

**Clinical Policy: ACEI and ARB Duplicate Therapy** 

Reference Number: CP.PMN.61

Effective Date: 08.01.14 Last Review Date: 05.25 Line of Business: Medicaid

**Revision Log** 

See <u>Important Reminder</u> at the end of this policy for important regulatory and legal information.

#### **Description**

Angiotensin-converting enzyme inhibitors (ACEIs) and angiotensin receptor blockers (ARBs) act on the renin-angiotensin system. Concurrent use of an ACEI and an ARB is considered duplicate therapy.

### FDA Approved Indication(s)

Most ACEIs and ARBs are indicated for the treatment of hypertension and heart failure. Some are also indicated for diabetic nephropathy, myocardial infarction prophylaxis, proteinuria, reduction of cardiovascular mortality, and stroke prophylaxis.

### Policy/Criteria

Provider must submit documentation (such as office chart notes, lab results or other clinical information) supporting that member has met all approval criteria.

It is the policy of health plans affiliated with Centene Corporation<sup>®</sup> that ACEI and ARB duplicate therapy is **medically necessary** when the following criteria are met:

### I. Initial Approval Criteria

- A. ACEI and ARB Duplicate Therapy (must meet all):
  - 1. Member meets one of the following (a or b):
    - a. All of the following (i, ii, and iii):
      - i. Member is currently receiving ACEI and ARB combination therapy for chronic heart failure:
      - ii. Member has received the combination for at least 30 days;
      - iii. Member is responding positively to therapy;
    - b. Member is being titrated to, or tapered from, another ACEI or ARB;
  - 2. Provider documents that member is aware of duplicative therapy;
  - 3. Dose does not exceed the FDA-approved maximum recommended dose for the relevant ACEI and ARB.

#### **Approval duration:**

**Chronic heart failure** – 12 months

**Cross-taper** – 3 months

#### B. Other diagnoses/indications: Not applicable



### **II. Continued Therapy**

### A. ACEI and ARB Duplicate Therapy (must meet all):

- 1. Currently receiving ACEI and ARB combination therapy via Centene benefit or member has previously met initial approval criteria;
- 2. Member is responding positively to therapy;
- 3. For cross-taper requests (titrated to, or tapered from, another ACEI or ARB), member has not received more than 6 months of ACEI and ARB combination therapy;
- 4. If request is for a dose increase, new dose does not exceed the FDA-approved maximum recommended dose for the relevant ACEI and ARB.

#### **Approval duration:**

**Chronic heart failure** – 12 months

Cross-taper – 3 months (limited to 6 months total)

#### B. Other diagnoses/indications: Not applicable

#### III. Diagnoses/Indications for which coverage is NOT authorized: Not applicable

#### IV. Appendices/General Information

Appendix A: Abbreviation/Acronym Key

ACEI: angiotensin-converting enzyme inhibitor

ARB: angiotensin receptor blocker FDA: Food and Drug Administration

Appendix B: Therapeutic Alternatives

Not applicable

#### V. Dosage and Administration

Dosage and Administration		
Drug Name	Maximum Dose	
ACEIs		
Benazepril (Lotensin®)	80 mg/day	
Captopril	450 mg/day	
Enalapril (Vasotec®, Epaned®)	40 mg/day	
Fosinopril	80 mg/day	
Lisinopril (Zestril <sup>®</sup> , Qbrelis <sup>®</sup> )	80 mg/day	
Moexipril	30 mg/day	
Perindopril	16 mg/day	
Quinapril (Accupril®)	80 mg/day	
Ramipril (Altace®)	20 mg/day	
Trandolapril (Mavik®)	8 mg/day	
ARBs		
Azilsartan (Edarbi®)	80 mg/day	
Candesartan (Atacand®)	32 mg/day	
Irbesartan (Avapro®)	300 mg/day	
Losartan (Cozaar®, Arbli®)	100 mg/day	
Olmesartan (Benicar®)	40 mg/day	
Telmisartan (Micardis®)	80 mg/day	



Drug Name	Maximum Dose
Valsartan (Diovan®)	320 mg/day
Neprilysin Inhibitor/ARB	
Entresto® (sacubitril/valsartan)	194/206 mg/day

VI. Product Availability

Drug Name	Availability	
ACEIs		
Benazepril (Lotensin)	Tablets: 5 mg, 10 mg, 20 mg, 40 mg	
Captopril	Tablets: 12.5 mg, 25 mg, 50 mg, 100 mg	
Enalapril (Vasotec, Epaned)	Tablets: 2.5 mg, 5 mg, 10 mg, 20 mg	
	Oral solution: 1 mg/mL	
Fosinopril	Tablets: 10 mg, 20 mg, 40 mg	
Lisinopril (Zestril, Qbrelis)	Tablets: 2.5 mg, 5 mg, 10 mg, 20 mg, 30 mg, 40 mg	
	Oral solution: 1 mg/mL	
Moexipril	Tablets: 7.5 mg, 15 mg	
Perindopril	Tablets: 2 mg, 4 mg, 8 mg	
Quinapril (Accupril)	Tablets: 5 mg, 10 mg, 20 mg, 40 mg	
Ramipril (Altace)	Capsules: 1.25 mg, 2.5 mg, 5 mg, 10 mg	
Trandolapril (Mavik)	Tablets: 1 mg, 2 mg, 4 mg	
ARBs		
Azilsartan (Edarbi)	Tablets: 40 mg, 80 mg	
Candesartan (Atacand)	Tablets: 4 mg, 8 mg, 16 mg, 32 mg	
Irbesartan (Avapro)	Tablets: 75 mg, 150 mg, 300 mg	
Losartan (Cozaar, Arbli)	Tablets: 25 mg, 50 mg, 100 mg	
	Oral suspension: 10 mg/mL	
Olmesartan (Benicar)	Tablets: 5 mg, 20 mg, 40 mg	
Telmisartan (Micardis)	Tablets: 20 mg, 40 mg, 80 mg	
Valsartan (Diovan)	Tablets: 40 mg, 80 mg, 160 mg, 320 mg	
Neprilysin Inhibitor/ARB		
Entresto (sacubitril/valsartan)	Tablets: 24/26 mg, 49/51 mg, 97/103 mg	

#### VII. References

- 1. Clinical Pharmacology [database online]. Philadelphia, PA: Elsevier. Updated periodically. Available at: http://www.clinicalkey.com/pharmacology. Accessed January 23, 2025.
- 2. Zhao M, Qu H, Wang R, et al. Efficacy and safety of dual vs single renin-angiotension system blockage in chronic kidney disease: an updated meta-analysis of randomized controlled trials. *Medicine*. 2021 Sept 3; 100(35): e26544.
- 3. Makani H, Bangalore S, Desouza KA, Shah A, Messerli FH. Efficacy and safety of dual blockade of the renin-angiotensin system: meta-analysis of randomised trials. *BMJ*. 2013;346:f360.
- 4. Fried LF, Emanuele N, Zhang JH, et al. Combined angiotensin inhibitor treatment of diabetic nephropathy. *N Engl J Med.* 2013; 369(20).



- 5. Mann JF, Schmieder RE, McQueen M, et al. Renal outcomes with telmisartan, ramipril, or both, in people at high vascular risk (the ONTARGET study); a multicentre, randomized, double-blind, controlled trial. *Lancet*. 2008;372(9638):547-53.
- 6. James PA, Oparil S, Carter BL, et al. 2014 evidence-based guideline for the management of high blood pressure in adults: report from the panel members appointed to the eighth Joint National Committee (JNC 8). *JAMA*. 2014;311(5):507-520. doi:10.1001/jama.2013.284427.

Reviews, Revisions, and Approvals	Date	P&T Approval Date
2Q 2021 annual review: no significant changes; references reviewed and updated.	02.15.21	05.21
2Q 2022 annual review: no significant changes; references reviewed and updated.	01.31.22	05.22
2Q 2023 annual review: no significant changes; references reviewed and updated.	01.19.23	05.23
2Q 2024 annual review: no significant changes; removed eprosartan due to product discontinuation; references reviewed and updated.	01.19.24	05.24
2Q 2025 annual review: no significant changes; removed discontinued brand products from section V and VI; references reviewed and updated.  RT4: added new oral suspension formulation of losartan, Arbli, to policy.	03.18.25	05.25

#### **Important Reminder**

This clinical policy has been developed by appropriately experienced and licensed health care professionals based on a review and consideration of currently available generally accepted standards of medical practice; peer-reviewed medical literature; government agency/program approval status; evidence-based guidelines and positions of leading national health professional organizations; views of physicians practicing in relevant clinical areas affected by this clinical policy; and other available clinical information. The Health Plan makes no representations and accepts no liability with respect to the content of any external information used or relied upon in developing this clinical policy. This clinical policy is consistent with standards of medical practice current at the time that this clinical policy was approved. "Health Plan" means a health plan that has adopted this clinical policy and that is operated or administered, in whole or in part, by Centene Management Company, LLC, or any of such health plan's affiliates, as applicable.

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#### Note:

**For Medicaid members**, when state Medicaid coverage provisions conflict with the coverage provisions in this clinical policy, state Medicaid coverage provisions take precedence. Please refer to the state Medicaid manual for any coverage provisions pertaining to this clinical policy.

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