



Spectrophotometric estimation of cefixime and ofloxacin from tablet dosage form

Avanija Dube*, Sujit Pillai, Sumit Sahu and Naina Keskar
GRY Institute of Pharmacy, Borawan, Khargone, (M.P.) -India

Abstract

Two simple, rapid, accurate and precise spectrophotometric methods have been developed for simultaneous estimation of Cefixime and Ofloxacin from tablet dosage form. Method I involves formation of 'simultaneous equations' at 234 nm (λ max of Cefixime) and 296 nm (λ max of Ofloxacin); while Method II involves, formation of 'Absorbance ratio equation' at 275(ISOABSORPTIVE POINT) and 296 nm (λ max of Ofloxacin) using methanol as a solvent. The linearity was observed in the concentration range of 4 - 20 μ g/ml for Cefixime and 2 - 10 μ g/ml for Ofloxacin. The results of analysis have been validated statistically and by recovery studies and were found satisfactory.

Key-Words: Cefixime(CEF), Ofloxacin(OFL), Simultaneous equation method, Absorbance ratio method

Introduction

Cefixime (CEF) is an oral third generation cephalosporin antibiotic. Chemically, it is (6R,7R)-7-[[2-(2-amino-1,3-thiazol-4-yl)-2-(carboxymethoxyimino) acetyl]amino}-3-ethenyl 8-oxo-5-thia-1-azabicyclo-[4.2.0]oct-2-ene-2- carboxylic acid, clinically used in the treatment of susceptible infections including gonorrhoea, otitis media, pharyngitis, lower respiratory-tract infections such as bronchitis, and urinary-tract infections. Ofloxacin (OFL) is a fluoroquinolone derivative. Chemically, it is (\pm)-9-fluoro-2, 3-dihydro-3-methyl-10- (4-methyl-1 piperazinyl)-7-oxo-7H-pyrido-[1,2,3-de]- 1,4-benzoxazine -6-carboxylic acid¹. Literature review reveals that Cefixime can be estimated spectrophotometrically² and Ofloxacin can be estimated spectrophotometrically³ in combination with other drugs only one method is reported for estimation of Cefixime and Ofloxacin in combination by RP-HPLC⁴. So here an attempt has been made to develop simple, accurate, sensitive, rapid and economic method for simultaneous estimation of Cefixime and Ofloxacin from tablet dosage forms using UV - Visible spectroscopy.

Material and Methods

Reagents and Chemicals

AR- grade methanol was used as solvent which was purchased from Loba Chem Pvt Ltd, Mumbai. Analytical pure standard sample of CEF and OFL were supplied as gift sample by Vapicare Pharma pvt ltd, Vapi, Gujarat and used without further purification. The pharmaceutical dosage form used in study was CEFIO 200 (label claim CEF 200 mg, OFL 200 mg) manufactured in India by Accent Pharma. India.

Instrumentation

The instrument used for the entire analysis was SIMADZU UV 1700 UV-VIS recording spectrophotometer. It is a double beam high speed scanning spectrophotometer with advanced quantitative software and provides full facilities for monochromators, a CRT display and a parallel head printer.

Preparation of standard stock solution

100 mg of Cefixime (CEF) and 100 mg of Ofloxacin (OFL) were weighed separately and transferred to two separate 100 ml volumetric flasks. Each drug was dissolved in 40 ml of methanol and shaken gently for 10 min. The volume was made up to the mark with same solvent and the final strength obtained was 1000 μ g/ml.

Methods

Method I: Simultaneous Equation Method⁵

In simultaneous equation method, when no region can be found free from overlapping spectra of two

* Corresponding Author:

E-mail: avanija12pharma@gmail.com

Mob. +919993987422

chromophores, it is still possible to devise a method based on measurements at two wavelengths. Two dissimilar chromophores must necessary have different powers of light absorption at some point or in linear absorption spectra.

If samples contain two absorbing drugs (X and Y), each of which absorbs at the λ_{max} of the other, it may be possible to determine both drugs by the technique of simultaneous equations.

From overlain spectra (Fig 1.) 234 nm λ_{max} for Cefixime and 296 nm λ_{max} for Ofloxacin were selected for formation of simultaneous equation of two drugs. The absorbance at 234 nm and 296 nm for CEF and OFL were measured. The absorptivity values of each drug at both wavelengths were determined. The absorbance and absorptivity at this wavelength were substituted in following equations to obtain the concentration of both drugs.

$$C_x = \frac{A_2 a_{y1} - A_1 a_{y2}}{a_{x2} a_{y1} - a_{x1} a_{y2}} \quad \text{I}$$

$$C_y = \frac{A_1 a_{x2} - A_2 a_{x1}}{a_{x2} a_{y1} - a_{x1} a_{y2}} \quad \text{II}$$

Where, A_1 and A_2 were absorbance of sample at 234 nm and 296 nm respectively, a_{x1} and a_{x2} are absorptivity of Cefixime at 234 nm and 296 nm, a_{y1} and a_{y2} are absorptivity of Ofloxacin at 234 nm and 296 nm.

Validity of above framed equation was checked by using mixed standard of pure drug sample of two drugs, measuring their absorbance at respective wavelength and calculating concentration of two components. Results of which are reported in Table.1.

Method II: Absorbance Ratio Method⁵

The absorbance ratio method is a modification of the simultaneous equation procedure. It depends on the property that, for a substance which obeys Beer's Law at all wavelengths is a constant value independent of concentration or Path length. In the quantitative assay of two components in admixture by the absorbance by the absorbance ratio method, absorbance's are measured at two wavelengths one being the λ_{max} of one of the components (λ_2) and the other being a wavelength of equal absorptivity of the two components (λ_1), that is, an isoabsorptive point.

From overlain spectra (Fig 1.) 275 nm (Isobestic point) and 296 nm λ_{max} for Ofloxacin were selected for formation of Absorbance ratio equation of two drugs.

The absorbance at 275 nm and 296 nm for CEF and OFL were measured. The absorptivity values of each drug at both wavelengths were determined. The absorbance and absorptivity at this wavelength were substituted in following equations to obtain the concentration of both drugs.

$$C_x = \frac{Q_M - Q_Y}{Q_X - Q_Y} * \frac{A_1}{a_{x1}} \quad \text{III}$$

$$C_x = \frac{Q_M - Q_x}{Q_y - Q_x} * \frac{A_1}{a_{y1}} \quad \text{VI}$$

Q_M , Q_X , and Q_Y were obtained as below:
 $Q_M = A_2/A_1$, $Q_X = a_{x2}/a_{x1}$, $Q_Y = a_{y2}/a_{y1}$

Where, A_1 and A_2 were absorbance of sample at 275 nm and 296 nm respectively, a_{x1} and a_{x2} are absorptivity of Cefixime at 275 nm and 296 nm, a_{y1} and a_{y2} are absorptivity of Ofloxacin at 275 nm and 296 nm.

Validity of above framed equation was checked by using mixed standard of pure drug sample of two drugs, measuring their absorbance at respective wavelength and calculating concentration of two components. Results of which are reported in Table.1

Assay of tablet formulation

Twenty tablets were weighed and crushed to obtain a fine powder. An accurately weighed sample equivalent to 100 mg of CEF and 100 mg of OFL was taken in a stoppered volumetric flask (100.0ml); 40ml of Methanol was added and sonicated for 10 min. The solution was filtered through Whatmann filter paper (No 41) and the volume was made up to the mark with the same solvent. The aliquot portions of above solutions were further diluted with solvent to get final concentration of about 6 $\mu\text{g/ml}$ CEF and 6 $\mu\text{g/ml}$ of OFL, respectively and absorbances were measured at 234.0 nm, 275 nm and 296.0 nm against blank. The concentrations of two drugs in sample were determined by using equations 1 and 2 and equation 3 and 4. The results are reported in the Table 2.

Recovery studies

The accuracy of the proposed method was checked by recovery studies, by addition of standard drug solution to pre analyzed sample solution at three different concentration levels (80%, 100%, and 120%) within range of linearity for both the drugs. Results are reported in table 3

Results and Conclusion

Table 1: Result of analysis of mixed standard for method I and method II

Conc Present (µg/ml)		Conc Found (%)			
		Method I		Method II	
CEF	OFL	CEF	OFL	CEF	OFL
4	8	101	99.67	97.25	100.25
6	6	102.98	103.88	100.66	104.66

Method I: Simultaneous Equation Method

UV-spectrophotometric method using simultaneous equation was developed. CEF showed absorbance maxima at 234 nm and OFL at 296.0 nm. Linearity was observed in the concentration range of 4 - 20 µg/ml for CEF and 2 - 10 µg/ml for OFL correlation coefficient was found to be 0.9996 and 0.9998 at 234 nm and 296 nm respectively. The proposed method was applied for pharmaceutical formulation and % label claim for CEF and OFL was found to be 97.00 and 102.20, respectively. The method is accurate and precise and can be used for routine pharmaceutical analysis.

Method II: Absorbance Ratio Method

UV-spectrophotometric method by using absorbance ratio method was developed. Absorbances selected were 275 nm (isoabsorptive point) and 296 nm (λ max of Ofloxacin) Linearity was observed in the concentration range of 4 - 20 µg/ml and 2 - 10 µg/ml correlation coefficient was found to be 0.9995 and 0.9998 respectively The proposed method was applied for pharmaceutical formulation; % label claim for CEF and OFL was found to be 95.16 and 102.83, respectively. The low % RSD indicates method is accurate and precise.

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Table 3: Result of recovery studies of CEF and OFL

Drug	% Recovery (Mean \pm S.D)	
	Method I	Method II
CEF	97.22 \pm 0.0016	100.75 \pm 0.16
OFL	102.22 \pm 0.08	104.66 \pm 0.16

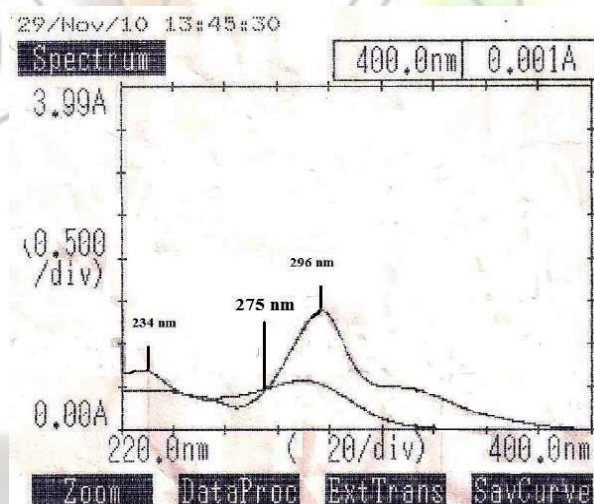


Fig 1: Overlain spectra of CEF and OFL

Table 2: Result of analysis of tablet formulation

Label Claimed mg/ml		% Label Estimated(Mean ± S.D)				% R.S.D			
		Method I		Method II		Method I		Method II	
CEF	OFL	CEF	OFL	CEF	OFL	CEF	OFL	CEF	OFL
200	200	97.00 (±.0016)	102.20 (±.0024)	95.16 (±.0006)	102.83 (±.0024)	0.31	0.26	0.17	0.26