DRUG QUANTITY MANAGEMENT POLICY – PER RX

POLICY: Pulmonary Arterial Hypertension – Sildenafil Drug Quantity Management Policy – Per Rx
- LiQrev® (sildenafil oral suspension – CMP)
- Revatio® (sildenafil tablets and powder for suspension – Pfizer, generic)

REVIEW DATE: 02/08/2023; selected revision 05/24/2023

INSTRUCTIONS FOR USE

THE FOLLOWING COVERAGE POLICY APPLIES TO HEALTH BENEFIT PLANS ADMINISTERED BY CIGNA COMPANIES. CERTAIN CIGNA COMPANIES AND/OR LINES OF BUSINESS ONLY PROVIDE UTILIZATION REVIEW SERVICES TO CLIENTS AND DO NOT MAKE COVERAGE DETERMINATIONS. REFERENCES TO STANDARD BENEFIT PLAN LANGUAGE AND COVERAGE DETERMINATIONS DO NOT APPLY TO THOSE CLIENTS. COVERAGE POLICIES ARE INTENDED TO PROVIDE GUIDANCE IN INTERPRETING CERTAIN STANDARD BENEFIT PLANS ADMINISTERED BY CIGNA COMPANIES. PLEASE NOTE, THE TERMS OF A CUSTOMER'S PARTICULAR BENEFIT PLAN DOCUMENT [GROUP SERVICE AGREEMENT, EVIDENCE OF COVERAGE, CERTIFICATE OF COVERAGE, SUMMARY PLAN DESCRIPTION (SPD) OR SIMILAR PLAN DOCUMENT] MAY DIFFER SIGNIFICANTLY FROM THE STANDARD BENEFIT PLANS UPON WHICH THESE COVERAGE POLICIES ARE BASED. FOR EXAMPLE, A CUSTOMER'S BENEFIT PLAN DOCUMENT MAY CONTAIN A SPECIFIC EXCLUSION RELATED TO A TOPIC ADDRESSED IN A COVERAGE POLICY. IN THE EVENT OF A CONFLICT, A CUSTOMER'S BENEFIT PLAN DOCUMENT ALWAYS SUPERSEDES THE INFORMATION IN THE COVERAGE POLICIES. IN THE ABSENCE OF A CONTROLLING FEDERAL OR STATE COVERAGE MANDATE, BENEFITS ARE ULTIMATELY DETERMINED BY THE TERMS OF THE APPLICABLE BENEFIT PLAN DOCUMENT. COVERAGE DETERMINATIONS IN EACH SPECIFIC INSTANCE REQUIRE CONSIDERATION OF 1) THE TERMS OF THE APPLICABLE BENEFIT PLAN DOCUMENT IN EFFECT ON THE DATE OF SERVICE; 2) ANY APPLICABLE LAWS/REGULATIONS; 3) ANY RELEVANT COLLATERAL SOURCE MATERIALS INCLUDING COVERAGE POLICIES AND; 4) THE SPECIFIC FACTS OF THE PARTICULAR SITUATION. COVERAGE POLICIES RELATE EXCLUSIVELY TO THE ADMINISTRATION OF HEALTH BENEFIT PLANS. COVERAGE POLICIES ARE NOT RECOMMENDATIONS FOR TREATMENT AND SHOULD NEVER BE USED AS TREATMENT GUIDELINES. IN CERTAIN MARKETS, DELEGATED VENDOR GUIDELINES MAY BE USED TO SUPPORT MEDICAL NECESSITY AND OTHER COVERAGE DETERMINATIONS.

CIGNA NATIONAL FORMULARY COVERAGE:

OVERVIEW

Sildenafil (Revatio, generic) and LiQrev are phosphodiesterase type 5 (PDE5) inhibitors indicated for the treatment of **pulmonary arterial hypertension** ([PAH] World Health Organization [WHO] Group I) **in adults** to improve exercise ability and delay clinical worsening.[1,10] Sildenafil (Revatio, generic) is also indicated for **PAH** (WHO Group I) in **patients 1 to 17 years of age** to improve exercise ability and, in pediatric patients too young to perform standardized exercise testing, pulmonary hemodynamics thought to underlie improvements in exercise.[1] Due to marketing exclusivity rights, LiQrev is not labeled with information for pediatric use.[10]

Dosing

Adult Dosing

The recommended dose of sildenafil (Revatio, generic) for the treatment of PAH in adults is 20 mg three times daily (TID). The dose may be titrated to a maximum of 80 mg TID, if required, based on symptoms and tolerability. In clinical trials,

sildenafil doses of 25 mg twice daily to 100 mg five times daily have been used for PAH.[2-5]

The recommended dose of LiQrev for the treatment of PAH in adults is 20 mg TID.[10]

Pediatric Dosing
The recommended dose of sildenafil (Revatio, generic) for the treatment of PAH in pediatric patients is based on weight (Table 1).[1]

Table 1. Sildenafil (Revatio, generic) Recommended Dosing in Pediatric Patients.[1]

Patient Weight	Recommended Dose
≤ 20 kg	10 mg TID
20 kg to 45 kg	20 mg TID
> 45 kg	20 mg TID[a]

TID – Three times daily; [a] A maximum dose in pediatric patients has not been identified. In patients weighing > 45 kg, the dose may be titrated to a maximum of 40 mg three times daily, if required, based on symptoms and tolerability.

Availability
Sildenafil (Revatio, generic) is available as 20 mg tablets and as 10 mg/mL powder for oral suspension in a 112 mL bottle (after reconstitution).[1] Revatio is also available as a 10 mg/12.5 mL vial which is not targeted in this policy.

LiQrev is available as a 10 mg/mL oral suspension in bottles of 122 mL.[10]

Off-Label Use
Sildenafil (Revatio, generic) has some data in patients with Raynaud's phenomenon at doses provided in strengths used for PAH.[6-9]

POLICY STATEMENT
This Drug Quantity Management program has been developed to promote the safe, effective, and economic use of sildenafil products for PAH. If the Drug Quantity Management rule is not met for the requested medication at the point of service, coverage will be determined by the Criteria below. All approvals are provided for 1 year in duration.

Drug Quantity Limits

Product	Strength and Form	Retail Maximum Quantity per Rx	Home Delivery Maximum Quantity per Rx
LiQrev® (sildenafil oral suspension)	10 mg/mL oral suspension (122 mL)	122 mL (1 bottle)	366 mL (3 bottles)
Revatio® (sildenafil tablets and oral suspension, generic)	20 mg tablets (bottles of 90 tablets)	90 tablets	270 tablets
	10 mg/mL oral suspension (when reconstituted) [112 mL bottle]	112 mL (1 bottle)	336 mL (3 bottles)

Pulmonary Arterial Hypertension – Sildenafil Drug Quantity Management Policy – Per Rx product(s) is(are) covered as medically necessary when the following criteria is(are) met.

CRITERIA

Sildenafil 20 mg tablets (Revatio, generic)

1. If the patient is prescribed greater than 20 mg three times daily for pulmonary arterial hypertension (PAH) or Raynaud's phenomenon, approve a quantity sufficient for a 30-day supply at retail or a 90-day supply at home delivery.

Sildenafil 10 mg/mL oral suspension (Revatio, generic) and LiQrev 10 mg/mL oral suspension

1. Approve a quantity sufficient for a 30-day supply at retail or a 90-day supply at home delivery if the patient meets the following criteria (A, B, and C):
Note: Round up to accommodate a whole package size (e.g., if the required dose is 20 mg three times daily [2 mL three times daily or 6 mL per day], 180 mL would be required for 30 days and 540 mL would be required for 90 days. Therefore, for sildenafil 10 mg/mL oral suspension [Revatio, generic], 224 mL [2 bottles] would be approved at retail or 560 mL [5 bottles] would be approved at home delivery. For LiQrev, 244 mL [2 bottles] would be approved at retail or 610 mL [5 bottles] would be approved at home delivery).
   A) Patient meets ONE of the following (i or ii):
      i. Patient has a diagnosis of pulmonary arterial hypertension (PAH); OR
      ii. Patient has a diagnosis of Raynaud's phenomenon; AND
   B) Patient is prescribed greater than 10 mg three times daily; AND
   C) Patient is unable to swallow a 20 mg sildenafil tablet (Revatio, generic).

CONDITIONS NOT COVERED

Any other exception is considered not medically necessary, including the following (this list may not be all inclusive):

1. Erectile dysfunction or sexual dysfunction.

REFERENCES

1. Revatio tablets, oral suspension [prescribing information]. Morgantown, WV: Viatris; January 2023.
2. Vazquez ZGS, Klinger JR. Guidelines for the treatment of pulmonary arterial hypertension. *Lung*. 2020;198:581-596.
3. Ruopp NF, Cockrill BA. Diagnosis and treatment of pulmonary arterial hypertension. A review. *JAMA*. 2022;327(14):1379-1391.
4. Klinger JR, Elliott CG, Levine DJ, et al. Therapy for pulmonary arterial hypertension in adults. Update of the CHEST guideline and Expert Panel Report. *CHEST*. 2019;155(3):565-586.
5. Humbert M, Kovacs G, Hoeper MM, et al, for the ESC/ERS Scientific Document Group. 2022 ESC/ERS guidelines for the diagnosis and treatment of pulmonary hypertension. *Eur Heart J*. 2022;43(38):3618-3731.

6. Roustit M, Blaise S, Allanore Y, et al. Phosphodiesterase-5 inhibitors for the treatment of secondary Raynaud's phenomenon: systematic review and meta-analysis of randomized trials. *Ann Rheum Dis*. 2013;72:1696-1699.
7. Shenoy PD, Kumar S, Jha LK, et al. Efficacy of tadalafil in secondary Raynaud's phenomenon resistant to vasodilatory therapy: a double-blind, randomized, crossover trial. *Rheumatol*. 2010;49:2420-2428.
8. Fernandez-Codina A, Canas-Ruano E, Pope JE. Management of Raynaud's phenomenon in systemic sclerosis-a practical approach. *J Scleroderm Relat Disord*. 2019;4(2):102-110.
9. Hinze AM, Wigley FM. Pharmacotherapy options in the management of Raynaud's phenomenon. *Curr Treat Opt Rheumatol*. 2018;4(3):235-254.
10. LiQrev oral suspension [prescribing information]. Farmville, NC: CMP Pharma; April 2023.

HISTORY

Type of Revision	Summary of Changes	Review Date
Annual Revision	**Revatio tablets (generic): Pulmonary Arterial Hypertension**. Criteria were modified to allow exception for patients who are prescribed greater than 20 mg three times daily. Criteria requiring documentation of a dose of greater than 20 mg three times daily for patients titrating their dose, criteria for patients already on a dose of 20 mg three times daily, and criteria for patients taking greater than 20 mg three times daily were removed. **Revatio for oral suspension (generic): Pulmonary Arterial Hypertension**. Criteria were modified to allow exception for patients who are prescribed greater than 10 mg three times daily. Criteria requiring documentation of a dose of greater than 10 mg three times daily for patients titrating their dose, criteria for patients already on a dose of 10 mg three times daily, and criteria for patients taking greater than 10 mg three times daily were removed.	03/09/2022
Early Annual Revision	Policy was updated to reflect the existing quantity limits when a product is obtained via home delivery. Approval duration was changed from 3 years to 1 year. **Sildenafil 20 mg tablets (Revatio, generic):** Clinical exception criteria were updated to approve for a patient who is prescribed greater than 20 mg three times daily for Raynaud's phenomenon (previously only approved for a patient with pulmonary arterial hypertension). **Sildenafil 10 mg/mL oral suspension (Revatio, generic):** Clinical exception criteria were updated to approve for a patient who is prescribed greater than 10 mg three times daily, is unable to swallow a 20 mg tablet, and has a diagnosis of Raynaud's phenomenon (previously only approved for a patient with a diagnosis of pulmonary arterial hypertension).	02/08/2023
Update	02/21/2023: No change to criteria. Policy updated to reflect expanded age indication of sildenafil (Revatio, generic) in pediatric patients.	NA
Selected Revision	**LiQrev:** New quantity limit added to the policy of 122 mL (1 bottle) per dispensing at retail and 366 mL (3 bottles) per dispensing at home delivery. An exception to provide a quantity sufficient for a 30-day supply at home delivery or a 90-day supply at retail was added for a patient with pulmonary arterial hypertension or Raynaud's phenomenon who requires a dose greater than 10 mg	05/24/2023

	three times daily and is unable to swallow a 20 mg sildenafil tablet (Revatio, generic).	

NA – Not applicable.