium-sparing diuretics, ARBS, ACE inhibitors, NSAIDS, certain nutritional supplements and many others.
3. Administer intravenous calcium gluconate if the patient is at no risk or low risk of developing digitalis toxicity.
4. Administer intravenously 300 to 500 mL/hr of 10% dextrose solution containing 10 to 20 units of crystalline insulin per 1000 mL.
5. Correct acidosis, if present, with intravenous sodium bicarbonate.
6. Use exchange resins, hemodialysis, or peritoneal dialysis.

In patients who have been stabilized on digitalis, too rapid a lowering of the serum potassium concentration can produce digitalis toxicity.

11 DESCRIPTION

Potassium chloride, USP is a colorless, elongated, prismatic, or cubical crystals, or white, granular powder. It is soluble in water and slightly soluble in alcohol. Chemically, potassium chloride is K-Cl with a molecular mass of 74.55 g/mol.

Oral Solution 10%: Each 15 mL of solution contains 1.5 g of potassium chloride, USP and the following inactive ingredients: citric acid anhydrous, FD&C Yellow #6, glycerin, methylparaben, natural orange flavor, propylene glycol, propylparaben, purified water, sodium citrate dihydrate, sucralose.

Oral Solution 20%: Each 15 mL of solution contains 3.0 g of potassium chloride, USP and the following inactive ingredients: citric acid anhydrous, FD&C Yellow #6, glycerin, methylparaben, natural orange flavor, propylene glycol, propylparaben, purified water, sodium citrate dihydrate, sucralose.

Natural orange flavor includes fruit extract and natural flavor.

12 CLINICAL PHARMACOLOGY

12.1 Mechanism of Action

The potassium ion (K^+) is the principal intracellular cation of most body tissues. Potassium ions participate in a number of essential physiological processes including the maintenance of intracellular tonicity; the transmission of nerve impulses; the contraction of cardiac, skeletal, and smooth muscle; and the maintenance of normal renal function.

The intracellular concentration of potassium is approximately 150 to 160 mEq per liter. The normal adult plasma concentration is 3.5 to 5 mEq per liter. An active ion transport system maintains this gradient across the plasma membrane.

Potassium is a normal dietary constituent, and under steady-state conditions the amount of potassium absorbed from the gastrointestinal tract is equal to the amount excreted in the urine. The usual dietary intake of potassium is 50 to 100 mEq per day.

12.3 Pharmacokinetics

Based on published literature, the rate of absorption and urinary excretion of potassium from KCl oral solution were higher during the first few hours after dosing relative to modified release KCl products. The bioavailability of potassium, as measured by the cumulative urinary excretion of K^+ over a 24 hour post dose period, is similar for KCl solution and modified release products.

16 HOW SUPPLIED/STORAGE AND HANDLING

Potassium chloride oral solution, USP is an orange solution available in two strengths as follows:

10%: 20 mEq/15 mL oral solution
NDC# 60505-6184-6 Bottle of 473 mL

20%: 40 mEq/15 mL oral solution
NDC# 60505-6185-6 Bottle of 473 mL

Storage

Store at 20°C to 25°C (68°F to 77°F) [see USP Controlled Room Temperature]. Dispense in a tight, light-resistant container as defined in the USP. PROTECT from LIGHT and FREEZING.

APOTEX INC.

POTASSIUM CHLORIDE ORAL SOLUTION, USP

10% and 20%

Manufactured by: Manufactured for:
Apotex Inc. Apotex Corp.
Toronto, Ontario Weston, Florida
Canada M9L 1T9 USA 33326

Revised: July 2022

Rev. 5

PRINCIPAL DISPLAY PANEL - 20 mEq per 15 mL Bottle Label

Representative sample of labeling (see HOW SUPPLIED section of complete listing):

APOTEX CORP.

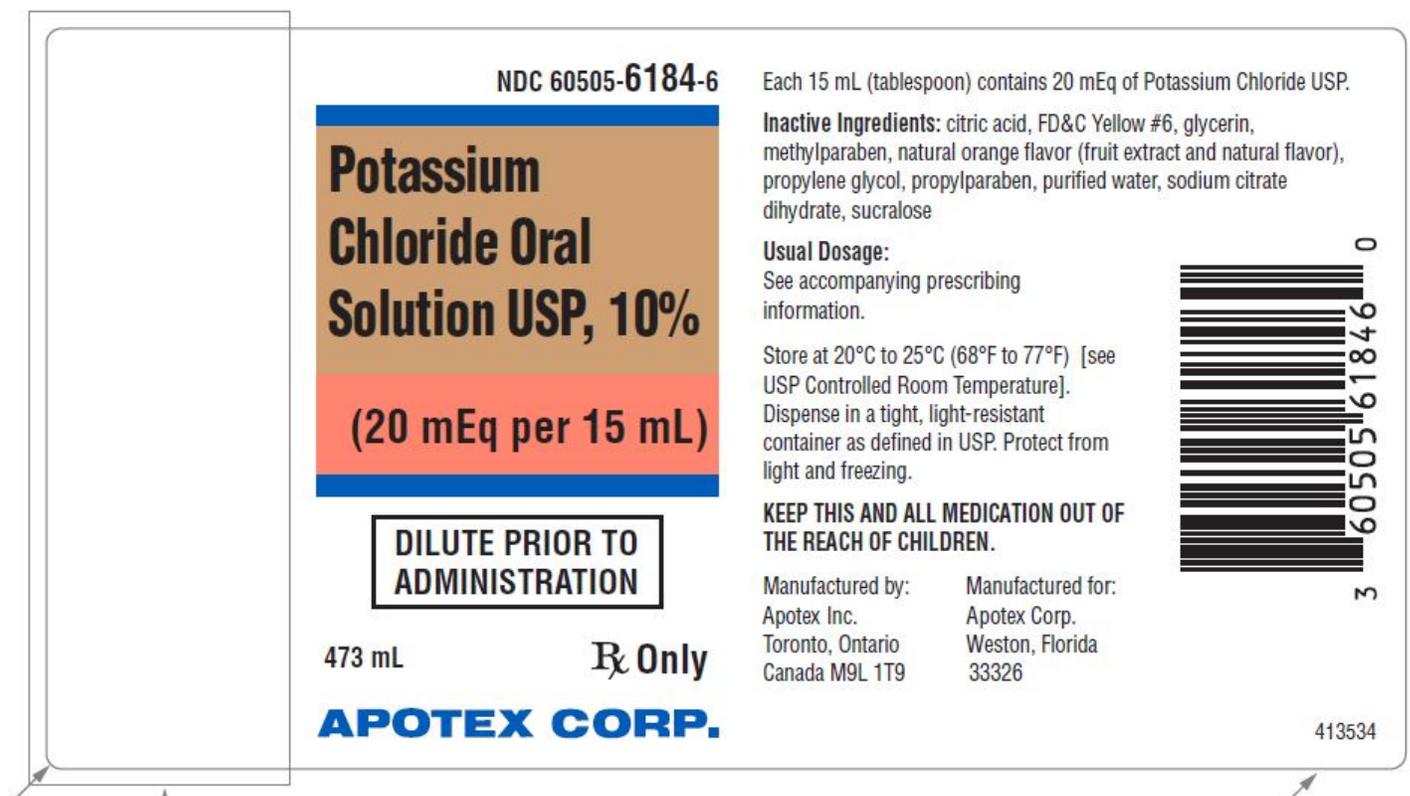
NDC 60605-6184-6

Potassium Chloride Oral Solution USP, 10%

20 mEq per 15 mL

Rx

473 mL



PRINCIPAL DISPLAY PANEL - 20 mEq per 15 mL Carton Label

Representative sample of labeling (see HOW SUPPLIED section of complete listing):

APOTEX CORP.

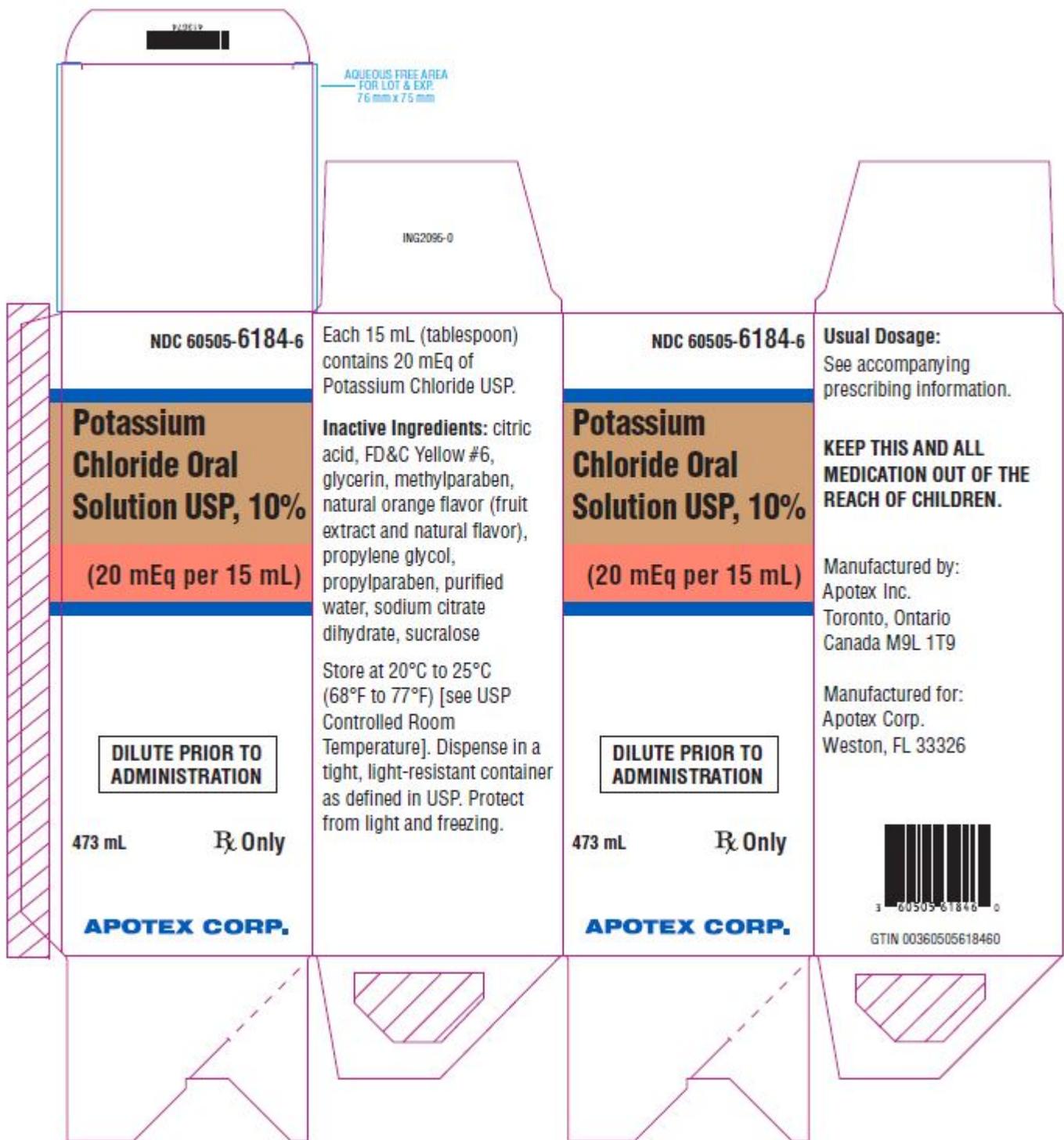
NDC 60605-6184-6

Potassium Chloride Oral Solution USP, 10%

20 mEq per 15 mL

Rx

473 mL



PRINCIPAL DISPLAY PANEL - 40 mEq per 15 mL Bottle Label

Representative sample of labeling (see HOW SUPPLIED section of complete listing):

APOTEX CORP.

NDC 60605-6185-6

Potassium Chloride Oral Solution USP, 20%

40 mEq per 15 mL

Rx

473 mL

NDC 60505-6185-6

Potassium Chloride Oral Solution USP, 20%

(40 mEq per 15 mL)

DILUTE PRIOR TO ADMINISTRATION

473 mL **Rx Only**

APOTEX CORP.

Each 15 mL (tablespoon) contains 40 mEq of Potassium Chloride USP.

Inactive Ingredients: citric acid, FD&C Yellow #6, glycerin, methylparaben, natural orange flavor (fruit extract and natural flavor), propylene glycol, propylparaben, purified water, sodium citrate dihydrate, sucralose

Usual Dosage:
See accompanying prescribing information.

Store at 20°C to 25°C (68°F to 77°F) [see USP Controlled Room Temperature].
Dispense in a tight, light-resistant container as defined in USP. Protect from light and freezing.

KEEP THIS AND ALL MEDICATION OUT OF THE REACH OF CHILDREN.

Manufactured by: Apotex Inc.
Toronto, Ontario
Canada M9L 1T9

Manufactured for: Apotex Corp.
Weston, Florida
33326

3 60505 61856 9

413573

PRINCIPAL DISPLAY PANEL - 40 mEq per 15 mL Carton Label

Representative sample of labeling (see HOW SUPPLIED section of complete listing):

APOTEX CORP.

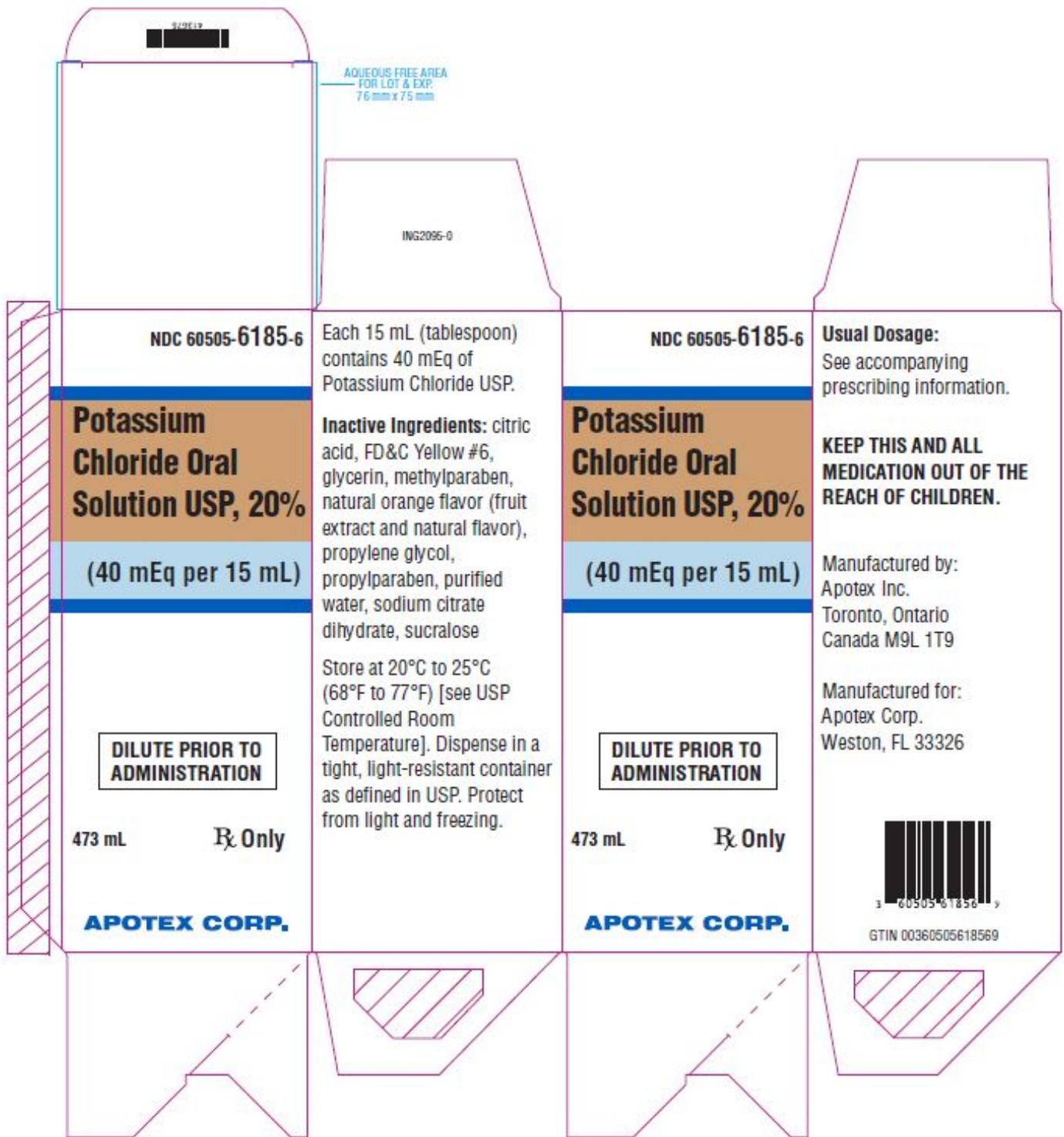
NDC 60605-6185-6

Potassium Chloride Oral Solution USP, 20%

40 mEq per 15 mL

Rx

473 mL



POTASSIUM CHLORIDE

potassium chloride solution

Product Information

| | | | |
|--------------------------------|-------------------------|---------------------------|----------------|
| Product Type | HUMAN PRESCRIPTION DRUG | Item Code (Source) | NDC:60505-6184 |
| Route of Administration | ORAL | | |

Active Ingredient/Active Moiety

| Ingredient Name | Basis of Strength | Strength |
|---|--------------------|-----------------|
| Potassium Chloride (UNII: 660YQ98I10) (Potassium Cation - UNII:295O53K152, Chloride Ion - UNII:Q32ZN48698) | Potassium Chloride | 20 meq in 15 mL |

Inactive Ingredients

| Ingredient Name | Strength |
|---|----------|
| Anhydrous Citric Acid (UNII: XF417D3PSL) | |
| FD&C Yellow NO. 6 (UNII: H77VEI93A8) | |
| Glycerin (UNII: PDC6A3C00X) | |
| Methylparaben (UNII: A2I8C7HI9T) | |
| Propylene Glycol (UNII: 6DC9Q167V3) | |
| Propylparaben (UNII: Z8IX2SC1OH) | |
| Trisodium Citrate Dihydrate (UNII: B22547B95K) | |
| Sucralose (UNII: 96K6UQ3ZD4) | |
| Water (UNII: 059QF0KO0R) | |

Product Characteristics

| | | | |
|-----------------|--------|---------------------|--|
| Color | ORANGE | Score | |
| Shape | | Size | |
| Flavor | ORANGE | Imprint Code | |
| Contains | | | |

Packaging

| # | Item Code | Package Description | Marketing Start Date | Marketing End Date |
|---|------------------|--|----------------------|--------------------|
| 1 | NDC:60505-6184-6 | 1 in 1 CARTON | 10/07/2021 | 04/30/2026 |
| 1 | | 473 mL in 1 BOTTLE, PLASTIC; Type 0: Not a Combination Product | | |

Marketing Information

| Marketing Category | Application Number or Monograph Citation | Marketing Start Date | Marketing End Date |
|--------------------|--|----------------------|--------------------|
| ANDA | ANDA211067 | 10/07/2021 | 04/30/2026 |

POTASSIUM CHLORIDE

potassium chloride solution

Product Information

| | | | |
|--------------------------------|-------------------------|---------------------------|----------------|
| Product Type | HUMAN PRESCRIPTION DRUG | Item Code (Source) | NDC:60505-6185 |
| Route of Administration | ORAL | | |

Active Ingredient/Active Moiety

| Ingredient Name | Basis of Strength | Strength |
|---|--------------------|-----------------|
| Potassium Chloride (UNII: 660YQ98I10) (Potassium Cation - UNII:295O53K152, Chloride Ion - UNII:Q32ZN48698) | Potassium Chloride | 40 meq in 15 mL |

Inactive Ingredients

| Ingredient Name | Strength |
|---|----------|
| Anhydrous Citric Acid (UNII: XF417D3PSL) | |
| FD&C Yellow NO. 6 (UNII: H77VEI93A8) | |
| Glycerin (UNII: PDC6A3C00X) | |
| Methylparaben (UNII: A218C7HI9T) | |
| Propylene Glycol (UNII: 6DC9Q167V3) | |
| Propylparaben (UNII: Z8IX2SC1OH) | |
| Trisodium Citrate Dihydrate (UNII: B22547B95K) | |
| Sucralose (UNII: 96K6UQ3ZD4) | |
| Water (UNII: 059QF0KO0R) | |

Product Characteristics

| | | | |
|-----------------|--------|---------------------|--|
| Color | ORANGE | Score | |
| Shape | | Size | |
| Flavor | ORANGE | Imprint Code | |
| Contains | | | |

Packaging

| # | Item Code | Package Description | Marketing Start Date | Marketing End Date |
|---|------------------|--|----------------------|--------------------|
| 1 | NDC:60505-6185-6 | 1 in 1 CARTON | 10/07/2021 | 08/31/2023 |
| 1 | | 473 mL in 1 BOTTLE, PLASTIC; Type 0: Not a Combination Product | | |

Marketing Information

| Marketing Category | Application Number or Monograph Citation | Marketing Start Date | Marketing End Date |
|--------------------|--|----------------------|--------------------|
| ANDA | ANDA211067 | 10/07/2021 | 08/31/2023 |

Labeler - Apotex Corp (845263701)**Registrant** - Apotex Inc (209429182)