

Methods of Analysis of Lisinopril: A Review

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Abstract

Lisinopril is an (ACE) angiotensin-converting enzyme inhibitor and is used in the treatment of high blood pressure, heart failure, and heart attacks. Numerous analytical methods have been developed for the determination of lisinopril. These methods include Chromatographic methods, UV spectrophotometric methods, IR, Polarographic, Stress degradation, titration and assay. This review outlines the applications of these techniques for the determination of lisinopril in pharmaceutical preparations.

Keywords: Chromatographic; Infrared (IR); Lisinopril; Polarographic; Spectrometric; Titration

Introduction

Lisinopril is an oral long-acting angiotensin converting enzyme inhibitor. It is a lysine-derivative of Enalaprilate and has structural similarity with its substrate. It differs from captopril by lacking the sulfhydryl group [1,2]. Lisinopril is chemically described as (S)-1-[N2-(1-carboxy-3-phenylpropyl)-L-lysyl]-L-proline dihydrate. Its empirical formula is C₂₁H₃₁N₃O₅·2H₂O.

Lisinopril is used in high blood pressure (hypertension), congestive heart failure, and in heart attack, also in renal and retinal complications of diabetes [2]. It also exhibits haemodynamic effects [3]. It is an active site directed inhibitor [4]. It promotes natriuresis and useful in preventing diabetic retinopathy in the patients of type II diabetes [4,5].

Its onset of action is 1-2 hours. Duration of action is 24 hours. Absorption of the lisinopril is slowly and moderately from GI tract (oral) and peak plasma concentration obtain after 7 hours. It has no interactions with drugs and foods. The drug distribution is up to 25%. It is excreted unchanged in urine and does not undergo metabolism. The drug is given orally in case of hypertension. Adult dose is initially 5-10 mg daily given at bedtime. Dose of drug in renovascular hypertension, volume depletion, severe hypertension in the beginning 2.5-5 mg once daily. In diuretic patients 5 mg once daily. Maintenance dose is 20 mg once daily up to 80 mg daily can be given. In case of children ≥ 6 years: initially up to 0.07 mg/kg (up to 5 mg once daily) can be given. Bioavailability of the drug is about 25% (Figure 1) [6].

Properties of lisinopril

Lisinopril is a white, crystalline powder, having a molecular weight of 441.52 and molar mass of 40.488 g/mol. Lisinopril is a white to off-white, crystalline powder, with a molecular weight of 441.53. It

is soluble in water and sparingly soluble in methanol and practically insoluble in ethanol, acetone, acetonitrile and chloroform [7].

Analytical Method

Literature review shows that there are developed methods including spectrophotometric, atomic absorption, HPLC and LC-MS method for the estimation of lisinopril [8].

The official analytical methods for Lisinopril described are potentiometric titration and HPLC various spectrophotometric methods, chromatographic methods of analysis such as micellar electrokinetic chromatography and gas liquid chromatography [9,10], Capillary electrophoresis, fluoroimmunoassay, radioimmunoassay and fluoroenzymatic assay have also been reported [11,12].

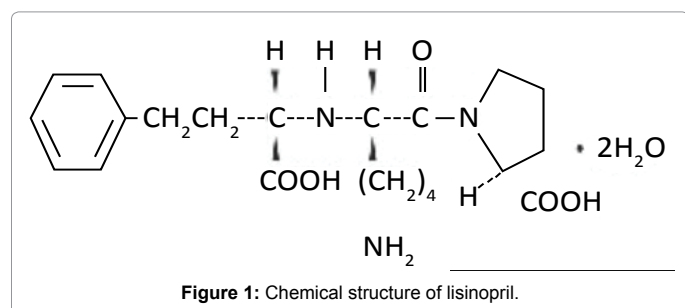
The available methods are associated with drawbacks such as low reliability due to isomerisation, less sensitive, measurement at lower wavelength, pH dependent, inaccessibility and requirement of expertise [13].

Literature survey reveals the availability of several methods by using various mixtures but no method was available on the mixture of distilled water and methanol 70:30 which was an unique method with better results [14].

Analytical data are used to screen the potential drugs in biological samples, support formulation. Studies assist in the development of medicine production, monitor the (API) bulk pharmaceuticals and finished products and also test final products for release [2].

Methods of Estimation for Lisinopril

There are various methods of analysis for lisinopril described in literature as,



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- 1) UV-visible spectroscopic methods
- 2) Spectrofluorimetric methods of estimation by reaction modifications
- 3) High performance liquid chromatographic methods
- 4) Combination of methods or techniques
- 5) Titrimetric methods
- 6) Miscellaneous methods

UV-visible spectroscopic methods

There are developed Spectrophotometric methods of analysis in single or in combination. Lisinopril shows absorption in UV-visible range in alkaline media produced by sodium hydroxide measured by absorption maxima method [15-17].

Quantitative estimation of Lisinopril in bulk and its marketed formulations by simple and sensitive UV spectrophotometric method have been developed which is accurate and economical.

Spectrophotometric method have been developed to determine the Lisinopril in unadulterated form or in formulation by coupling with o-phenyldiamine to convert the Lisinopril to a more sensitive form which can be measured in Visible region its by using aqueous solvents.

The spectrophotometric method developed includes the degradation of Lisinopril to produce diketopiperazine, by the sodium hypochlorite solution in the alkaline media. Then subjected to condensation with o-Phenyldiamine to form complex showing reddish-yellow color. The diketone-o phenyldiamine complex formed is soluble in acidic and basic solvents. It has been determined by spectrophotometrically at λ_{max} 399.5 nm in alkaline media and displays linearity with Beer's range of 2 μ g to 30 μ g, the factors like temperature, reaction duration, pH of solvent are optimized. The method has been validated according to ICH Guidelines [18].

Spectrofluorimetric methods of estimation by reaction modifications

Lisinopril is non-fluorescent drug which is converted into fluorescent derivative. Fluorimetric method is based upon the condensation reaction between primary amino group of Lisinopril and Fluorescein to form fluorescent derivative (LSFN) in methanol at 60°C for 5 min. The formation of fluorescent derivative can be determined by the UV (λ_{max} 227 nm), NMR, Mass and IR spectra [13]. Spectrofluorimetric methods are used to estimate lisinopril present in small quantities i.e. in nanogramms in blood or plasma; these are frequently based upon coupling reactons with fluorephores. The saga of the development of analytical methods for lisinopril has shown the various phases of technical advancements in analytical field with respect to Lisinopril [15].

Reported estimation of Lisinopril in pharmaceutical tablets using sequential injection analysis by spectrofluorimetric method based upon the reaction between Lisinopril and o-phthalaldehyde in presence of 2-mercaptoethanol (borate buffer medium, pH=10.6), calculated at excitation wavelength of 346 nm and emission at 455 nm [19].

High performance liquid chromatographic methods

HPLC methods have been widely been used for the determination of lisinopril. We used HPLC method because these are simple, accurate, and precise and time consuming. Determination of lisinopril in dosage

forms in addition to spiked human plasma by using solid phase extraction was carried out by spectrophotometry and reversed-phase HPLC. The recommended methods were applied without any intrusion in dosage forms, either alone or co-formulated for the determination of lisinopril with hydrochlorothiazide. In addition, HPLC method was expanded to the in vitro determination of the drug in spiked human plasma [17,20,21].

Lisinopril is very useful in antihypertensive and statins as cholesterol lowering drug. By using RP-HPLC method simultaneous determination of lisinopril with pravastatin, atorvastatin, and rosuvastatin was done. Application of the proposed method was successfully applied to the determination of these compounds in pharmaceutical preparations because of their high percentage of recovery, good accuracy and precision [22].

Beasle et al. described and validate a stability-indicating HPLC method for lisinopril, lisinopril degradation product, methyl and propyl paraben in extemporaneous formulation of lisinopril [23].

RP- HPLC procedures were developed to provide a single, symmetric peak for each drug like angiotensin-converting enzyme (ACE), ramipril, enalapril maleate, benazepril lisinopril and quinapril. The HPLC method using Hypersil silica column and mobile phase consisting of methanol-water-triethylamine (50:50:0.1 v/v) and quantitation evaluation was achieved with UV detection at 210 nm [24,25,18].

HPLC method was validated and applied for the simultaneous determination of lisinopril and NSAIDs in bulk, pharmaceuticals formulations and human serum. The quantitative was achieved at 225 nm having flow rate of 1.0 ml/min. Suitability of this method for the quantitative determination of the drugs was proved by validation in accordance with International Conference on Harmonization (ICH) guidelines. The method is precise and accurate and can be used for analysis of pharmaceutical preparations in quality control (QC) and clinical laboratories [26].

The aim of this research was to study and to standardize a HPLC and absorbance ratio method for the determination of hydrochlorothiazide and lisinopril in commercially available pharmaceutical dosage forms. An HPLC method has been developed for the simultaneous determination of hydrochlorothiazide and lisinopril from formulations, using acetonitrile:water (20:80 v/v) as a mobile phase and flow rate of 1.0 ml/min. The determination of hydrochlorothiazide and lisinopril was performed by using the absorbance read at 272.0 nm, 258.8 nm and 262.7 nm in the zero-order spectra of their mixture [27].

An isocratic RP-HPLC method has been developed for the simultaneous determination of lisinopril and H2 antagonists in bulk, dosage formulations, and human serum at 225 nm. Limit of detection (LOD) and limit of quantitation (LOQ) were in the ranges of 0.07-10.4 ng/ml. Both intra- and inter-day precision and accuracy results were 98.0-102% [28].

Arayne et al. developed HPLC method for simultaneous determination of Metformin and ACE inhibitors like lisinopril, captopril, etc and its degradable product in bulk drugs, pharmaceutical products and in human serum. The analytes peaks were observed at 218 nm. The developed method was successfully applied to quantitate metformin, lisinopril, captopril, and enalapril in pharmaceutical formulations and human serum [29].

In this study, a sensitive, specific, precise and accurate method for lisinopril quantitative determination in human serum was developed

and validated. The mean recovery of lisinopril from serum samples was 88%. The LOQ for lisinopril was 6 ng/ml [30].

Combination of methods or techniques

By using HPLC (high-performance liquid chromatography) a new method was developed in which impurities in the bulk lisinopril drug was differentiated. In which mobile phase used in a ratio of 5% (v/v) acetonitrile and 95% (v/v) aqueous buffer and Detection was carried out at 210 nm later on by merge with the FT-ICRMS and NMR data, structures of unknown impurities were also found [17]. In another method isomerization of lisinopril has been studied by using HPLC, NMR, and DFT. It shows that 77% Trans and 23% cis was eluted [21].

Another sensitive and precise HPLC method with fluorimetric detection has been developed for the assay of lisinopril in human plasma and urine. The reaction mixture was chromatographed on C18-column with gradient elution, using methanol and 0.02 M phosphate buffer pH 3.2. The mean recovery of lisinopril from plasma and urine was 63.41 and 74.08%, respectively [25].

Another simple and sensitive liquid chromatography tandem multiple-stage mass spectrometry (HPLC/MS/MS) method that is suitable for bulk lisinopril analysis was developed, by which lisinopril and its RSS isomer were separated and differentiated. LC-UV scan of lisinopril revealed the presence of an unknown impurity (~0.14%) at a relative retention time RT of 3.26 m using phosphate buffer-acetonitrile as binary gradient system. The impurity was isolated by HPLC employing a linear gradient of water and acetonitrile [31,32].

The establishment of structures as well as quantification and characterizations of degraded products were carried out by sophisticated instrumental methods like UV, IR, and RP-HPLC. Furthermore, the mechanism of degradation was verified by GC-MS studies [33].

Titrimetric methods

Different titrimetric method for analysis of Lisinopril is carried out in which include following: In the middle of 18 century there was origin of titrimetric method of analysis. In 1835 Gay-Lussac invented the volumetric methods which actually lead to the foundation of the term titration. While the assay method is very old yet but certain signs of some renovation may including as dispersion to non-aqueous titrations, intensifying the field of application to weak acids and bases as well as to detection of potentiometric end point improving the accuracy of the methods. Simultaneously detection of group analysis procedures, in which titrimetric methods play a vital role in establish reaction rates. The main advantages of this method are time saving and labor with high precision and the fact that there is no need of using reference standards. Titrimetric methods have been used for the determination of lisinopril 1 in commercial dosage forms. Furthermore, it may widely uses for drug estimation; titrimetry has also been used for the estimation of degradation products of the pharmaceuticals [34-37].

Different simple titrimetric procedures are described for the determination of lisinopril (LNP) in bulk and in pharmaceutical that depend on the neutralization of basic-amino and acidic carboxylic acid groups present in LNP. In titrimetric studies have been applied in different range of concentration as 2.0-15 mg of lisinopril (LNP) this established method for determination LNP in tablets and as well as the results were validated statistically by comparing the results with those of the reference method by applying the Student's t-test and F-test. The accuracy precision of the methods were evaluated further determine by recovery studies via standard addition technique [9,38].

Another advance method in which using copper (II) phosphate for the determination of lisinopril by anodic stripping voltammetric assay. In this method of analysis different experimental parameters have been carefully premeditated and fully validated. The method has been helpful effectively for the determination of the drugs in their pharmaceutical preparations as well as in plasma. The acquired results were compared statistically with those obtained from the official USP methods as well as in case of AS published method [39].

Miscellaneous methods

Lisinopril have been finding out in different dosage form and also in biological fluids by highly sensitive and simple polarographic method. In this method, dealing of the compound with nitrous acid as a result measuring the cathodic current produced by the resulting nitroso derivative. The polarographic behavior was calculated as adopting direct current (DCt), differential pulse (DP) and alternating current (ACt) polarography along with well define pH range of 1.0-8.0 was achieved in Britton-Robinson buffers (BRb). This method may also be performed for determination of lisinopril in spiked human urine and plasma [18].

Another approach has been developed for analysis of lisinopril (LIS) in pharmaceutical formulations using a tris(2,2'-bipyridyl)-ruthenium (II) (Ru (bipy) 32+) peroxydisulfate chemiluminescence (CL) system in a two chip device. This is an ideal, rapid, selective and sensitive method including parameters like flow rates, pH, and concentration of reagents were carefully optimized under influence the CL signal intensity. This method is consider as one the best method for analysis of LIS in pharmaceutical products and was establish to be free from any hindrance of acid-induced degradation (AID) products and other ingredients frequently present in these preparations [40].

By using Spectrophotometric and polarographic technique determination of lisinopril has been established using 2, 4-dinitrofluorobenzene by formation of colored products and polarographically active derivatives. In this study methods have been validated and optimized by different experimental conditions which may be pertain to the determination of lisinopril in their commercial scale tablets. Comparative statistical analysis has been recognized by means of the authorized HPLC methods [41].

A novel method for the quantitative determination of the angiotensin-converting enzyme inhibitor lisinopril in human plasma have been established by mean of negative ion chemical ionization mass spectrometry and gas chromatography [42].

Other method with respect to chemical nature the drug may also act as photosensitive behavior as go through hydrolysis and oxidized in presence of oxygen. Due to respect of stress degradation of lisinopril certain studies have been conducted under different conditions recommended by International Conference on Harmonization (ICH) in which objective of study as to find out the pathway for stress degradation of Lisinopril in bulk [43].

Conclusion

The analytical methods always accentuate for the simplicity, accuracy, precision, and specificity. Analytical method development mainly involves developing method having advantages over existing methods. High pressure liquid chromatography HPLC methods required costly equipment, labor intensive sample preparation procedure and the personal skilled in chromatographic techniques most of the HPLC methods reviewed have the prospective application

to clinical research of multi-drug pharmacokinetics, drug combination studies and also for interaction studies.

The analytical methods developed for lisinopril show that the former analytical methods developed for lisinopril evaluation in single or in combination formulations reveal the development of spectrophotometric and fluorimetric methods based coupling reactions followed by chromatographic methods. The other methods such as capillary electrophoresis, fluoroimmunoassay, radio immunoassay and fluoro enzymatic assay have also been reported. The development of each method indicates the overcoming of the limitations of the existing methods and making availability of better methods.

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