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# Liquid chromatographical methods for determination of selected antihypertensive drugs: a review

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# ABSTRACT

This work summarizes of the analytical methods reported in the literature for the separation and quantification using several liquid chromatographic methods for the determination of antihypertensives in biological matrices and pharmaceutical formulations have been discussed in the course of this review, namely RPHPLC, HPTLC and LC-MS methods; their selectivity, sensitivity, accuracy and reproducibility make them a good choice for analysis.

**KEY WORDS:** Hypertension, analysis, Liquid chromatography.

# INTRODUCTION

Hypertension is one of the most serious diseases of the XXI century concerning about 20-30% of the world population of adults. Early detection and proper pharmacotherapy of hypertension could decrease the risk of stroke, left ventricular hypertrophy, cerebral hemorrhage, cerebral vessel disease or peripheral artery disease (Szanajderman, 1998; Kupershmit, 1998). Diuretics, particularly thiazides and thiazide-like, loop diuretics and potassium sparing diuretics are applied in the hypertension treatment. From treatment perspective, complexes consisting of the selective and nonselective  $\beta$ -adrenergic receptor antagonists and  $\alpha$ -adrenergic receptor antagonists, vasodilators, calcium channel blockers, ACE inhibitors (from angiotensin-converting enzyme) and angiotensin receptor antagonists play significant role. Diversity of chemical structures present in a group of hypotensive drugs encourage for searching new methods useful in their quantitative analysis.In this review by taking into account, the number of liquid chromatography methods discussed previously by Stolarczykin (2010), that leads to the availability of fast, selective, sensitive, precise and accurate analytical methods for the quantitative determination of selected antihypertensive drugs in biological fluids and pharmaceutical formulations.

Liquid chromatographical methods: Generally in aqueous solutions, Antihypertensive drugs are likely to exist in the ionic form, use of an acidic mobile phase/addition of ion-pair reagents to the mobile phase is recommended in most cases present in this review. The use of other chromatographical procedures extended along with reversedphase liquid chromatography are High-performance thin-layer chromatography (HPTLC) is a different chromatographic technique that can also be successfully applied to separate analytes in a mixture. Paul (2011) developed High performance liquid Chromatography method for the simultaneous determination of Telmisartan and Amlodipine besylate combination in tablets.Patel (2011) developed High performance liquid Chromatography method for the simultaneous determination of Eprosartan and Hydrochlorothiazide combination in tablets. Liu (2011) introduced a HPLC method using MS/MS detection for Hydarlazine quantification in BALB/C mouse plasma and brain. Wankhede (2010) described two spectroscopic methods and High performance liquid Chromatography for the simultaneous determination of Losartan, Amlodipine besylate and Hydrochlorothiazide combination in tablets.Kavitha (2010) developed and validated a RP-HPLC method for the analysis of Atenolol, Hydrochlorothiazide and Losartan potassium in tablet formulation. Mustafa celebier (2010) developed and validated a HPLC method for the simultaneous estimation of Amlodipin and Valsartan in their combined dosage forms and for drug dissolution studies. Farouk (2010) developeda isocratic RP-HPLC method and subsequent validated for the analysis of Torasemide, Irbesartan and Olmesartanmedoxomil. Suresh Kumar (2010) developeda reversed-phase liquid chromatographic method for the simultaneous estimation of Atorvastatin Calcium and Telmisartan in tablet formulations.Patil (2010) developed a High-performance thin-layer chromatographic method for analysis of Telmisartan and Atorvastatin calcium in fixed dose combination. Bahia Moussa (2010) developed a reversed phase liquid chromatography and thin layer chromatography methods as a stability indicating assays of OlmesartanMedoxomil in presence of its acid or alkaline induced degradation products. Della Grace Thomas Parambi (2010) developeda quantitative HPTLC method for determination of OlmesartanMedoxomil in tablet dosage form. ZaveriMaitreyi (2010) developed a precise and validated RP-HPLC method for the simultaneous determination of Atenolol and Hydrochlorothiazide in tablet formulation. Wankhede (2009)described two

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spectroscopic methods and High performance liquid Chromatography for the simultaneous determination of Olmesartanmedoximil and Amlodipine besylate combination in tablets. Shah (2008) developed High performance liquid Chromatography method for the simultaneous determination of Atorvastatin calcium and Amlodipine besylate combination in tablets. Wankhede (2007) developed High performance liquid Chromatography for the simultaneous determination of Telmisartan and Hydrochlorothiazide combination in tablets. Palled (2005) developed High performance liquid Chromatography method for the determination of Telmisartan in tablets. Manes (1990) described HPLC method for determination of Hydralazine and its metabolite in human plasma using Methyl Red as Internal standard. Wong (1987) developed a specific and sensitive method for extraction and analysis of Hydralazine by High-performance liquid chromatography with electrochemical detection. Molles (1985) reported HPLC method which utilizes the derivatization product of Hydralazine with p-Hydroxybenzaldehyde or p-Anisaldehyde as an internal standard.

Ahs	Metho	Detection	Sample	Column	Mobile phase
	d		-		-
Telmisartan and Amlodipine besylate	HPLC	UV=237n m	Pharmaceutica ls	Symmetry C18 column (250 mm x 4.6 mm i.d, 5µm)	methanol, Acetonitrile, Potassium dihydrogen phosphate buffer (60:40; pH 4.0)
Eprosartan and Hydrochlorothiazide	HPLC	UV=240n m	Pharmaceutica ls	Phenomenex C18 column (250 mm x 4.6 mm i.d, 5µm)	0.5% formic acid, methanol, Acetonitrile (80:25:20; pH 2.8)
Hydralazine	LC-MS -MS	m/z 225.2 →129.5	mouse plasma and brain	Agilent ZORBAX SB- C18	0.01 mol/l methanol: ammonium acetate (60:40, v/v)
Losartan, Amlodipine besylate and Hydrochlorothiazide	HPLC	UV=238n m	Pharmaceutica ls	Kromasil C-18 (5µm, 250*4.6 mm)	Acetonitrile, 0.025 mol/L Potassium dihydrogen phosphate buffer (43:57; pH 3.7)
atenolol, hydrochlorothiazide and losartan potassium	HPLC	UV=270n m	Pharmaceutica ls	Phenomenex C18 (250 mm x 4.6 mm i.d, 5µm)	acetonitrile: 50mM potassium dihydrogenortho phosphate (pH-3.5) ratio 50:50v/v
Amlodipine and valsartan	HPLC	UV=240n m	Pharmaceutica ls	Waters Atlantis d C18 (250 mm x 4.6 mm. i.d., 5.0µm)	phosphate buffer (pH-3.6): acetonitrile: methanol (46:44:10v/v/v),
Torasemide, Irbesartan and Olmesartanmedoxomil	HPLC	UV=280n m	Pharmaceutica 1	Waters Atlantis d C18 (250 mm x 4.6 mm. i.d., 5.0µm)	phosphate buffer pH 3:acetonitrile (60:40,v/v), phosphate buffer pH 3.2:acetonitrile (60:40,v/v)
atorvastatin calcium and telmisartan	HPLC	UV=254n m	Pharmaceutica 1	Waters Symmetry C18 (250 mm x 4.6 mm. i.d., 5.0µm)	ammonium acetate (0.02M, pH 4.0 adjusted with glacial acetic acid) and acetonitrile in ratio (40:60 v/v)
Telmisartan and Atorvastatin calcium	HPTLC	Densitome tric	Pharmaceutica ls	Silica gel G 60 F <sub>254</sub> , HPTLC plates	toluene: methanol (7: 3, v/v)
Olmesartan	HPLC HPTLC	UV=257n m Densito- metric	Pharmaceutica ls	Agilent, Exclipse XDB- C18 column Silica gel G 60 F <sub>254</sub> , HPTLC plates	acetonitrile: methanol: water: glacial acetic acid (40:35:25:0.1 v/v/v/v) chloroform: methanol: formic acid (8:1.5:0.5 v/v/v)
Olmesartan	HPTLC	Densito- metric	Pharmaceutica ls	Silica gel G 60 F <sub>254</sub> HPTLC plates	chloroform: acetonitrile: toluene: glacial acetic acid, (1:8:1:0.1(v/v/v/v)
Atenolol and Hydrochlorothiazide	HPLC	UV=286n m	Pharmaceutica ls	Zorbax SB-CN (250 x 4.6 mm), 5µm	Water: Buffer: Methanol (50:35:15)
Olmesartan and Amlodipine besylate	HPLC	UV=238n m	Pharmaceutica ls	Kromasil C-18(5µm, 250*4.6 mm)	Acetonitrile 0.05 mol/L, Potassium dihydrogen phosphate buffer (50:50)

Table.1.Liquid	chromatography	methods for	determination	of Anti-Hypertensives

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Atorvastatin calcium and	HPLC	UV=240n	Pharmaceutica	Phenomenex C18	methanol, Acetonitrile,
Amlodipine besylate		m	ls	column (250 mm x 4.6	0.05mol/L, Potassium
				mm i.d, 5µm)	dihydrogen phosphate buffer
					(20:50:30; pH 3.5)
Telmisartan and	HPLC	UV=238n	Pharmaceutica	ODS Hypersil C-18	Acetonitrile 0.05 mol/L,
Hydrochlorothiazide		m	ls	(5µm,250*4.6 mm)	Potassium dihydrogen
					phosphate buffer (60:40; pH
					3.0)
Telmisartan	HPLC	UV=245n	Pharmaceutica	Hypersil BDS C-18	methanol, Acetonitrile (40:60)
		m	ls	(5µm,250*4.6 mm)	
Hydralazine	HPLC	UV=408n	Human	ODS-2 column packed	acetonitrile: aqueous
		m	plasma	with spherisorb	triethylamine phosphate buffer
				(250 × 4, 3 μm)	(80:20, v/v - pH 3)
Hydralazine	HPLC	Electro	Human	Supelcosil LC-18-DB	66% methanol in 0.055 <i>M</i>
		chemical	plasma	(5µm)	citric acid/0.02 M dibasic
		detection			sodium phosphate (pH 2.5).
Hydralazine	HPLC	UV=295n	Human	µBondapak Phenyl	methanol: 2% acetic acid
		m	plasma	column (30 cm x 3.9	solution (60:40, v/v)
				mm I.D.10 μm)	

#### CONCLUSION

This work is a comprehensive and critical review of the analytical methods reported in the literature for the determination of selected antihypertensive drugs in biological matrices and pharmaceutical formulation. Overall, it should be noted that a large number of liquid chromatographic methods have been reported. These methods constitute useful tools for pharmacokinetic and toxicological studies or for quality control tests. Moreover, some of them may support the routine therapeutic drug monitoring of these antihypertensive drugs in clinical practice.

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