## Recommendation for drug combination in uncomplicated hypertension<sup>†</sup> \*Preferred Combination Acceptable Combination NOT Preferred Combination

• Thiazide + K+-sparing diuretic

β-blocker + diuretic or dihydropyridine CCB

tahs

tabs

tabs

tabs

tabs

tabs

tabs

tabs

tabs

ANGIOTENSIN II RECEPTOR BLOCKER + CALCIUM CHANNEL BLOCKER (DIHYDROPYRIDINE) +

1 tab daily.

Form Usual Adult Dose

Aliskiren + thiazide or CCB

CCB + thiazide

**COMBINATION HYPERTENSION TREATMENTS** (Part 1 of 4)

 ACEI + thiazide ACEI + dihydropyridine CCB ARB + thiazide ARB + dihvdropvridine CCB

Generic

benazepril/HCTZ

captopril/HCTZ

enalapril/HCTZ

lisinopril/HCTZ

moexipril/HCTZ

quinapril/HCTZ

trandolapril/

olmesartan/

amlodipine/HCTZ

verapamil (ext-rel)

THIAZIDE DIURETIC

Brand

**HCT** 

Lotensin

Capozide

Vaseretic

Zestoretic

Accuretic

Tarka

Tribenzor

Strenath \*ACE INHIBITOR + THIAZIDE DIURETIC 5mg/6.25mg,

10mg/12.5mg, 20mg/12.5mg, 20mg/25mg

25mg/15mg, 25mg/25mg, 50mg/15mg, 50mg/25mg

10mg/25mg

10mg/12.5mg, 20mg/12.5mg+

10mg/12.5mg,

20mg/12.5mg,

7.5ma/12.5ma.

15mg/12.5mg,

10mg/12.5mg+,

20mg/12.5mg+,

ACE INHIBITOR + CALCIUM CHANNEL BLOCKER (DIPHENYLALKYLAMINE)

20mg/5mg/12.5mg,

40mg/5mg/12.5mg,

40mg/10mg/12.5mg, 40mg/10mg/25mg

40mg/5mg/25mg,

1mg/240mg,

2mg/180mg,

2mg/240mg, 4mg/240mg

15mg/25mg

20mg/25mg

20mg/25mg

enalapril/day and 50mg HCTZ/day. Allow 2-3wks for titration of HCTZ component. Or, substitute for individually titrated components. Not for initial therapy. Initially 10mg/12.5mg or 20mg/12.5mg; increase HCTZ dose 2-3wks after. Max 80mg/50mg daily. CrCl <30mL/min: not recommended. Switching from monotherapy with either component:

initally 10/12.5 or 20/12.5 once daily, then after 2-3wks

titrate HCTZ component. If on diuretic: if possible, suspend diuretic for 2-3 days, then adjust. Or, substitute for

individually titrated components.

scored : Take 1hr before a meal. Switching from monotherapy

with either component: 1 tab once daily; adjust at

Not for initial therapy. Previously titrated: use same

doses as individual components. Switching from

Titrate individual components, Take with food.

One tab once daily. May titrate at 2-week intervals;

impairment: start amlodipine 2.5mg.

max one 40/10/25mg tab daily. ≥75yrs or severe hepatic

2-3-week intervals; usual max 30mg/50mg per day. Or, substitute for individually titrated components.

quinapril monotherapy: initially one 10/12.5 tab or one 20/12.5 tab once daily; allow 2-3wks before increasing HCTZ component. Switching from HCTZ 25mg/day monotherapy: initially one 10/12.5 tab daily or one 20/12.5 tab once daily. Adjust based on response and serum potassium. CrCl ≤30mL/min: not recommended.

ACEI + ARB

scored : To switch from benazepril monotherapy: initially

β-blocker + ACEI or ARB

10/12.5mg once daily. Or, titrate individual components.

scored: Take 1hr before meals. As initial therapy: one 25/15 tab

daily; adjust at 6wk intervals. Previously titrated:

β-blocker + nondihydropyridine CCB

β-blocker + central acting (clonidine, etc)

Switching from monotherapy with either component: start with 10/25 once daily, then adjust; max 20mg

use same doses as individual components. Usual max 150mg captopril, 50mg HCTZ daily.

(continued)

<b>COMBINATION HYPERTENSION TREATMENTS</b> (Part 2 of 4)						
Generic	Brand	Strength	Form	Usual Adult Dose		
ANGIOTENSIN II RECEPTOR BLOCKER + CALCIUM CHANNEL BLOCKER (DIHYDROPYRIDINE) + THIAZIDE DIURETIC (continued)						
valsartan/ amlodipine/HCTZ	Exforge HCT	160mg/5mg/25mg, 160mg/10mg/12.5mg, 160mg/10mg/25mg, 320mg/10mg/25mg	tabs	One tab daily. Titrate at 2-week intervals; max one 320mg/10mg/25mg tab daily. <i>Replacement therapy:</i> may be substituted for individually titrated components. <i>Addon/switch therapy:</i> may be used to provide additional BP lowering for patients not adequately controlled on doses of any two antihypertensive classes: ARBs, CCBs, and diuretics.		
*ANGIOTENSIN II RECEPTOR BLOCKER + THIAZIDE DIURETIC						
azilsartan/ chlorthalidone	Edarbyclor	40mg/12.5mg, 40mg/25mg	tabs	Initially 40/12.5mg once daily. May increase to 40/25mg after 2–4wks as needed. <i>Max</i> : 40/25mg. <i>Patients titrated to the individual components:</i> may give corresponding dose of Edarbyclor. See full labeling.		
candesartan/HCTZ	Atacand HCT	16mg/12.5mg, 32mg/12.5mg, 32mg/25mg	scored tabs	Not for initial therapy. May be substituted for titrated components. <i>BP not controlled on HCTZ 25mg once daily, or controlled but serum potassium decreased</i> : one 16/12.5 tab once daily. <i>BP not controlled on candesartan 32mg per day</i> : initially one 32/12.5 tab once daily; may increase to 32/25 once daily. <i>CrCl ≤30mL/min</i> : not recommended.		
irbesartan/HCTZ	Avalide	150mg/12.5mg, 300mg/12.5mg	tabs	Take once daily. Not controlled on monotherapy: initially 150/12.5mg, titrate to 300/12.5mg then 300/25mg if needed. Replacement therapy: may be substituted for titrated components. Initial therapy: start at 150/12.5mg for 1−2wks, then titrate as needed up to max 300/25mg. Maximum effects within 2−4wks after dose change. CrCl ≤30mL/min: not recommended.		
losartan/HCTZ	Hyzaar	50mg/12.5mg, 100mg/12.5mg, 100mg/25mg	tabs	One 50/12.5mg tab once daily; may increase after about 3wks (2–4wks for severe HTN) to two 50/12.5mg tabs once daily or one 100/25mg tab once daily. <i>Titrate components</i> : see literature. <i>HTN with LVH</i> : switch from losartan monotherapy (see literature). <i>CrCl&lt;30mL/min</i> : not recommended.		
olmesartan/HCTZ	Benicar HCT	20mg/12.5mg, 40mg/12.5mg, 40mg/25mg	tabs	Not for initial therapy. May be substituted for titrated components. Individualize. BP not controlled on olmesartan or HCTZ alone: one tab once daily; may titrate at 2–4-week intervals; usual max 40mg/25mg once daily. CrCl <30mL/min: not recommended. Volume depleted: reduce dose.		
telmisartan/HCTZ	Micardis HCT	40mg/12.5mg, 80mg/12.5mg, 80mg/25mg	tabs	Not for initial therapy. May be substituted for titrated components. <i>BP not controlled on telmisartan</i> 80mg/day: one 80mg/12.5mg tab once daily; may titrate to 160mg/25mg. <i>BP not controlled on HCTZ 25mg/day</i> : one 80mg/12.5mg tab or 80mg/25mg tab once daily; may titrate to 160mg/25mg if BP uncontrolled after 2–4wks.		

valsartan/HCTZ

BP controlled on HCTZ 25mg/day but hypokalemic: One 80mg/12.5mg tab once daily. CrCl ≤30mL/min: not recommended. Hepatic insufficiency or biliary obstruction: initially one 40mg/12.5mg tab once daily; monitor closely. Severe renal or hepatic impairment: not recommended.

Take once daily. Add-on or initial therapy and not

volume-depleted: Initially 160mg/12.5mg; may increase after 1-2wks up to max 320mg/25mg. Replacement

therapy: may be substituted for the titrated components.

Maximum effects within 2-4wks after dose change.

CrCl ≤30mL/min: not recommended. (continued)

tabs

Diovan HCT 80mg/12.5mg,

160mg/12.5mg,

320mg/12.5mg,

160mg/25mg,

320mg/25mg

COMBINATION HYPERTENSION TREATMENTS (Part 3 of 4)							
Generic	Brand	Strength	Form	Usual Adult Dose			
BETA-BLOCKER + THIAZIDE DIURETIC							
atenolol/ chlorthalidone		50mg/25mg+, 100mg/25mg		Switching from monotherapy: initially one 50mg/25mg tab daily, may increase to one 100mg/25mg tab daily.  CrCl 15—35mL/min: max 50mg atenolol/day. CrCl <15mL/min: max 50mg atenolol every other day.			
bisoprolol/HCTZ		2.5mg/6.25mg, 5mg/6.25mg, 10mg/6.25mg		Initially one 2.5mg/6.25mg tab once daily. Adjust at 14- day intervals; max two 10mg/6.25mg tabs once daily.			
	HCT	50mg/25mg, 100mg/25mg, 100mg/50mg	scored tabs	Titrate individual components.			
metoprolol succinate extended- release/HCTZ	-	25mg/12.5mg, 50mg/12.5mg, 100mg/12.5mg		Individualize. Take once daily. Titrate as needed every 2wks up to max 200/25mg. Severe renal impairment: not recommended. Moderate hepatic impairment: consider initiating with lower HCTZ component.			

tahs

caps

tabs

tabs

\*CALCIUM CHANNEL BLOCKER (DIHYDROPYRIDINE) + ANGIOTENSIN II RECEPTOR BLOCKER tahs

scored: Titrate components. Renally impaired: increase dosing

Not adequately controlled with dihydropyridine CCB.

2.5mg/10mg once daily; may titrate up to 10mg/40mg

once daily if BP remains uncontrolled. Replacement

therapy: may be substituted for titrated components.

Initially 3.5mg/2.5mg once daily. Adjust at 7–14 day

Take once daily. Initial therapy: initially 5/20mg;

may increase after 1-2wks up to max 10mg/40mg;

≥75yrs old or hepatic impairment: not recommended.

Replacement therapy: may be substituted for titrated components. Add-on therapy: may be used if not controlled on monotherapy. Individualize; titrate at 2-week intervals up to max 10/40mg once daily, usually by increasing dose of one or both components if BP not controlled on prior therapy. Maximum effects within 2wks

Take once daily. Initial therapy: 5/40mg or 5/80mg; may

therapy: may be used if not controlled on monotherapy; if dose-limiting adverse reactions with amlodipine 10mg,

(continued)

titrate at 2-week intervals to max 10/80mg. Add-on

switch to 5/40mg tab. Replacement therapy: may be substituted for the titrated components. Renal and/or hepatic impairment: titrate slower. ≥75yrs, or hepatic impairment: not for initial use (initially use amlodipine alone, or add amlodipine 2.5mg to telmisartan).

ACE inhibitor, unable to achieve BP control with

amlodipine without developing edema: Initially

CrCl < 30ml/min: not recommended

intervals: max 14mg/10mg once daily.

interval: see literature.

scored: Titrate individual components.

after dose change.

nadolol/

bendroflumethiazide

propranolol/HCTZ

amlodipine/

amlodipine/

perindopril

amlodipine/

olmesartan

amlodipine/

telmisartan

benazepril

Corzide

Inderide

Lotrel

**Prestalia** 

Azor

Twynsta

40ma/5ma.

80mg/5mg

40ma/25ma

\*CALCIUM CHANNEL BLOCKER (DIHYDROPYRIDINE) + ACE INHIBITOR

2.5mg/10mg,

5ma/10ma.

5mg/20mg,

5mg/40mg,

10mg/20mg,

10ma/40ma

2.5mg/3.5mg,

5ma/7ma.

10ma/14ma

5ma/20ma.

5mg/40mg,

10mg/20mg, 10mg/40mg

5mg/40mg,

5mg/80mg,

10mg/40mg,

10mg/80mg

**COMBINATION HYPERTENSION TREATMENTS** (Part 4 of 4) Generic Brand Strenath Form Usual Adult Dose \*CALCIUM CHANNEL BLOCKER (DIHYDROPYRIDINE) + ANGIOTENSIN II RECEPTOR BLOCKER (continued) Exforae amlodipine/ 5mg/160mg, tahs Take once daily. Initial therapy and not volume depleted: 5mg/320mg, valsartan Initially 5/160mg; may increase after 1-2wks up to 10ma/160ma. max 10/320mg. Add-on therapy: may be used if not 10mg/320mg controlled on monotherapy; if inadequate response after 3-4wks, may titrate up to max 10/320mg, Replacement therapy: may be substituted for the titrated components. Maximum effects within 2wks after dose change. Elderly. hepatic impairment: initial therapy not recommended. CENTRAL ALPHA-AGONIST + THIAZIDE DIURETIC methyldopa/HCTZ 250mg/15mg, tabs Titrate individual components. Initially one 250mg/15mg tab 2-3 times daily or one 250mg/25mg tab 2 times 250ma/25ma DIRECT RENIN INHIBITOR + CALCIUM CHANNEL BLOCKER (DIHYDROPYRIDINE) aliskiren/amlodipine : Tekamlo Take once daily, consistently with regard to meals. Initially 150mg/5mg, tabs

DIRECT RENIN INHIBITOR + CALCIUM CHANNEL BLOCKER (DIHYDROPYRIDINE) + THIAZIDE DIURETIC

tabs

tabs

scored tabs

tabs

caps

tahs

Not an inclusive list of medications, official indications, and/or dosing details. Please see drug monograph at www.eMPR.com

scored

150mg/10mg,

300mg/5mg,

300mg/10mg

150/5/12.5mg,

300/5/12.5mg,

300/10/12.5mg,

150mg/12.5mg

300ma/12.5ma

150mg/25mg

300mg/25mg

5mg/50mg

50mg/50mg+

37.5mg/25mg

37.5ma/25ma.

75mg/50mg

Aldactazide 25mg/25mg,

300/5/25mg,

300/10/25mg

Amturnide

DIRECT RENIN INHIBITOR + THIAZIDE DIURETIC

Tekturna

K+ SPARING DIURETIC + THIAZIDE DIURETIC

Dyazide

Maxzide

**Key:** HCTZ = hydrochlorothiazide; + = scored.

and/or contact company for full drug labeling.

**HCT** 

aliskiren/

amlodipine/HTCZ

aliskiren/HCT7

amiloride/HCTZ

spironolactone/

triamterene/HCTZ

**HCTZ** 

**NOTES** 

one 150mg/5mg tab daily. Add-on: switch when BP is not controlled with aliskiren or any DHP CCB alone. Replacement therapy: switch from previously-titrated components. Titrate at 2-4-week intervals (slow

titration in hepatic impairment or heart failure); max one

Take once daily. Titrate at 2-week intervals; max one

300/10/25mg tab daily. Replacement: may substitute

for individually titrated components. Add-on/switch: if

not adequately controlled on any two of the following:

aliskiren, dihydropyridine CCB, thiazide diuretics. May switch with a lower dose of any component that causes dose-limiting ADRs. ≥75yrs or severe hepatic impairment: initially amlodipine 2.5mg/day (not available). Concomitant simvastatin: see Interactions.

Take consistently with regard to meals. 1 tab once daily.

initially 150mg/12.5mg; may increase after 2-4wks up to

Initially 1 tab daily with food. May increase to 2 tabs daily

Usual maintenance: 50/100mg each of spironolactone

(Rev. 10/2017)

1-2 tabs of 37.5/25 daily or 1 tab of 75/50 daily.

and HCTZ daily in single or divided doses.

Add-on or initial therapy and not volume-depleted:

Replacement therapy: substitute for the titrated

300mg/10mg tab daily.

max 300mg/25mg.

in single or divided doses.

1-2 caps once daily.

components.