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# Acoustic and volumetric properties of ciprofloxacin hydrochloride in dioxane-water mixture at 303.3 K

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## Abstract

The acoustical properties have been investigated from the ultrasonic velocity and density measurements of substituted heterocyclic drug (ciprofloxacin hydrochloride) in 1, 4-dioxane-water at 303.3K. The measurements have been performed to evaluate acoustical parameters such as adiabatic compressibility ( $\beta_s$ ), Partial molar volume ( $\phi_v$ ), intermolecular free length ( $L_f$ ), apparent molar compressibility ( $\phi_K$ ), specific acoustic impedance (Z), relative association ( $R_A$ ), and solvation number (Sn).

Key word: Ultrasonic velocity, ciprofloxacin hydrochloride, apparent molar volume

## Introduction

In the recent years, measurements of the Ultrasonic velocity are helpful to interpret solute-solvent and ion-solvent interaction in aqueous and non aqueous medium [1-2]. Ultrasonic is a versatile nondestructive technique and highly useful for the investigation of various physical properties [5-7]. Recent developments have found use of ultrasonic energy in medicine, engineering and agriculture. Drug-macromolecular interactions are an important phenomenon in physiological media, such as blood, membranes. intraand extracellular fluids. Physicochemical investigations play an important role in understanding the nature and the extent of the patterns of molecular aggregation that exist in binary liquid mixtures and their sensitivities to variations in composition and the molecular structure of the pure components. The drug-solvent molecular interaction play an important role in the understanding of drug action [7]. Viscometric properties provide valuable clues for solute-solvent interactions in the solution phase [8].

Ciprofloxacin hydrochloride (Sip-roh-floxass-in hi-droh-clor-ride) is a medicine which is used in bacterial infections. Ciprofloxacin hydrochloride (HCl) is a fluoroquinolone antibiotic. It is effective against a wide range of gram positive and gram negative bacteria, and is most well known for its effectiveness against mycoplasma. Ciprofloxacin HCl works by interfering with the bacterial enzyme DNA gyrase, an enzyme necessary for bacterial synthesis, replication, and transcription in both the active and non-active growth phases of the bacterial life cycle [9].The effective concentration for Ciprofloxacin HCl, as with many antibiotics varies with the organism or the cell type, environmental conditions, and stage of growth cycle. However, a concentration of 10  $\mu$ g/mL is active against most strains of gram

negative and gram positive bacteria, as well as several species of mycoplasma. In comparison to other antibiotics, Ciprofloxacin HCl has many advantages, including lack of known resistant strains, minimal side effects, a lower effective working concentration, non-cytotoxicity, a low reoccurrence rate for contamination, and the ability to kill bacteria in both the active and non-active growth phases [9,10].

In this, the present research Acoustical and volumetric properties have been measured for Ciprofloxacin hydrochloride drug in dioxane-water mixture at 303.3K and different parameters such as adiabatic compressibility ( $\beta_s$ ), Partial molar volume ( $\phi_v$ ), intermolecular free length ( $L_f$ ), apparent molar compressibility ( $\phi_K$ ), specific acoustic impedance (Z), relative association ( $R_A$ ), solvation number (Sn).limiting apparent molar compressibility ( $\phi^0\kappa$ ), limiting apparent molar volume( $\phi^0v$ ) and their constant (Sk, Sv). Viscosity coefficient (A, B)were studied [12-15].

## **Material and Method**

Ciprofloxacin hydrochloride (HCl) and dioxane were purchased from Sigma Aldrich. The double distilled dioxane was used for preparation of different concentration of drug solution.

## Preparation of Ciprofloxacin hydrochloride (HCl) in water-Dioxane

The drug Ciprofloxacin hydrochloride (HCl) was used. Dioxane was purified by Vogel's standard method. The double distilled dioxane was used for preparation of different concentration of drugs solution. The densities were determined by using specific gravity bottle by relative measurement method with accuracy  $\pm 1x10-5$  gm/cm<sup>3</sup>.

#### Theory

Sound-velocity is given by:  $U = (\phi \kappa.d)^{-1/2}$ Apparent molar compressibility ( $\phi \kappa$ ) has been calculated by using the relation,

 $\varphi \kappa = 1000\beta_{s}C^{-1} - \beta_{s} d_{0}^{-1}(1000C^{-1}dM)$ 

Where,  $\beta_{s_i} d$  and  $d_0$  are the adiabatic compressibility and density of solution and solvent respectively. C is molar concentration of solute; M is molecular weight of solute.

Apparent molar volume  $(v_{\phi}) = 1000(Cd_0)^{-1} (d_0 - d) + M_2 d_0^{-1}$ 

Specific acoustic impedance (Z) = Us ds Intermolecular free length ( $L_f$ ) = K $\sqrt{\beta_s}$ Relative association ( $R_A$ ) = (ds / d0) x (U0 /Us)<sup>1/3</sup> The entire viscosity data have been analyzed by using

Jones-Dole equation  $\eta_r - 1 / \sqrt{c} = A + B \sqrt{c}$ 

#### **Result and Discussion**

In the present investigation, different acoustical properties such as Relative viscosity( $\eta_r$ ), Density(ds), Apparent molar volume(vo), Solvation number(Sn) are listed in table-1. limiting apparent molar Limiting apparent volume( $\varphi_0 v$ ), molar compressibility ( $\varphi_0 \kappa$ ), their constant (Sk, Sv) and viscosity coefficient (A, B) are listed in table-2. Ultrasonic velocity (Us), apparent molar compressibility ( $\varphi \kappa$ ), relative association ( $R_A$ ), adiabatic compressibility(Ks), Intermolecular free length (L<sub>f</sub>),specific acoustic impedance (Z) are listed in table-3. It was found that the ultrasonic velocity decreased with the increase in concentration for dioxane-water system. Variation of ultrasonic velocity in solution depends upon the increase or decrease of molecular free length after mixing the component. Intermolecular free length increased linearly on increase in concentration of Ciprofloxacin hydrochloride in 1,4dioxane. .It is happened because there was significant interaction between ions and solvent molecules suggesting a structure promoting

behavior of the added electrolyte. The specific acoustic impedance (Z) decreased with the increase in concentration. When concentration of electrolyte was increased, the thickness of oppositely charged ionic atmosphere increases due to decrease in ionic strength. This is suggested by decrease in acoustic impedance with increase in concentration. It was seen that the intermolecular free length increased very little with the increase in concentration in dioxanewater system. This intermolecular free length variation is due to greater force of attraction between solute and solvent by forming hydrogen bonding. The adiabatic compressibility increased with the increase in concentration of solution. It is happened due to collection of solvent molecule around ions, this supporting weak ion-solvent interaction. This indicates that there is significant solute-solvent interaction. It was observed that apparent molar volume increased with concentration. It indicates the existence of strong ion-solvent interaction. It was found that the value of adiabatic compressibility was increased with the increase in concentration of Ciprofloxacin hydrochloride in dioxane-water system. It shows strong electrostatic attractive force in the vicinity of ions. From the data, we were concluded that strong molecular association was found in dioxane-water system. The value of relative association increased with the increase in concentration the solvation number in Ciprofloxacin hydrochloride -dioxane system is increased with concentration, it indicates the solvent molecule forms strong coordination bond in primary layer. The limiting molar compressibility was positive. The value of  $S_k$  exhibits positive. It indicates the existence of ion-ion or solute-solute interactions in system. From table-2, it was found that the value of limiting apparent molar volume was positive in systems. It indicates that the ion-dipolar interaction in Ciprofloxacin hydrochloride in 1,4-dioxane-water mixture. The positive value of  $S_v$  indicates the strong solute-solvent interaction. These value indicates an induced effect of 1, 4- dioxane on solute-solvent

determined in table-2. From Table-2 it was observed that the value of 'A' (Falkenhagen coefficient) was positive. 'A' measures of ionic interaction. It indicates that there is a strong solute-solute interaction in solute molecules. The Jones–Dole coefficient measures solute –solvent interaction. The value of "B"coefficient was negative in which measures the solute-solvent interactions. But with the increase in concentration of dioxanewater,the B value becomes less negative indicating stronger ion-solvent interaction. The value of A and B are determined.figure.1 and Table 2

interaction. The value of  $S_k$  and  $S_v$  have been

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Concentration (mole.lit <sup>-1)</sup>	Relative viscosity (ηr)	Density (ds) kg m <sup>-3</sup>	Apparent molarvolume (vφ) m <sup>3</sup> mole <sup>-1</sup>	Solvation number (S <sub>n</sub> )		
10% dioxane-w	ater mixture at 303.3	K				
0.00	1.512	1021.11	0.5072	0.32743		
$1 \times 10^{-3}$	1.444	1021.23	0.5166	0.32975		
$2x10^{-3}$	1.382	1021.32	0.5264	0.33256		
$3x10^{-3}$	1.303	1021.46	0.5356	0.33451		
$4x10^{-3}$	1.232	1021.58	0.5392	0.33696		
5x10 <sup>-3</sup>	1.221	1021.62	0.5458	0.33936		
6x10 <sup>-3</sup>	1.206	1021.67	0.5506	0.34239		
7x10 <sup>-3</sup>	1.189	1021.70	0.5544	0.34461		
8x10 <sup>-3</sup>	1.172	1021.77	0.5586	0.34714		
9x10 <sup>-3</sup>	1.159	1021.81 0.5618		0.34925		
20% dioxane-water mixture at 303.3 K						
0.00	1.521	1021.32	0.5297	0.32806		
$1 \times 10^{-3}$	1.461	1021.41	0.5335	0.33017		
$2x10^{-3}$	1.404	1021.49	0.5366	0.33206		
$3x10^{-3}$	1.345	1021.54	0.5405	0.33402		
$4x10^{-3}$	1.295	1021.59	0.5441	0.33697		
$5 \times 10^{-3}$	1.232	1021.66	0.5480	0.33986		
6x10 <sup>-3</sup>	1.212	1021.69	0.5512	0.34249		
$7x10^{-3}$	1.195	1021.73	0.5548	0.34499		
8x10 <sup>-3</sup>	1.181	1021.76	0.5589	0.34736		
9x10 <sup>-3</sup>	1.166	1021.82	0.5677	0.35034		
30% dioxane-w	ater mixture at 303.3	K				
0.00	1.531	1021.47	0.5302	0.32840		
$1 \times 10^{-3}$	1.468	1021.54	0.5339	0.34042		
$2x10^{-3}$	1.412	1021.59	0.5373	0.34183		
$3x10^{-3}$	1.355	1021.62	0.5409	0.34314		
$4x10^{-3}$	1.304	1021.68	0.5449	0.34520		
5x10 <sup>-3</sup>	1.243	1021.72	0.5483	0.34713		
6x10 <sup>-3</sup>	1.218	1021.75	0.5514	0.34985		
$7x10^{-3}$	1.207	1021.79	0.5552	0.35120		
8x10 <sup>-3</sup>	1.189	1021.81	0.5589	0.35366		
9x10 <sup>-3</sup>	1.173	1021.89	0.5087	0.35517		

## Table-2: Physical properties of ciprofloxacin hydrochloride in dioxane-water mixture at 303.3 K

В.					
Parameter	10%	20%	30%		
Limiting Apparent molar volume ( $\phi_v$ )m <sup>3</sup> mole	0.5053	0.5056	0.5058		
Limiting Apparent molar compressibility ( $\phi0\kappa)x\;10^{-9}\;m^2\;N^{-1}$	1.0425	1.0427	1.0428		
$S_v m^3 kg^{1/2} mole^{-3/2}$	6.322	6.327	6.329		
$S_k M^3$ mole-2kg. N <sup>-1</sup>	51.885	51.886	51.889		
A B	10.501 -55.453	10.504 -55.451	10.506 -55.449		

 $Table - 2: Limiting \ Apparent \ molar \ compressibility \ (\phi\kappa), \ Limiting \ Apparent \ molar \ volume \ (v\phi), \ S_{v}, S_{k}, A \ and B \ and and B \ and$ 

## Table-3:

Ultrasonic velocity (Us) m s<sup>-1</sup>, Apparent molar compressibility ( $\phi k x 10^{-9}$ ) m<sup>2</sup>N<sup>-1</sup>, Relative association (R<sub>A</sub>), Adiabatic compressibility (K<sub>S</sub>)  $x 10^{-10}$  m<sup>2</sup>N<sup>-1</sup>, Intermolecular free length (L<sub>f</sub>)  $x 10^{-11}$  m, Specific acoustic impedance (Zx10<sup>6</sup>) kg m<sup>-2</sup> s<sup>-1</sup>

Concentration	Ultrasonic	Apparent	Relative	Adiabatic	Intermolec	Specific
	velocity (Us)	molarcompress	associati	compressibiliy	ular	acoustic
	<b>m</b> s <sup>-1</sup>	ibiity(øk x10 <sup>-9</sup> )	on	$(\beta_{s}) x 10^{-10}$	free length	impedance
		$m^2 N^{-1}$	( <b>R</b> <sub>A</sub> )	$m^2 N^{-1}$	$(L_f) x 10^{-11}$	$(Zx10^{6})$
					m	kg m <sup>-2</sup> s <sup>-1</sup>
10% dioxane-water mixture at 303.3 K						
0.00	1322.21	3.5603	1.00164	5.60432	4.76121	1.3503
$1 \times 10^{-3}$	1318.24	3.5852	1.00359	5.63804	4.77563	1.3462
$2x10^{-3}$	1313.62	3.6165	1.00582	5.67706	4.79206	1.3424
$3x10^{-3}$	1307.85	3.6372	1.00735	5.72666	4.81293	1.3367
$4x10^{-3}$	1303.93	3.6643	1.00933	5.76068	4.82715	1.3328
$5 \times 10^{-3}$	1299.19	3.6895	1.01195	5.80314	4.84492	1.3274
6x10 <sup>-3</sup>	1294.54	3.7231	1.01359	5.84395	4.86196	1.3227
$7x10^{-3}$	1286.67	3.7472	1.01527	5.89693	4.88937	1.3168
8x10 <sup>-3</sup>	1287.43	3.7751	1.01716	5.93554	4.89993	1.3126
$9x10^{-3}$	1277.91	3.5603	1.00164	5.97965	4.91804	1.3078
20% dioxane-w	ater mixture at	303.3 K				
0.00	1322.28	3.564	1.00168	5.60439	4.76125	1.3504
$1 \times 10^{-3}$	1318.25	3.5853	1.00357	5.63806	4.77564	1.3463
$2x10^{-3}$	1313.64	3.6167	1.00583	5.67704	4.79207	1.3425
$3x10^{-3}$	1307.88	3.6374	1.00734	5.72662	4.81294	1.3368
$4x10^{-3}$	1303.81	3.6647	1.00938	5.76064	4.82716	1.3329
5x10 <sup>-3</sup>	1299.13	3.6894	1.01192	5.80313	4.84494	1.3275

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6x10 <sup>-3</sup>	1294.45	3.7237	1.01354	5.84399	4.86195	1.3228	
7x10 <sup>-3</sup>	1286.61	3.7479	1.01528	5.89696	4.88936	1.3169	
8x10 <sup>-3</sup>	1287.42	3.7753	1.01712	5.93551	4.89994	1.3127	
9x10 <sup>-3</sup>	1277.84	3.5602	1.00163	5.97963	4.91805	1.3079	
30% dioxane-v	vater mixture at 3	303.3 K					
0.00	1322.30	3.5606	1.00174	5.60441	4.76125	1.3504	
1x10 <sup>-3</sup>	1318.29	3.5855	1.00364	5.63807	4.77565	1.3463	
$2x10^{-3}$	1313.67	3.6167	1.00589	5.67707	4.79207	1.3425	
3x10 <sup>-3</sup>	1307.88	3.6378	1.00736	5.72668	4.81295	1.3369	
$4x10^{-3}$	1303.97	3.6644	1.00939	5.76069	4.82717	1.3329	
5x10 <sup>-3</sup>	1299.21	3.6896	1.01198	5.80315	4.84495	1.3278	
6x10 <sup>-3</sup>	1294.55	3.7235	1.01360	5.84397	4.86197	1.3229	
7x10 <sup>-3</sup>	1286.68	3.7476	1.01527	5.89698	4.88935	1.3169	
8x10 <sup>-3</sup>	1287.44	3.7756	1.01717	5.93559	4.89997	1.3129	
9x10 <sup>-3</sup>	1277.92	3.5606	1.00165	5.97967	4.91808	1.3086	



Fig.1: Relative viscosity of drug using Jones-Dole equation

## Conclusion

In the present study mentions the experimental data for ultrasonic velocity and density at 303.3K for ciprofloxacin in 1,4-dioxane-water mixture. From experimental data calculated acoustical parameters and studied to explain solute-solvent interaction and ion-ion / solute-solute interaction are existing between drug and organic solvent mixture.

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