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Spectrophotometric determination of chlorpheniramine maleate and phenylpropanolamine hydrochloride in dosage forms

*Arun Kaura¹, Vikas Gupta¹, G S Roy², Monika Kaura²

¹University Institute of Pharmacy, Baba Farid University of Health Sciences, Faridkot, Punjab, India

²B.I.S. College of Pharmacy, Gagra, Moga, Punjab, India

ABSTRACT

A rapid and simple method for simultaneous determination of Chlorpheniramine Maleate (CPM) and Phenylpropanolamine Hydrochloride (PPM) by first derivative UV spectrophotometry has been developed in combined pharmaceutical dosage forms. The proposed method was successfully applied for the determination of drugs in physical mixture and commercial formulations and results showed good linearity, precision and reproducibility.

Key Words: Derivative, UV absorption, spectral overlap, principle maxima, wavelength range, analytical signal.

INTRODUCTION

Chlorpheniramine maleate (CPM) inhibits the effects of histamine on capillary permeability and bronchial smooth muscles. It is an anti-allergic drug, widely used in cough-cold preparations. Phenylpropanolamine hydrochloride (PPM) is indirectly acting sympathomimetic agent and it is used in the symptomatic relief of nasal congestion. These drugs are used either alone or in combination. The simultaneous determination of these drugs is not possible by direct UV absorption measurement method because of spectral overlap of their principle maxima. The present work consists of taking a derivative of convenient order of the analytical signal i.e. absorbance in the wavelength domain. Besides the official methods (IP & USP) the other analytical methods available in literature for determination of Chlorpheniramine Maleate (Alaa El-Gindy *et al.*, 2005; Cieri *et al.*, 2006; Fried *et al.*, 2002; Liao *et al.*, 2008; Marin *et al.*, 2002; Rouhollah Heydari *et al.*, 2008; Senyuva *et al.*, 2002), Phenylpropanolamine Hydrochloride (Abbasi *et al.*, 2006; Azhagvuel *et al.*, 2007; Ferreyra *et al.*, 2002; Issa *et al.*, 2005; Kaddoumi *et al.*, 2004; Nakashima *et al.*, 2002) and combination of

Chlorpheniramine Maleate & Phenylpropanolamine Hydrochloride (Fabrizio *et al.*, 1980) have been mentioned. These methods are time consuming; therefore an alternative method for selective derivative UV spectrophotometry is rendered.

MATERIALS AND METHODS

Materials

Authentic specimens of CPM and PPM were provided as a gift samples from M/S Plethico Pharmaceuticals, Indore. All other reagents used were of analytical grade.

Analysis of commercial tablets & recovery studies

Spectrophotometric analysis was carried out on a Shimadzu UV 160-A spectrophotometer in 1cm matched quartz cells. The amplitudes were obtained directly from the display of the first derivative in the wavelength range 200-325 nm with the derivation interval 5nm at a medium scan speed for determination of CPM and PPM. The standard calibration curves of both the drugs were prepared to ascertain that the drugs solutions obey the Beer's Law in the concentration range to be utilized for the preparation of mixed standards, it was found that both the drugs followed Beer's Law in the working range of concentrations i.e. 0-24 µg/ml for CPM and 0-450 µg/ml for PPM. The solutions of both the drugs were prepared in distilled water.

*Corresponding Author:

Arun Kumar Kaura
University Institute of Pharmacy,
Baba Farid University of Health Sciences,
Sadiq Road, Faridkot-151203 (Pb.) India
E-mail: arunkaura70@rediffmail.com
Contact No.: +919888010178

Table 1: Statistical evaluation of linearity of standard solutions of PPM and CPM.

Analyte	Slope	Intercept	Coefficient of Correlation
PPM	0.0010394	-0.3131	0.999
CPM	0.0141781	0.0271	0.999

PPM: indicates Phenylpropranolamine hydrochloride
 CPM: indicates Chlorpheniramine maleate

The correlation among slope, intercept and coefficient of correlation were estimated for both the drugs using standard calibration curves for PPM and CPM. The results of statistical data are given in table 1. An accurately weighed portion (243.26 mg) of the powder (mixed contents of 20 tablets) equivalent to about 4 mg of CPM and 25 mg of PPM were transferred to a 100 ml calibrated flask. 50mg of PPM standard drug was weighed and transferred to the same volumetric flask. 60ml of distilled water was added and drugs were dissolved by shaking vigorously for 10 minutes. The resultant solution was filtered in 100 ml calibrated flask and volume

was made upto mark with distilled water. The final solution was labeled to claim 40 µg/ml of CPM and 750 µg/ml of PPM. From this stock solution different dilutions were made such that it falls in the working range of concentrations i.e. 0-24 µg/ml for CPM and 0-450 µg/ml for PPM. These diluted solutions were directly subjected to spectrophotometric analysis.

To study the recovery of CPM and PPM, pre-analysed tablet sample solutions were taken and to these solutions, different concentrations of standard drugs (CPM and PPM) were added. The resultant solutions were analysed by proposed first derivative UV spectrophotometric method. The results (table 2) were found to be accurate, precise and reproducible.

RESULTS AND DISCUSSION

The normal or zero order overlain absorption UV spectra of CPM (40 µg/ml) and PPM (250 µg/ml) in the 200-325 nm wavelength region (figure 1), PPM exhibited three absorption peaks and it showed

Table 2: (a) Statistical estimation of results of CPM and PPM in authentic, commercial and recovery studies samples (b) Variations in performed statistical analysis.

Type of sample	Expected Concentration (µg/ml)		Found Concentration (µg/ml)		% found		Mean % Found	
	CPM	PPM	CPM	PPM	CPM	PPM	CPM	PPM
Authentic samples	4	50	4.0829	50.0885	102.07	100.17	99.958	99.374
	6	100	5.9065	98.2961	98.44	98.29		
	10	200	9.9183	200.0677	99.18	100.03		
	15	300	15.764	296.4829	100.50	98.82		
	20	400	19.9219	398.2545	99.60	99.56		
Commercial samples	4	75	4.0308	76.8705	100.77	102.49	100.69	100.126
	8	150	8.0427	146.5037	100.53	97.66		
	12	225	12.2108	226.8497	101.75	100.82		
	16	300	16.1706	301.8393	101.06	100.61		
	20	375	19.8698	371.4725	99.34	99.05		
Recovery studies samples	10	150	10.1105	149.9799	101.10	99.98	99.6075	100.07
	4	125	3.9606	123.1972	99.01	98.55		
	6	100	5.9402	101.77	99.0	101.77		
	3	75	2.9797	74.9896	99.32	99.98		

(a)

Type of sample	Standard Deviation		Standard Error		Coefficient of Variation	
	CPM	PPM	CPM	PPM	CPM	PPM
Authentic Samples	1.2484	0.7184	0.5583	0.3213	1.2489	0.7229
Commercial Samples	0.7893	1.6459	0.3529	0.7360	0.7839	1.6437
Recovery Studies Samples	0.8713	1.1419	0.4356	0.5709	0.8747	1.1412

(b)

PPM: indicates Phenylpropranolamine hydrochloride
 CPM: indicates Chlorpheniramine maleate

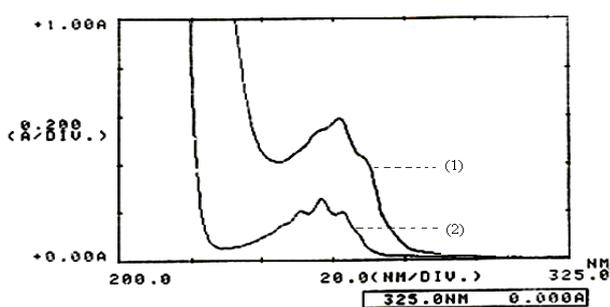


Figure 1: Normal and zero order overlain spectrum of CPM (1) and PPM (2).

maximum absorbance at 257 nm, CPM also absorbed near this wavelength region with a peak at 261.6 nm. The extensive overlap of spectral bands of these two drugs refrained conventional UV spectrophotometry for their individual determination in a mixture.

The first derivative spectra recorded the sharp bands of large amplitude of PPM (250 $\mu\text{g/ml}$) and CPM (40 $\mu\text{g/ml}$) (figure 2), which permitted very clear identification and determination of these drugs. The choice of the optimum wavelength is based on the fact that the contribution of one component to the overall derivative signal is zero at the wavelength at which the other component exhibited maximum absorption. From the first derivative overlain spectra of CPM and PPM (figure 2), it was found that at 231.4 nm, PPM showed zero contribution. Therefore, this point was selected as zero crossing for PPM and at this point CPM showed remarkable absorbance. So, 231.4 nm was selected for CPM absorbance measurement and likewise 259.2 nm for PPM absorbance measurement for the simultaneous determination of CPM and PPM in a mixture form.

CONCLUSION

The derivative UV spectrophotometry appears to be a suitable technique for the reliable analysis of commercial formulations containing combination of CPM and PPM. The most striking features of the derivative method are its simplicity, sensitivity and rapidity. It is also an easier and economical method than HPLC separation technique and does not require the use of any expensive or toxic reagent. These advantages make it especially suitable for routine quality control.

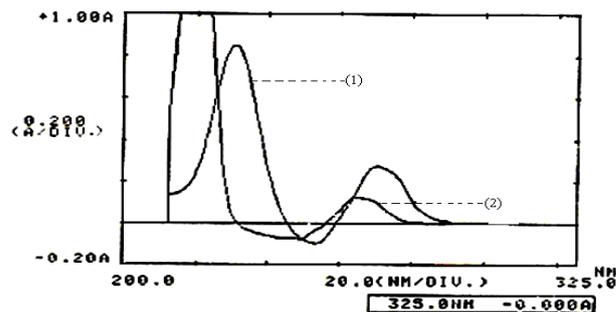


Figure 2: First derivative overlain spectrum of CPM (1) and PPM (2).

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