

Experimental Study and Correlation of the Solid-liquid Equilibrium of Some Amino Acids in Binary Organic Solvents

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Abstract – Under ordinary atmospheric circumstances, the gravimetric technique was used to measure the solubility of L-cysteine (L-Cys) and L-alanine (L-Ala) in various solvents, including methyl alcohol, ethyl acetate, and mixtures of the two, in the range of 283.15 K to 323.15 K. Both individual solvents and their combinations showed a rise in the solubility of L-Cys and L-Ala with increasing temperature, according to the analyzed data but when analyzed at a constant temperature in the selected mixed solvents, the solubility declined with decreasing of initial mole fractions of methyl alcohol. To further assess, the relative utility of the four solubility models, we fitted the solubility data using the Jouyban-Acree (J-A), van't Hoff-Jouyban-Acree (V-J-A), Apelblat-Jouyban-Acree (A-J-A), and Ma models followed by evaluation of the values of the RAD information criteria and the RMSD were. The dissolution was also found to be an entropy-driven spontaneous mixing process in the solvents since the thermodynamic parameters of the solvents were determined using the van't Hoff model. In order to support the industrial crystallization of L-cysteine and L-alanine and contribute to future theoretical research, we have determined the experimental solubility, correlation equations, and thermodynamic parameters of the selected amino acids during the dissolution process.

Key words: L-cysteine, L-alanine, Solubility, Mixed solvents, Thermodynamic modeling, Thermodynamic properties

1. Introduction

Amino acids such as L-cysteine and L-alanine are integral part of human body and found in many proteins [1,2]. Their chemical structures are shown in Figure 1, and comprehensive information is also given in Table 1. Because of their versatility, amino acids are considered essential in many fields, including cosmetics, food, medicine, and fine chemicals [3-6]. The solubility and thermodynamic properties of biomolecules are crucial physicochemical features that influence the efficacy of purification and separation techniques from solutions. Chemical structure, ionic strength [7], experimental temperature [8], solvent polarity [9], media acidity or alkalinity [10,11], and concentration of electrolytes in aqueous solutions [12], are some variables that affect the solubility of a biomolecule. The Thermodynamics of solvation is also greatly affected by these parameters.

Proteins rely on amino acids as their building blocks, which are essential for the metabolism of all living organisms [4,13-15]. Crystallization is often used in commercial amino acid synthesis procedures to purify the synthesized product. While designing crystallization procedures, understanding of the solubility of the solute is crucial, since it determines both the starting concentration

and the final concentration of the solute in the solvent. L-Cys and L-Ala solubility have recently been shown in various solvents and aqueous solvent combinations [16,17]. Our comprehensive literature search revealed no information on the solubility of L-Cys and L-Ala in methyl alcohol and ethyl ethanoate mixed binary solutions at the indicated temperatures.

The solubility of L-Cys and L-Ala in methyl alcohol plus ethyl ethanoate solvent mixtures was measured gravimetrically at atmospheric pressure and temperatures ranging from 283.15 to 323.15 K. The experimental solubilities of L-Cys and L-Ala were correlated using the J-A, V-J-A, A-J-A, and Ma models to determine the impact of temperature and solvent composition on solubility. Additionally, the dissolving processes of L-Cys and L-Ala had their thermodynamic parameters computed, which included changes in molar enthalpy (ΔH_{sol}°), molar entropy (ΔS_{sol}°), and Gibbs free energy (ΔG_{sol}°).

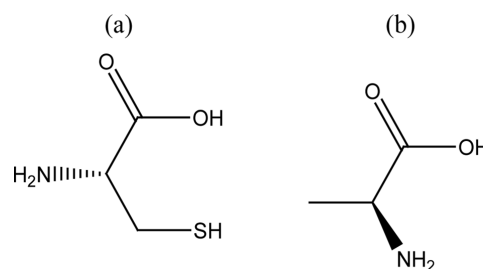


Fig. 1. Molecular structure of L-cysteine (a) and L-alanine (b).

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Table 1. Description of solutes and solvents used in the experiments

Chemicals	Purity (%)	Source	CAS no.
L-cysteine	≥ 99.0	Heibei Huayang Biological technology Co.,Ltd.	52-90-4
L-alanine	≥ 0.990	Heibei Huayang Biological technology Co.,Ltd.	56-41-7
methyl alcohol	≥ 99.5	Sinopharm Chemical Regent Co. Ltd	67-56-1
Ethyl acetate	≥ 99.5	Sinopharm Chemical Regent Co. Ltd	141-78-6

2. Experimental

2-1. Materials

L-cysteine and L-alanine was purchased from Heibei Huayang Biological Technology Co., Ltd. with their mass fraction purity more than 0.98. Reagents of analytical quality, including methyl alcohol and ethyl ethanoate, were obtained from Sinopharm Chemical Reagent Co. Ltd. and used without further purification. Their mass fraction purities exceeded 0.995. Comprehensive details regarding all materials employed in the experiments are listed in Table 1.

2-2. Apparatus and procedures

Within the temperature range of 283.15 K to 323.15 K and pressures of 0.1 MPa, this research investigated the solid-liquid equilibrium of L-Cys and L-Ala using the gravimetric technique, previously published in details [18-22].

The process involved, placing the excess drug of the solid sample (AA) and a predetermined quantity of chosen solvents into a double-jacketed glass container with a specified temperature control (± 0.05 K). Water circulation was maintained throughout the glass container to uphold the specified temperature. The mixed solution was continuously stirred for over 12 hours to attain dissolution equilibrium. Subsequently, the solid-liquid solution was allowed to stratify for 12 hours, stirring was halted, and precipitation occurred. Following this, a 5 mL sample was extracted from the upper liquid layer, transferred to a double dish, weighed, and subjected to a 12-hour drying process at 323.15 K. The experimental solubility was measured thrice to minimize errors. The validity of the entire setup has been verified with previously published works [18-22].

The mole fraction solubility of AA (designated as x_{AA}^{exptl}) in the chosen binary solvents was determined using Eq. (1), while the methyl alcohol mole fraction (denoted as x_M) used in Eq. (2) to calculate in every system including a mixture of solvents.

$$x_{AA}^{exptl} = \frac{m_2/M_1}{m_1/M_1 + m_2/M_2 + m_3/M_3} \quad (1)$$

$$x_M = \frac{m_2/M_2}{m_2/M_2 + m_3/M_3} \quad (2)$$

In this case, m_1 , m_2 and m_3 stand for the masses of AA, methyl alcohol, and ethyl ethanoate, independently; M_1 , M_2 , M_3 designate the molar masses of AA, methyl alcohol, and ethyl ethanoate, respectively.

3. Results and Discussion

3-1. Solubility data

Prior research has shed light on the solubility of L-Cys and L-Ala in methyl alcohol and ethyl ethanoate between 283 K and 323 K [16, 17]. Figure 2 displayed the study findings and observations. With a significant relative error of around 2% between them, these numbers align with our findings.

Tables 2-3 and Figure 3 showed resulting L-Cys and L-Ala solubility data of each binary system. The results show that at a given temperature, the experimental solubility of selected amino acids increased monotonically with an increasing mole fraction of methyl alcohol. There was a temperature-dependent increase in the solubility of L-Cys and L-Ala, even when the solvent composition remains constant.

3-2. Thermodynamic modelling

Several models have successfully correlated the solubility of solid solutes in mixtures of solvents. The experimental solubilities of AA in methyl alcohol, ethyl ethanoate, and (methyl alcohol + ethyl ethanoate) solvent mixtures were associated in this work while using the J-A model, V-J-A model, A-J-A model and Ma model. The J-A model, V-J-A model, A-J-A model and Ma model are related to solubility or solution thermodynamics. Their physical implications may involve predicting and understanding the interaction pattern of

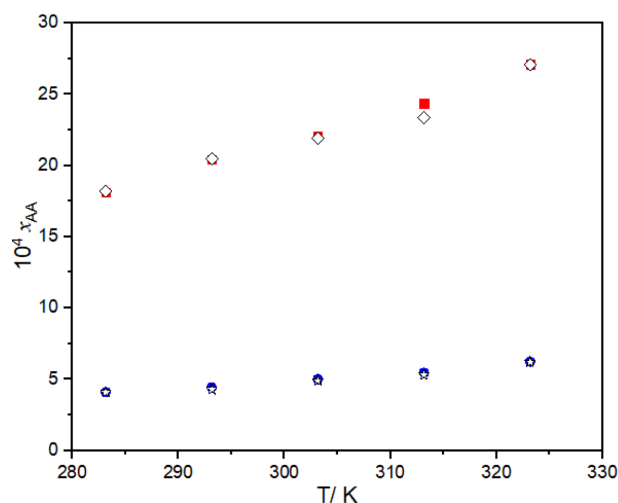


Fig. 2. Graphical comparison of experimental solubilities of selected amino acids in methyl alcohol with literature values [16,17] at various temperatures: this work (■, L-cysteine; ●, L-alanine) and literature (◇, L-cysteine; ☆, L-alanine).

Table 2. Experimental (x_{explt}) and calculated (x_{calcd}) mole fraction solubility of L-cysteine in (methyl alcohol + ethyl acetate) mixture at several temperatures and $p=101.3$ kPa^a

T /K	J-A model		V-J-A model		A-J-A model		Ma Model		J-A model		V-J-A model		A-J-A model		Ma Model	
	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$
	0.0								0.20							
283.15	0.240	0.240	0.240	0.235	0.228	0.686	0.679	0.665	0.670	0.656						
293.15	0.290	0.290	0.290	0.331	0.336	0.831	0.804	0.914	0.891	0.906						
303.15	0.580	0.580	0.580	0.469	0.482	1.304	1.415	1.229	1.192	1.225						
313.15	0.670	0.670	0.670	0.666	0.673	1.626	1.610	1.621	1.604	1.625						
323.15	0.890	0.890	0.890	0.949	0.918	2.174	2.055	2.103	2.168	2.117						
	0.4								0.6							
283.15	2.080	2.037	1.989	2.030	1.994	5.683	5.590	5.445	5.622	5.533						
293.15	2.508	2.360	2.583	2.538	2.574	6.457	6.354	6.708	6.631	6.695						
303.15	3.262	3.645	3.298	3.202	3.277	8.278	8.641	8.151	7.914	8.049						
313.15	4.160	4.081	4.145	4.073	4.120	9.688	9.543	9.782	9.541	9.619						
323.15	5.087	4.994	5.136	5.214	5.121	11.392	11.227	11.607	11.599	11.431						
	0.8								1.0							
283.15	12.161	12.140	11.626	12.323	12.151	18.120	18.120	18.120	18.562	18.382						
293.15	13.673	13.650	13.702	13.824	13.891	20.410	20.410	20.410	20.060	20.110						
303.15	16.249	16.465	15.973	15.721	15.889	22.100	22.100	22.100	21.998	22.143						
313.15	18.090	18.061	18.440	18.088	18.175	24.350	24.350	24.350	24.429	24.517						
323.15	20.592	20.561	21.100	21.021	20.787	27.107	27.107	27.107	27.424	27.270						

^aStandard uncertainty u is $u(T) = 0.01$ K. The relative standard uncertainty u is $u_r(p) = 0.05$, $u_r(x_{explt}) = 0.02$.

Table 3. Experimental (x_{explt}) and calculated (x_{calcd}) mole fraction solubility of L-alanine in (methyl alcohol + ethyl acetate) mixture at several temperatures and $p=101.3$ kPa^a

T /K	J-A model		V-J-A model		A-J-A model		Ma Model		J-A model		V-J-A model		A-J-A model		Ma Model	
	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$	$10^4 x_{explt}$	$10^4 x_{calcd}$
	0.0								0.20							
283.15	0.810	0.810	0.810	0.825	0.797	0.997	1.0478	0.972	1.069	1.046						
293.15	1.108	1.108	1.108	1.069	1.090	1.358	1.370	1.304	1.337	1.361						
303.15	1.451	1.451	1.451	1.395	1.442	1.732	1.746	1.715	1.686	1.735						
313.15	1.810	1.810	1.810	1.831	1.855	2.172	2.125	2.217	2.142	2.170						
323.15	2.319	2.319	2.319	2.415	2.326	2.782	2.665	2.821	2.737	2.670						
	0.4								0.6							
283.15	1.606	1.608	1.605	1.645	1.621	2.588	2.470	2.582	2.532	2.512						
293.15	2.070	2.000	2.007	1.974	1.996	2.976	2.919	3.014	2.915	2.929						
303.15	2.474	2.466	2.474	2.393	2.437	3.346	3.484	3.484	3.396	3.425						
313.15	2.895	2.912	3.009	2.926	2.951	3.842	3.992	3.989	3.997	4.015						
323.15	3.564	3.557	3.615	3.604	3.547	4.673	4.751	4.529	4.747	4.713						
	0.8								1.0							
283.15	3.626	3.418	3.701	3.512	3.512	4.101	4.101	4.101	4.222	4.247						
293.15	3.901	3.852	4.048	3.892	3.890	4.430	4.430	4.430	4.528	4.491						
303.15	4.350	4.466	4.402	4.372	4.372	5.010	5.010	5.010	4.928	4.873						
313.15	4.956	4.981	4.762	4.971	4.974	5.462	5.462	5.462	5.432	5.407						
323.15	5.659	5.792	5.126	5.708	5.721	6.231	6.231	6.231	6.055	6.116						

^aStandard uncertainty u is $u(T) = 0.01$ K. The relative standard uncertainty u is $u_r(p) = 0.05$, $u_r(x_{explt}) = 0.02$.

solutes with solvents at various conditions. It included examining the effects of temperature, pressure, and composition on solubility in mixed solvents. The subsequent section provides detailed insights into the specifics of the selected models.

3-2.1 Jouyban-Acree model

To accurately depict dissolution in binary solvents, it was crucial

to consider not only temperature but also the influence of solvent composition on solubility. Hence, the Jouyban-Acree method was recommended [23,24].

$$\ln x_{AA} = x_B^0 \ln(x_{1,T})_B + x_C^0 \ln(x_{2,T})_B + x_B^0 x_C^0 \sum_0^J \frac{2^j}{T} (x_B^0 - x_C^0)^i \quad (3)$$

Here, x_B^0 and x_C^0 measure the proportion of ethyl ethanoate and

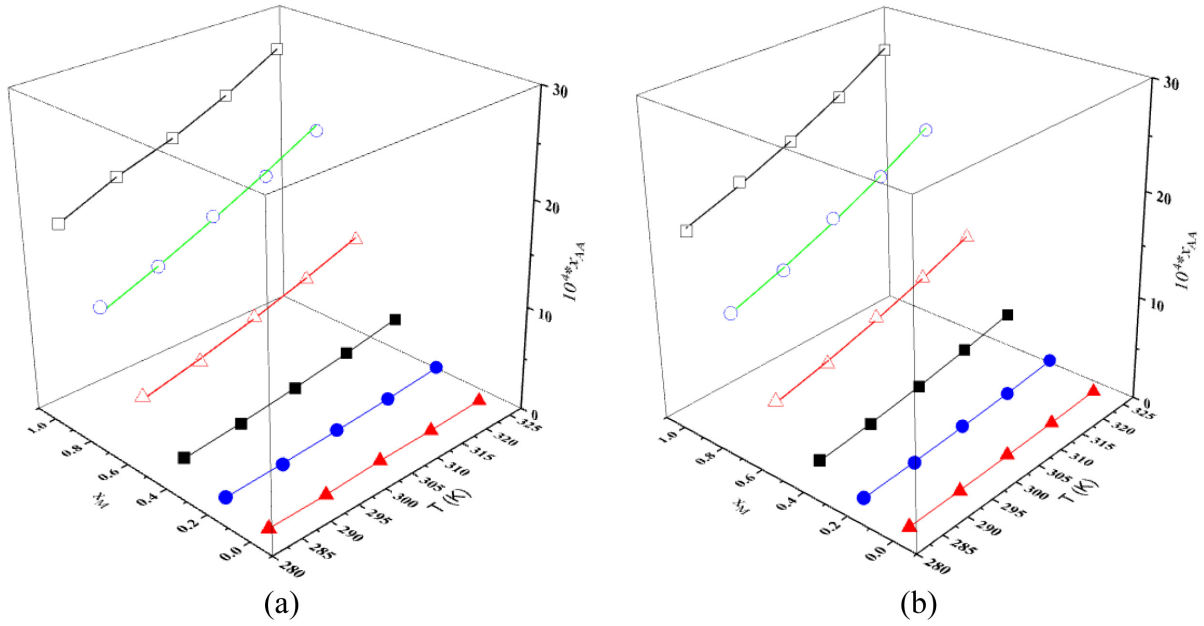


Fig. 3. Mole fraction solubility of L-cysteine in (methyl alcohol + ethyl acetate) binary solvent mixtures at various temperatures: \square , $x_M = 0.0$; \bullet , $x_M = 0.2$; \blacklozenge , $x_M = 0.4$; \star ; Δ , $x_M = 0.6$; \circ , $x_M = 0.8$; \diamond , $x_M = 1.00$; solid line, calculated with van't Hoff – Acree model (a) and Ma model (b).

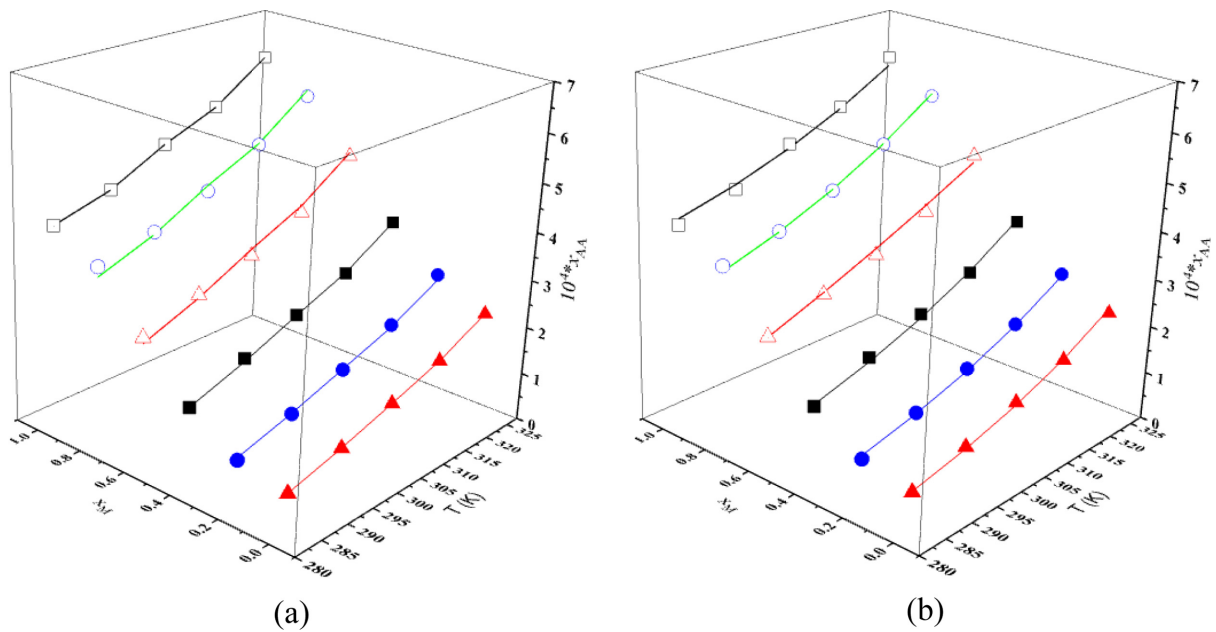


Fig. 4. Mole fraction solubility of L-alanine in (methyl alcohol + ethyl acetate) binary solvent mixtures at various temperatures: \square , $x_M = 0.0$; \bullet , $x_M = 0.2$; \blacklozenge , $x_M = 0.4$; \star ; Δ , $x_M = 0.6$; \circ , $x_M = 0.8$; \diamond , $x_M = 1.00$; solid line, calculated with Acree model (a) and Apelblat–Jouyban–Acree (b).

alcohol in a binary solvent that was free of a solute $x_{1,T}$ and $x_{2,T}$, the AA molar fraction solubility in pure ethyl ethanoate and alcohol at a given temperature T , respectively; J_i was the model parameter.

3-2-2. van't Hoff-Jouyban-Acree model

The van't Hoff model was the most straightforward equation describing the association between solubility and temperature based on the thermodynamic principles of solid-liquid equilibrium [25].

$$\ln x_{AA} = \frac{A}{T/K} + B \quad (4)$$

Eq. (5) represents the V-J-A model, which was derived by combining the van't Hoff equation with Eq. (3) [26,27].

$$\ln x_{AA} = x_B^0 \ln \left(A_1 + \frac{B_1}{T/K} \right) + x_C^0 \ln \left(A_2 + \frac{B_2}{T/K} \right) + x_B^0 x_C^0 \sum_{i=0}^{2J_i} \frac{J_i}{T} (x_B^0 - x_C^0)^i \quad (5)$$

3-2-3. Apelblat-Jouyban-Acree model

Apelblat introduced the Apelblat equation, a popular semi-empirical equation [28]. Eq. (3) can be used to explain it [29,30].

$$\ln x_{AA} = A + \frac{B}{T/K} + C \ln(T/K) \quad (6)$$

A , B , and C in this equation are the empirical constants that may be adjusted in the model. The solubility mole fraction of an amino acid in the solvents being studied at each absolute temperature T , was denoted as x_{AA} . Regarding the activity coefficient's volatility, the parameters A and B indicated that the solution was not optimum. C parameter represented the impact of temperature on the fusion enthalpy.

Similarly, the Apelblat model was constructed by combining the J - A model (Eq. 3) with the modified Apelblat equation (Eq. 6) [26, 27].

$$\ln x_{AA} = x_C^0 \ln \left(A_1 + \frac{B_1}{T/K} + C_1 \ln(T/K) \right) + x_C^0 \ln \left(A_2 + \frac{B_2}{T} + C_2 \ln(T/K) \right) + x_B^0 x_C^0 \sum_{i=0}^2 \frac{J_i}{T} (x_B^0 - x_C^0)^i \quad (7)$$

Using Eqs. (5) and (7) at different temperatures, we can determine how the solute dissolved in binary solvent mixtures.

3-2-4. Ma model

To better correlate, the experimental solubility data in solvent mixed at different temperatures, Ma [31,32] suggested the following alternative adjustment to Eq. (7):

$$\ln x_{AA} = E_0 + \frac{E_1}{T} + E_2 \ln T + E_3 x_B^0 + E_4 \frac{x_B^0}{T} + E_5 \frac{(x_B^0)^2}{T} + E_6 \frac{(x_B^0)^3}{T} + E_7 \frac{(x_B^0)^4}{T} + E_8 x_B^0 \ln T \quad (8)$$

In Eq. (8), E_0 to E_8 represent the parameters of the equation.

Using experimental solubility as input, the model parameters in Eqs. (3), (5), (7), and (8) can be derived using the nonlinear least squares technique in the MATLAB. The following was the objective function formulation:

$$F = \sum_{i=1}^n (\ln x_i^{exptl} - \ln \gamma x_i^{calcd})^2 \quad (9)$$

Also, the associated outcomes were evaluated using the RAD and RMSD, described by Eqs. (10) and (11), respectively.

$$RAD = \frac{1}{n} \times \sum_{i=1}^n \left| \frac{x_{exptl} - x_{calcd}}{x_{exptl}} \right| \quad (10)$$

$$RMSD = \left[\frac{1}{n} \times \sum_{i=1}^n (x_{calcd} - x_{exptl})^2 \right]^{1/2} \quad (11)$$

The measured value and the computed value were represented by x_{exptl} and x_{calcd} in Eqs. (10) and (11), respectively, and n was the number of experimental points.

Table 4. Model parameters, ARD and RMSD of AA in mixtures of methyl alcohol + ethyl acetate at 101 k Pa

	L-cysteine	L-alanine
J-A model		
Paramter		
J ₀	567.992	110.433
J ₁	426.614	308.540
J ₂	-6.223	-121.846
10 ² × RAD	2.257	1.731
10 ³ × RMSD	0.686	0.173
V-J-A model		
Paramter		
A ₁	-2.484	-6.817
B ₁	-1105.010	-246.849
A ₂	0.223	0.918
B ₂	-3070.124	-2983.019
J ₀	572.375	184.818
J ₁	472.850	265.467
J ₂	-39.395	-109.480
10 ² × RAD	2.173	2.080
10 ³ × RMSD	0.528	0.465
A-J-A model		
Paramter		
A ₁	-120.882	-121.090
B ₁	4408.532	4408.544
C ₁	17.539	17.313
B ₂	-161.758	-146.745
B ₂	4116.725	4117.083
C ₂	24.187	21.750
J ₀	567.992	110.562
J ₁	426.614	308.514
J ₂	-6.223	-121.818
10 ² × RAD	4.067	2.840
10 ³ × RMSD	1.539	0.463
Ma Model		
Paramter		
E ₀	20.285	66.877
E ₁	-4077.183	-5495.510
E ₂	-2.936	-10.079
E ₃	-97.549	-246.670
E ₄	6658.950	12231.800
E ₅	720.480	1378.510
E ₆	-891.787	-1529.670
E ₇	25.917	458.744
E ₈	13.981	36.142
10 ² × RAD	3.850	2.018
10 ³ × RMSD	1.640	0.307

The J-A, V-J-A, A-J-A, and Ma models were employed while fitting solubility data in binary solvent systems. The appropriate model-specific estimated values can be found in Tables 2 and 3. Table 4 displayed all correlation model's parameters, RAD, and RMSD values. For L-cysteine, the four models exhibited the following overall RAD (percent) and RMSD (×103): J-A model (2.257 and 0.686), V-J-A model (2.173 and 0.528), A-J-A model (4.067 and 1.539), Ma model (3.850 and 1.640), and for L-alanine, J-A model (1.731 and 0.173), V-J-A model (2.080 and 0.465), A-J-A model (2.840 and 0.463), and Ma model (2.018 and 0.307). Regarding L-cysteine and L-alanine, respectively, the J-A and V-J-

A models performed the best among the four models used in this research.

3-3. Thermodynamic parameters

In the solid-liquid phase equilibrium system, the thermodynamic properties of AA in binary solvent mixtures were notably significant. Eq. (12) [33,34] represented the van't Hoff analysis, which can be employed to determine the molar enthalpy (ΔH_{sol}°), molar entropy (ΔS_{sol}°), and Gibbs free energy change (ΔG_{sol}°) of AA during the dissolution process:

$$\Delta H_{sol}^{\circ} = -R \left(\frac{\partial \ln x_{AA}}{\partial (1/T - 1/T_{hm})} \right) \quad (12)$$

In this expression, T_{hm} denoted the mean harmonic temperature, calculated to be 302.338 K using Eq. (13) whereas x_{AA} represented the AA solubility.

$$T_{mean} = N / \sum_{i=1}^N \frac{1}{T_i} \quad (13)$$

In this equation, the experimental temperature, denoted as T_i , N equals the number of the temperature points measured.

The relationship between the absolute temperature's reciprocal ($1/T$) and the ($\ln x_{AA}$) was linear. The slope of the plots of $\ln x_{AA}$ versus ($1/T - 1/T_{hm}$) was equal to $-\Delta H_{sol}^{\circ}/R$. By utilizing the following equations, the change in molar enthalpy may be determined (ΔH_{sol}°) from the slope and the Gibbs free energy change (ΔG_{sol}°) from the intercept.

$$\Delta H_{sol}^{\circ} = -R \times \text{slope} \quad (14)$$

$$\Delta G_{sol}^{\circ} = -RT_{mean} \times \text{intercept} \quad (15)$$

Eq. (16) can be applied to ascertain the molar entropy change (ΔS_{sol}°) during the dissolution process:

$$\Delta S_{sol}^{\circ} = \frac{\Delta H_{sol}^{\circ} - \Delta G_{sol}^{\circ}}{T_{mean}} \quad (16)$$

Eqs. (17) and (18) [35, 36] offered the respective contributions of entropy ($\% \xi_{TS}$) and enthalpy ($\% \xi_H$) to the Gibbs free energy change (ΔG_{sol}°) of AA during the dissolution process.

$$\% \xi_H = \frac{|\Delta H_{sol}^{\circ}|}{|\Delta H_{sol}^{\circ}| + |T \cdot \Delta S_{sol}^{\circ}|} \times 100 \quad (18)$$

$$\% \xi_{TS} = \frac{|T \cdot \Delta S_{sol}^{\circ}|}{|\Delta H_{sol}^{\circ}| + |T \cdot \Delta S_{sol}^{\circ}|} \times 100 \quad (19)$$

In this paper, ΔH_{sol}° , ΔS_{sol}° , ΔG_{sol}° , $\% \xi_H$, and $\% \xi_{TS}$ for AA were calculated for saturated solution systems by using the above Eqs. (14)-(19). The summarized information was presented in Table 5. It was evident that $\Delta S_{sol}^{\circ} > 0$, $\Delta G_{sol}^{\circ} > 0$ and $\Delta H_{sol}^{\circ} > 0$ for all systems. These results pointed to heat absorption and a rise in disorder as defining features of the breakdown process of the selected amino acids (L-Cys and L-Ala). Moreover, in all chosen binary systems of

Table 5. Thermodynamic functions relative to the dissolution process of L-cysteine and L-alanine in different (methyl alcohol + ethyl acetate) mixed solvents at $T_{hm} = 302.338$ K

x_M	$\Delta_{sol}H^{\circ}$ (kJ·mol ⁻¹)	$\Delta_{sol}G^{\circ}$ (kJ·mol ⁻¹)	$\Delta_{sol}S^{\circ}$ (J·mol ⁻¹ ·K ⁻¹)	$\% \xi_H$	$\% \xi_{TS}$
L-cysteine					
0	26.541	1.913	81.388	51.869	48.109
0.2	22.306	0.450	72.290	50.488	49.490
0.4	17.495	2.941	48.095	54.588	45.412
0.6	13.803	5.218	28.394	61.634	38.346
0.8	10.209	6.945	10.795	75.759	24.225
1	7.497	5.729	5.847	80.907	19.079
L-alanine					
0	20.384	0.837	64.597	51.070	48.930
0.2	17.838	1.327	54.563	51.954	48.046
0.4	14.901	2.209	41.942	54.025	45.982
0.6	11.964	3.094	29.312	57.447	42.553
0.8	9.268	3.733	18.292	62.628	37.372
1	6.919	2.964	13.072	63.646	36.385

(methyl alcohol + ethyl ethanoate), $\% \xi_H$ was observed to exceed $\% \xi_{TS}$, signifying that the impact of heat during dissolution outperformed the contribution of entropy.

Conclusions

Between 283.15 and 323.15 K, the solid-liquid equilibrium of L-Cys and L-Ala in methyl alcohol, ethyl acetate, and (methyl alcohol + ethyl ethanoate) was examined in this work. The experimental findings showed that solvent composition and temperature, altered L-cysteine and L-alanine solubility values across all the solvent systems. A rising mole fraction of methyl alcohol increased solubility. The J-A, V-J-A, A-J-A, and Ma models were used to calculate and correlate the solubility of L-Cys and L-Ala concerning temperature and solvent composition. Neither the RAD nor the RMSD values were higher than 3.22×10^{-2} and 3.09×10^{-4} . Results from the correlation analysis were good for all the models. In addition, $\Delta G_{sol}^{\circ} > 0$, $\Delta H_{sol}^{\circ} > 0$, $\Delta S_{sol}^{\circ} > 0$, showed that L-Cys and L-Ala dissolved in an entropic and heat-absorbing manner. The crystallization technique for L-Cys and L-Ala might be optimized and designed using the experimental solubility data.

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Declarations

Conflicts of interest: The authors state that they have no competing financial interests.

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