



A

# Al-Azhar Bulletin of Science Basic Science Sector

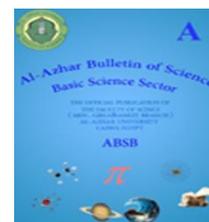
THE OFFICIAL PUBLICATION OF  
THE FACULTY OF SCIENCE  
( MEN , GIRLS & ASSUIT BRANCH )  
AL-AZHAR UNIVERSITY  
CAIRO, EGYPT

ABSBS



$\pi$





## THE ELECTROCHEMICAL SENSING OF NALBUPHINE HYDROCHLORIDE DRUG SUBSTANCE IN VITRO BY THE CYCLIC VOLTAMMETRIC AND CONDUCTOMETRIC TITRATION TECHNIQUES.

Ammar A. El-Fayoumi<sup>1</sup>, Essam A. Gomaa<sup>2</sup>, Badr A. Elsayed<sup>3\*</sup>, Magdy M. B. Awad<sup>3</sup>.

<sup>1</sup>M.Sc. Student, Chemistry Department, Faculty of Science, Al-Azhar University, Nasr City Cairo, 11884, Egypt.

<sup>2</sup>Chemistry department, Faculty of Science, Mansoura University, Mansoura, Egypt.

<sup>3</sup>Chemistry department, Faculty of Science, Al-Azhar University, Nasr City Cairo, 11884, Egypt.

\*Corresponding Author: Badr A. Elsayed.

E-mail: badrelsayed@gmail.com, [badrelsayed@azhar.edu.eg](mailto:badrelsayed@azhar.edu.eg).

Received: 08 Jan 2021; Revised: 06 Feb 2021; Accepted: 30 Mar 2021; Published: 28 Sep 2021

### ABSTRACT

The electrochemical sensing of Nalbuphine hydrochloride drug substance in vitro by the cyclic voltammetry technique using the activated glassy carbon electrode. The solvation of CdCl<sub>2</sub> alone or in the presence of Nalbuphine HCl was studied by CV technique in 0.05 M KCl supporting electrolyte and different concentrations from CdCl<sub>2</sub> at 305.15K. The cyclic voltammograms were preceded at different scan rates 0.1, 0.05, 0.02 and 0.01 V Sec<sup>-1</sup>. Also, different Nalbuphine HCl concentrations were utilized for studying their effect as electrochemical sensors on the solvation and kinetics parameters of CdCl<sub>2</sub>. The redox mechanism of the system was determined from the resulted data. Moreover, the Gibbs free energies of the complex formation were evaluated. The formation constants and Gibbs free energies were calculated from the conductometric titration curves. The molar ratios of the complexes were obtained indicating the formation of 1:2 and 1:1 (M:L). The formation constants of different complexes in water: methanol solvent followed the order: K<sub>f</sub> (1:2) > K<sub>f</sub> (1:1) for (M:L). The (ΔH) and (ΔS) of formation and association of complexes were also estimated and discussed. The solvation ΔG°, ΔHs and ΔSs were calculated from solubility measurements for nalbuphine HCl at different temperatures.

**Keywords:** Electrochemical sensor; Solvation; Cyclic voltammetry; Nalbuphine.

### 1. INTRODUCTION

Nalbuphine hydrochloride (NP.HCl) is a phenanthrene derivative. It is an opioid analgesic where it has mixed opioid agonist and antagonist activity. It is used for the relief of moderate to severe pain and as adjunct to anaesthesia. Nalbuphine has the IUPAC name (5α,6α)-17-(cyclobutyl-methyl)-4, 5 Epoxymorphinan -3,6,1 4-triol hydrochloride; N-cyclobutylmethyl-14-hydroxydihydro-normorphine (Fig. 1)[1].

Cadmium is a well-known heavy metal with extra toxicity impact on human organs. It is widely exist in human's body, the main sources of cadmium toxicity being smoking and welding.

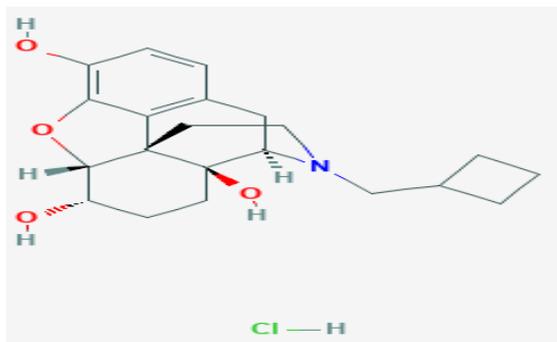
The electrochemical redox demeanor of cadmium had been determined by various electrochemical methods like hydrodynamic voltammetry, chronoamperometry, [2] coulometry [3], and polarography. There are many studies about cadmium cyclic voltammetry with many ligands.

Many literatures have been established for the sensing of Nalbuphine hydrochloride in pharmaceuticals using HPLC with UV or electrochemical detection [4].

In current work glassy carbon electrode was utilized in a sensing of Nalbuphine hydrochloride with cadmium salt using cyclic voltammetric determination and conductometric titrations.

Moreover, electroanalytical methods are less expensive, more portable and ease of handling than

the aforesaid techniques. Specifically, cyclic voltammetry is more officially used in industrial, environmental work and in the drug detection in their dosage forms and especially in biological samples since it combines excellent sensitivity, selectivity, accuracy, and precision with low cost of instrumentation and maintenance.



**Fig. 1.** Structure of Nalbuphine hydrochloride.

## 2. OBJECTIVES

Thermodynamic study of complexation reactions of Nalbuphine hydrochloride with cadmium ions not only result an interested data on the thermodynamics of complexation reaction, but also gives a better understanding of the high sensitivity of this ligand towards cadmium ions.

This work focuses on the determination of the stability constants and thermodynamic functions of complexation reactions between Cadmium ions and nalbuphine hydrochloride using the cyclic voltammetric and conductometric techniques.

## 3. EXPERIMENTAL

### 3.1 Materials

Water used in the preparation of solutions was bidistilled with a specific conductivity of  $0.07 \mu\text{S cm}^{-1}$  at 298.15 K, all salts ( $\text{CdCl}_2$  and  $\text{KCl}$ ) were purchased from merck, The used Methanol solvent ( $\text{MeOH}$ ) was obtained from CHEM-LAB nv Co.

### 3.2 Cell and Instrument

For cyclic voltammetry, the cell used containing three electrodes connected to potentiostat DY 2000,  $\text{Ag}/\text{AgCl}$  in saturated  $\text{KCl}$  solution was used as the reference electrode, glassy carbon electrode (GCE) used as the working electrode and Platinum wire electrode was used as the auxiliary electrode. The glassy carbon electrode was prepared in the laboratory from pure carbon piece and

polished with fine aluminum oxide on the wet woolen piece and its surface area is  $0.0314 \text{ cm}^2$ . The conductometric titration of the  $\text{CdCl}_2$  ( $1 \times 10^{-4}$  mole/L) against the ligand (Nalbuphine HCl  $1 \times 10^{-3}$  mole/L) in Methanol: water 30: 70 v/v was performed with 0.2 ml interval addition of Nalbuphine HCl solution. The specific conductance values were recorded using conductivity bridge ADWA, AD 3000 with a cell constant equal to  $1 \text{ cm}^{-1}$ . The temperature was adjusted at 293.15 K, 298.15 K, 303.15 K and 308.15 K respectively.

## 3. RESULTS AND DISCUSSIONS

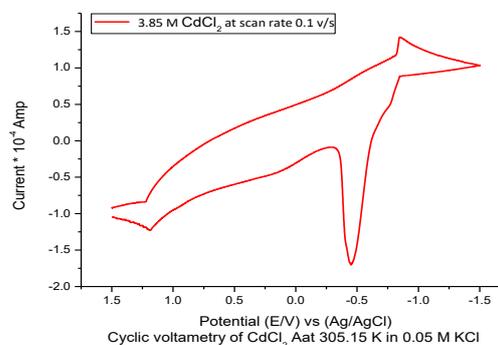
### 4.1 Cyclic voltammetry measurements

#### 4.1.1 Cyclic voltammetry of $\text{CdCl}_2$ with Nalbuphine

##### 4.1.1.1 Oxidation and reduction mechanism of $\text{CdCl}_2$ in absence of Nalbuphine HCl.

The redox behavior of  $\text{Cd}$  (II) in  $\text{CdCl}_2$  was examined in 0.05 M of  $\text{KCl}$  as a supporting electrolyte by cyclic voltammetry on GCE at temperatures (305.15 K). This process was measured from 1.5 to -1.5 V of potential window, the resulted current measured in Amp. and 0.1 V/S scan rate. The  $\text{CdCl}_2$  solution is added step wisely to reach the final concentration ( $3.85 \times 10^{-3}$  M) as shown in Fig.2 which illustrate that  $\text{Cd}^{2+}$  solution is electroactive since it gives one anodic peak ( $\text{Cd}^{2+}/\text{Cd}^{\text{I}}$ ). At 305.15 K the anodic peak current of the anodic peak is  $4.91 \times 10^{-4}$  Amp., and its anodic potential is -0.4975 V. These results indicate that the  $\text{Cd}^{2+}$  ion system is irreversible system involving the transfer of one electron.

Cyclic voltammetry behavior of  $\text{Cd}$  (II) scan between 1500 mv to - 1500mv is introduced in (Fig.2).



**Fig. 2.** Cyclic voltammograms of  $\text{CdCl}_2$  in absence of ligand (Nalbuphine HCl) at 303.15 K

The diffusion coefficient was estimated by using Randles-Sevcik equation (1) [5] which indicates the relation among the peak current, scan rate and diffusion coefficient.

$$i_p = 0.4463 n F A C (n F D v / RT)^{1/2} \quad (1)$$

Where ( $i_p$ ) is the peak current in Ampere, ( $n$ ) is number of electrons transferred in the redox event (usually 1), ( $F$ ) is Faraday's constant which equals 96485.33 C.mol<sup>-1</sup>, ( $A$ ) is the area of the working electrode in cm<sup>2</sup>, ( $C$ ) is the bulk concentration of metal ion (mol.L<sup>-1</sup>), ( $D$ ) is the diffusion coefficient in cm<sup>2</sup>/sec, ( $v$ ) is the scan rate in volts/sec, ( $R$ ) is the gas constant which equals 8.314J.mol<sup>-1</sup>.K<sup>-1</sup> and ( $T$ ) is the absolute temperature.

Additionally, the surface coverage  $\Gamma$  (mol.cm<sup>-2</sup>) and the Quantity of charge ( $Q$ ) [6] were estimated from equations (2, 3).

$$\Gamma = i_p / 4RT/n^2 F^2 A v \quad (2)$$

$$Q = n F A \Gamma \quad (3)$$

#### 4.1.1.2 Effect of different concentrations of CdCl<sub>2</sub> at 305.15 K

The impact of several concentrations of CdCl<sub>2</sub> (1.61, 2.38, 2.68, 3.13 & 3.85) x 10<sup>-3</sup> M have been studied in 0.05 M KCl at 305.15 K

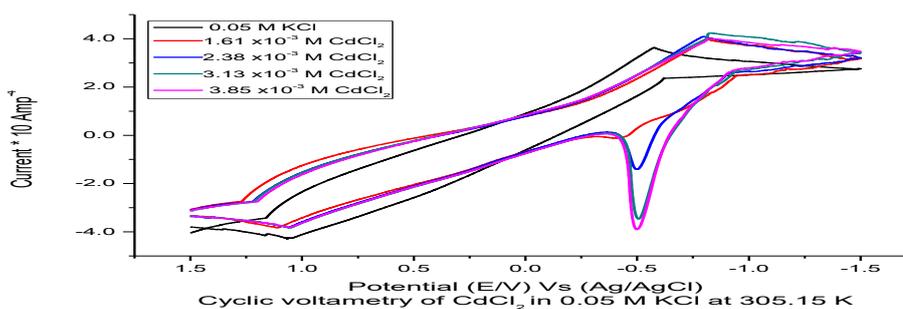


Fig. 3. Cyclic voltammograms of different of concentration for CdCl<sub>2</sub> in absence of Nalbuphine HCl.

Table 1: Solvation and kinetic parameters ( $D$ ,  $\Gamma$  and  $Q$ ) of CdCl<sub>2</sub> in the absence of Nalbuphine HCl at 305.15 K, scan rate 0.1 V/Sec

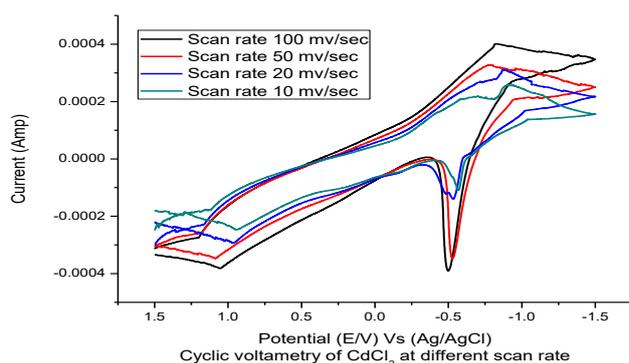
[M] x 10 <sup>-3</sup> mol.L <sup>-1</sup>	Ep,a Volt	Ip,a x 10 <sup>-4</sup> Amp	Da x 10 <sup>-10</sup> cm <sup>2</sup> .s <sup>-1</sup>	Γ a x 10 <sup>-8</sup> mol.cm <sup>-2</sup>	(-) Qa x 10 <sup>-4</sup> C
1.61	-0.434	0.42	0.93	1.45	0.44
2.38	-0.501	2.01	10.00	6.99	2.12
3.13	-0.512	4.36	27.32	15.13	4.58
3.85	-0.498	4.91	22.81	17.03	5.16

as displayed in Fig.3. The anodic peak gradually increased by raising the concentration of Cd ions which gives a proof that the reaction is governed by diffusion processes. Moreover, the solvation parameters like ( $\Gamma_a$  &  $Q_a$ ) increased as reported in Table (1). This behavior could be explained as the increase in concentrations of the electroactive species.

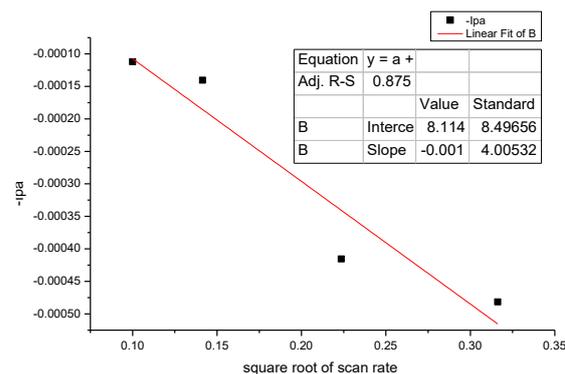
Increase in the anodic diffusion coefficient  $D_a$ , anodic surface coverages and Quantity of charge for anodic parts was increased by increasing cadmium chloride concentration indicating diffusion controlled reactions.

#### 4.1.1.3 Effect of different scan rates for CdCl<sub>2</sub>

The influence of scan rates (0.1, 0.05, 0.02 & 0.01) V.S<sup>-1</sup> was determined at 305.15 K as shown in Fig.4. The different solvation parameters increased by the decreasing of the scan rate supporting the diffusion mechanism of solvation as reported in Table (2). Also, the redox peak current decreased by decreasing the scan rates. The linear relation between  $-I_{pa}$  and the square root of scan rate in Fig. (5). This demonstrated that the reaction was governed by the surface diffusion process [7].



**Fig.4.** Effect of different scan rate of  $3.85 \times 10^{-3} \text{M CdCl}_2$ .



**Fig.5.** Relation ( $i_p$  vs  $\sqrt{v}$ ) for  $\text{CdCl}_2$  at final addition in absence of Nalbuphine HCl at 305.15 K and different scan rate.

**Table 2:** Correlation coefficient measurements

<b>Number of Points</b>	<b>4</b>
<b>Degrees of Freedom</b>	<b>2</b>
<b>Residual Sum of Squares</b>	<b>8.79E-09</b>
<b>Adj. R-Square</b>	<b>0.87597</b>

**Table 3:** Solvation and kinetic parameters ( $D$ ,  $\Gamma$  and  $Q$ ) of effect of scan rate on final addition of  $\text{CdCl}_2$  at 305.15 K.

Scan rate V/sec	$E_{p,a}$ volt	$i_{pa} \times 10^{-4}$ Amp	$D_a \times 10^{-9}$ $\text{cm}^2/\text{Sec}$	$\Gamma_a \times 10^{-7}$ $\text{mol}/\text{cm}^2$	$(-)Q_a \times 10^{-4}$
<b>0.10</b>	-0.498	4.91	2.28	1.70	5.16
<b>0.05</b>	-0.529	4.18	3.31	2.90	8.80
<b>0.02</b>	-0.533	1.40	0.94	2.44	7.39
<b>0.01</b>	-0.571	1.12	1.19	3.89	11.8

#### 4.1.1.4 Effect of Nalbuphine HCl on the redox behavior of $\text{CdCl}_2$

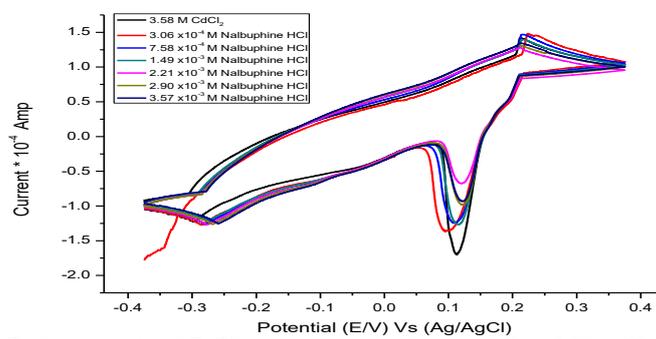
The influence of Nalbuphine HCl different concentrations (0.31, 0.76, 1.49, 2.21, 2.90, 3.57)  $\times 10^{-3}$  M on the electrochemical behavior of  $\text{CdCl}_2$  was determined at 305.15 K and potential range from 1.5 V to -1.5 V as shown in Fig.6.

As reported in Table (3) the addition of Nalbuphine hydrochloride to the solution and stepwisely increasing its concentration, the redox peaks current decreased than observed with  $\text{CdCl}_2$  alone as mentioned in Table (1).

#### 4.1.1.5 Effect of different scan rates in presence of Nalbuphine HCl

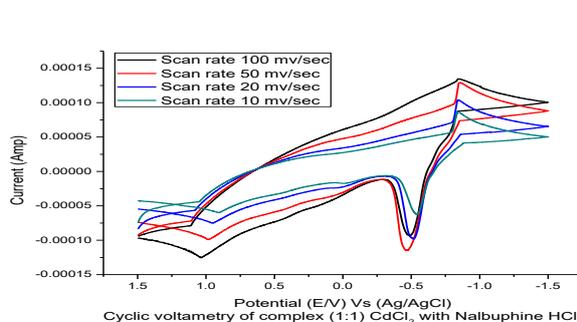
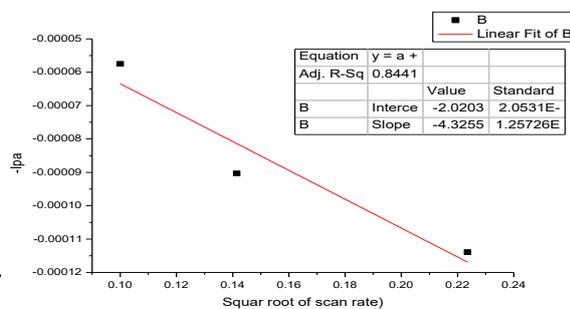
The influence of different scan rates (0.1, 0.05, 0.02, 0.01)  $\text{V}\cdot\text{S}^{-1}$  for 1:1 molar ratio between cadmium and nalbuphine had been studied as shown in Fig. 7. The decrease of the scan rate was followed by the decrease of the redox peaks currents, but the solvation parameters like ( $\Gamma_a$  &  $Q_a$ ) increased as shown in Table (4).

The linear relation between the peak current and square root of scan rate confirmed that the reaction was governed by diffusion processes as shown in Fig. (8).

Cyclic voltammetry of CdCl<sub>2</sub> in presence of nalbuphine HCl in 0.05 M KCl at 305.15 K**Fig. 6** Cyclic Voltammogram of CdCl<sub>2</sub> in the presence of Nalbuphine HCl at 305.15K**Table 4:** Solvation and kinetic parameters (D,  $\Gamma$  and Q) of CdCl<sub>2</sub> in the presence of Nalbuphine HCl at 305.15 K, scan rate 0.1 V/Sec

[L] x10 <sup>-3</sup> mol.L <sup>-1</sup>	E <sub>p,a</sub> Volt	I <sub>p,a</sub> x10 <sup>-4</sup> Amp	Dax10 <sup>-10</sup> cm <sup>2</sup> .s <sup>-1</sup>	$\Gamma_a$ x10 <sup>-8</sup> mol.cm <sup>-2</sup>	Q <sub>a</sub> x10 <sup>-4</sup> C
0.31	-0.406	1.35	1.74	4.68	1.42
0.76	-0.461	1.24	1.51	4.32	1.31
1.49	-0.465	1.27	1.61	4.39	1.33
2.21	-0.482	0.74	0.57	2.57	0.78
2.90	-0.491	1.01	1.09	3.51	1.06
3.57	-0.495	0.95	0.98	3.29	0.997

Most of the analysis of the redox processes data are smaller in case of CdCl<sub>2</sub> plus Nalbuphine HCl ligand than CdCl<sub>2</sub> alone indicating the complex reaction between metal and ligand, hence the method is sensitive to detect Nalbuphine hydrochloride.

**Fig.7.** Effect of different scan rate of 1:1 rate of scan rate for (Cd-Nalbuphine) complex.**Fig.8.** The relation between  $-I_{p_a}$  and square Cadmium chloride in presence of Nalbuphine HCl.**Table 5:** Correlation coefficient measurements:

Number of Points	3
Degrees of Freedom	1
Residual Sum of Squares	1.25E-10
Adj. R-Square	0.84419

**Table 6:** Solvation and kinetic parameters (D,  $\Gamma$  and Q) of effect of scan rate on complex at 305.15 K.

Scan rate V/sec	E <sub>p,a</sub> volt	i <sub>p,a</sub> x10 <sup>-5</sup> Amp	D <sub>a</sub> x10 <sup>-10</sup> cm <sup>2</sup> /Sec	$\Gamma_a$ x10 <sup>-8</sup> mol/cm <sup>2</sup>	Q <sub>a</sub> x10 <sup>-4</sup> C
0.10	-0.49	9.48	0.987	3.29	0.997
0.05	-0.47	11.4	2.85	7.91	2.40
0.02	-0.51	9.03	4.48	15.70	4.75
0.01	-0.55	5.75	3.63	20.00	6.05

#### 4.1.1.6 The stability constant for (Cd-Nalbuphine) complex

The values of stability constant ( $\log \beta_{MX}$ ) and Gibbs free energy ( $\Delta G^\circ$ ) increased by increasing the  $j$  (L/M) ratio, indicating the tendency towards the formation of the complex as reported in Table (5). The decrease of scan rate was followed with the decrease of the stability constant of the complex ( $\log \beta_{MX}$ ) and the Gibbs free energy as reported in Table (6).

The stability constant ( $\log \beta_{MX}$ ) of the Cd-Nalbuphine complex was calculated from equation (4) [8].

$$\Delta E^\circ = E^\circ_c - E^\circ_m = 2.303(RT/nF) * (\log \beta_{MX} + j \log C_x) \quad (4)$$

Where ( $E^\circ_c$ ) is the potential of the complex after every addition of (ligand) Nalbuphine HCl, ( $E^\circ_m$ ) the potential of the last addition of metal (cadmium) before adding the ligand (Nalbuphine HCl), ( $C_x$ ) the ligand concentration in the solution,  $j$  (the coordination number of the complex)

The Gibbs free energy for the interaction between cadmium and the Nalbuphine HCl was determined from the stability constant by applying equation (5) [9].

$$\Delta G^\circ = -2.303 RT \log \beta_{MX} \quad (5)$$

## 4.2 Conductometric measurements

The specific conductance values ( $K_s$ ) of the solutions of different concentrations of  $CdCl_2$  solution in (MeOH-H<sub>2</sub>O) mixtures were measured experimentally in absence and in presence of ligand at different temperatures (293.15, 298.15, 303.15 and 308.15 K).

The molar conductance ( $\Lambda_m$ ) values were calculated [10] using equation (6):

$$\Lambda_m = \frac{(K_s - K_{solv}) \cdot K_{cell} \cdot 1000}{C} \quad \dots\dots\dots (6)$$

Where  $K_s$  and  $K_{solv}$  are the specific conductance of the solution and the solvent, respectively;  $K_{cell}$  is the cell constant and  $C$  is the molar concentration of the metal salt solution.

#### 4.2.1 Formation constant for the complexation of the $CdCl_2$ with Nalbuphine HCl

The experimental data of ( $\Lambda_m$ ) were analyzed for the determination of formation constants for each type of the stoichiometric complexes. The formation constants ( $K_f$ ) for  $CdCl_2$  complex were calculated for each type of complexes (1:2) and (1:1) (M:L) [11] by using the equations:

$$M^{2+} + L \rightleftharpoons ML^{2+} \quad \dots (7)$$

$$K_f = \frac{[ML]}{[M][L]} = \frac{\Lambda_M - \Lambda_{obs}}{(\Lambda_{obs} - \Lambda_{ML})[L]} \quad \dots (7)$$

$$[L] = C_L - \left\{ C_M * \frac{\Lambda_M - \Lambda_{obs}}{(\Lambda_M - \Lambda_{ML})} \right\} \quad \dots(8)$$

Where  $\Lambda_m$  is the molar conductance of the metal before adding the ligand,  $\Lambda_{obs}$  is the molar conductance of solution during titration and  $\Lambda_{ml}$  is the molar conductance of the complex. The obtained values of  $\log (K_f)$  for the metal-ligand stoichiometric complexes are presented in Table (7) for  $CdCl_2$  in (MeOH-H<sub>2</sub>O) mixture.

The relation between  $\Lambda_m$  and the  $[M]/[L]$  molar ratio for  $CdCl_2$  in presence of Nalbuphine HCl in 30 % (MeOH-H<sub>2</sub>O) mixture as shown in Fig. (9).

The inflections in Fig.9 indicate the formation of different complexes.

Increasing temperature is followed by decrease in  $\log K_f$  favouring less solvation for interaction of  $CdCl_2$  with Nalbuphine HCl indicating migration of ions away from the collecting area.

It was obviously that complex formation between  $CdCl_2$  and Nalbuphine HCl in (Methanol: Water) as a solvent:

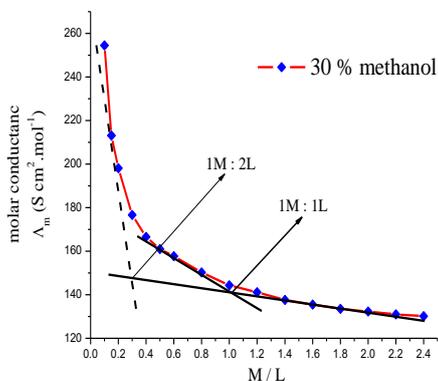
- 293.15 K is favorable temperature in the formation of 1:1  $[M]/[L]$
- 298.15 K is favorable temperature in the formation of 1:2  $[M]/[L]$
- 303.15 K is favorable temperature in the formation of 1:2  $[M]/[L]$
- 298.15 K is the same in the formation of 1:2 and 1:1  $[M]/[L]$

The Gibbs free energies of formation for 1:1 and 1:2 (M:L) stoichiometry complexes ( $\Delta G_f^\circ$ ) were calculated [12] by using the equation (9):

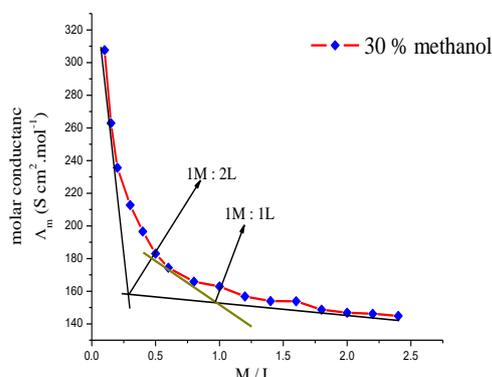
$$\Delta G_f^\circ = - 2.303 RT \log K_f \dots\dots (9)$$

The enthalpy ( $\Delta H_f$ ) for the metal salt complexes were calculated for each type of complexes, (1:2) and (1:1) (M:L) by using van't Hoff equation:

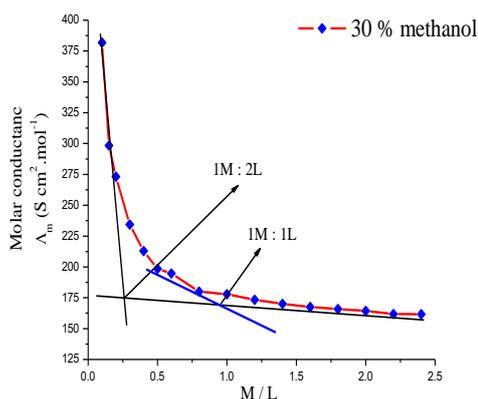
Where  $R$  is the gas constant and  $T$  is the absolute temperature. On plotting of  $\log K_f$  versus  $1/T$  different lines are obtained for the formation of 1:2 and 1:1 (M:L) stoichiometric complexes for  $CdCl_2$  with Nalbuphine HCl as shown in Fig. 10 .



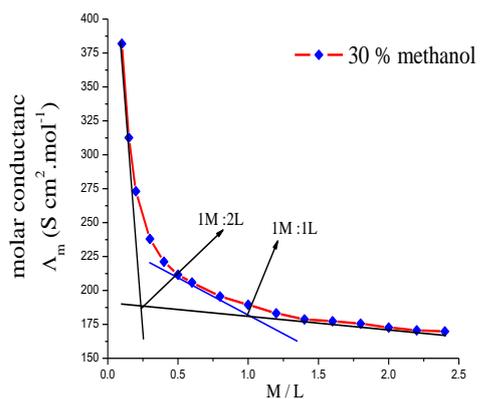
The relation between  $\Lambda_m$  and the [M]/[L] molar ratio for  $CdCl_2$  in 30 % MeOH at 289.15



The relation between  $\Lambda_m$  and the [M]/[L] molar ratio for  $CdCl_2$  in 30 % MeOH at 298.15 K



The relation between  $\Lambda_m$  and the [M]/[L] molar ratio for  $CdCl_2$  in 30 % MeOH at 308.15 K



The relation between  $\Lambda_m$  and the [M]/[L] molar ratio for  $CdCl_2$  in 30 % MeOH at 318.15 K

**Fig 9.** The relation between  $\Lambda_m$  and the [M]/[L] molar ratio for  $CdCl_2$  in presence of Nalbuphine in 30 % (MeOH-H<sub>2</sub>O) mixture.

**Table 7:**  $\Lambda_{ml}$  is the molar conductance of the complex, formation constant ( $K_f$ ), for  $Cd^{+2}$  – Nalbuphine HCl complex formation in (MeOH-H<sub>2</sub>O) mixture at different temperatures.

T (K)	M:L	[L] <sub>t</sub> x10 <sup>-5</sup>	[M] <sub>t</sub> x10 <sup>-5</sup>	$\Lambda_{ml}$ (S cm <sup>2</sup> .mol <sup>-1</sup> )	$\Lambda_{obs}$ (S cm <sup>2</sup> .mol <sup>-1</sup> )	Log $K_f$
293.15	1: 2	9.524	4.762	176.645	160.965	3.076
	1: 1	9.091	9.091	160.970	144.375	3.208
298.15	1: 2	9.524	4.762	212.695	183.020	3.345
	1: 1	9.091	9.091	183.000	162.860	3.249
303.15	1: 2	9.524	4.762	234.300	198.500	3.406
	1: 1	9.091	9.091	198.500	177.900	3.211
308.15	1: 2	9.524	4.762	237.900	211.680	3.200
	1: 1	9.091	9.091	211.700	189.420	3.220

$$\frac{d \ln K_f}{dT} = \frac{\Delta H_f^\circ}{RT^2} \dots\dots (10)$$

From the relation between  $\log K_f$  and  $1/T$ ,  $\Delta H_f$  can be calculated for each type of complexes from

the slope of each line ( $-\Delta H_f/2.303R$ ). The entropy values ( $\Delta S_f$ ) for complexes were calculated for each type of complexes (1:2) and (1:1) (M:L) by using the equation :

$$\Delta G_f^\circ = \Delta H_f - T\Delta S_f \quad \dots\dots\dots (11)$$

Where (S) is the entropy of system.

Formation thermodynamic parameters ( $\Delta G_f$ ,  $\Delta H_f$ ,  $T\Delta S_f$ ,  $\Delta S_f$ ) were collected in Table (8).

From the previous Table decreasing in  $\Delta G_f^\circ$  by increasing in temperatures indicating more spontaneous process.

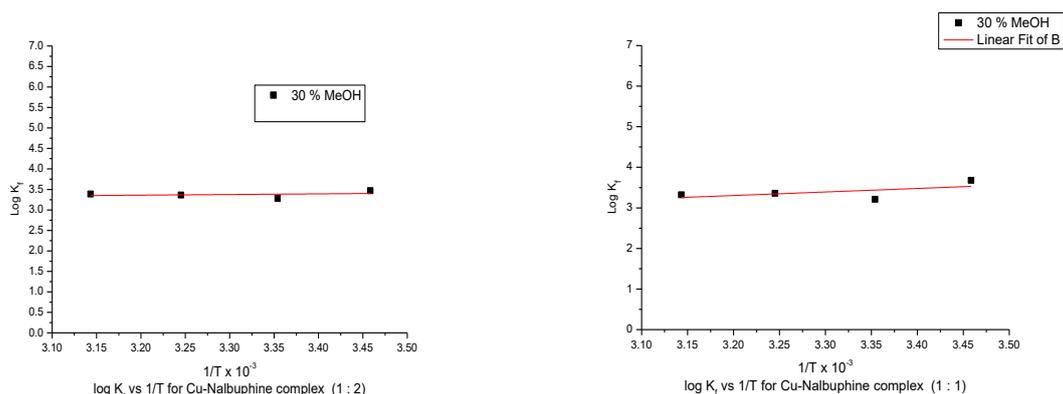
Most complex formation data are bigger in forming 1:2 complex than that of 1:1, indicating that the formation of 1: 2 complex is favorable than 1:1 complex.

## 5 CONCLUSION

- Most of the analysis of the redox processes data are smaller in case of CdCl<sub>2</sub> plus Nalbuphine HCl ligand than CdCl<sub>2</sub> alone indicating the complex reaction between metal and ligand, hence the method is sensitive to detect Nalbuphine hydrochloride.
- All scan rate measurements proved the diffusion controlled reactions.
- On drawing the relation between molar conductance and the ratio of metal to ligand concentrations, Different lines are obtained indicating the formation of 1:1 and 2:1 [L]/[M] stoichiometric complexes.
- The formation constants and Gibbs free energies of different complexes were determined. Negative values of Gibbs free energy of complexation indicate that the reaction is spontaneous.

### Declaration:

The authors declare that they have no conflict with anybody.



**Fig. 10.**  $\log K_f$  v.s  $1/T$  for Cd-Nalbuphine complexes (1:2) and (1:1)

**Table 8:** Complex formation thermodynamic parameters for Cd<sup>+2</sup> – Nalbuphine HCl complex formation in (MeOH-H<sub>2</sub>O) mixture at different temperatures.

T(K)	M:L	$\Delta G_f^\circ$ (kJ mol <sup>-1</sup> )	$\Delta H_f$ (kJ mol <sup>-1</sup> )	$T\Delta S_f$ (kJ mol <sup>-1</sup> )	$\Delta S_f$ (kJ mol <sup>-1</sup> K <sup>-1</sup> )
293.15	1:2	-17.030	7.973	25.003	0.086
	1:1	-17.759	-0.025	17.735	0.061
298.15	1:2	-19.094	7.973	27.067	0.091
	1:1	-18.547	-0.025	18.522	0.062
303.15	1:2	-20.097	7.973	28.069	0.091
	1:1	-18.949	-0.025	18.924	0.061
308.15	1:2	-19.493	7.973	27.466	0.086
	1:1	-19.616	-0.025	19.591	0.062

## 5. REFERENCES

- [1] Elqudaby HM, Hendawy HA, Souaya ER, Mohamed GG, & Eldin GM, Utility of activated glassy carbon and pencil graphite electrodes for voltammetric determination of nalbuphine hydrochloride in pharmaceutical and biological fluids. *Int J Electrochem*, 2016; <https://doi.org/10.1155/2016/8621234>
- [2] Coutinho CFB, Silva MO, Calegario ML, Machado SAS, & Mazo LH, Investigation of copper dissolution in the presence of glyphosate using hydrodynamic voltammetry and chronoamperometry. *Solid State Ionics*, 2007;178(1-2): 161-4. <https://doi.org/10.1016/j.ssi.2006.10.027>
- [3] Van Eenoo M, Labar C, and Lamberts L, A Simple Coulometric Method for Determining Cu(II) and Cu(I) Contents in Various Solid Samples, *Anal Lett*, 1986;19(21-22): 2063-78 <https://doi.org/10.1080/00032718608080866>.
- [4] Brownson DA, and Banks CE, *The handbook of graphene electrochemistry*, 2014. <https://doi.org/10.1007/978-1-4471-6428-9>
- [5] Wang J, *Analytical Electrochemistry*, 3rd ed, Wiley-VCH John Wiley & Sons publishers Inc., Hoboken, New Jersey, 2006. <https://doi.org/10.1002/0471790303>.
- [6] Christian GD, *Analytical Chemistry*, 6th Edition, John Wiley & Sons, Inc, 2004.
- [7] Gupta R, Gamare J, Sharma MK, and Kamat J V, Electrochemical investigations of Pu (IV)/Pu (III) redox reaction using graphene modified glassy carbon electrodes and a comparison to the performance of SWCNTs modified glassy carbon electrodes. *Electrochim Acta*, 2016; 191: 530-5. <https://doi.org/10.1016/j.electacta.2017.02.122>
- [8] Palmer KL, and Gilmore M, Multidrug-resistant enterococci lack CRISPR-cas. *MBio*, 2010; 1(4), <https://doi.org/10.1128/mBio.00227-10>
- [9] Barnard GM, Boddington T, Gregor JE, Pettit LD, and Taylor N, An investigation into the determination of stability constants of metal complexes by convolution—deconvolution cyclic voltammetry. *Talanta*, 1990;37(2): 219-28. [https://doi.org/10.1016/0039-9140\(90\)80026-c](https://doi.org/10.1016/0039-9140(90)80026-c)
- [10] Gomaa E, and Al-Jahdalli BM, Electrical Conductance of Cu (NO<sub>3</sub>)<sub>2</sub> with Kryptofix - 222 in Mixed ( MeOH - DMF) Solvents at Different Temperatures. *Am J Environ Eng*, 2012;2: 6-12, <https://doi.org/10.5923/j.ajee.20120202.02>
- [11] Ibrahim KM, Gomaa EA, Zaki RR, & Abd El-Hadi MN, The Association and Complex Formation Constants for CuSO<sub>4</sub>, NiSO<sub>4</sub> Stiochiometric Complexes with (E)-N'-(2-hydroxy-3H-indol-3-ylidene)-3-oxo-3-(Thiazol-2-Ylamino) Propanehydrazide in Ethanol Solutions at 294.15 K. *Structure*, 1, H2IH.
- [12] Helmy ET, Gomaa EA, & Abou Eleef EM, Complexation of 2-mercaptoimidazol with some barium salts conductometrically in various solvents at different temperatures. *Int J Modern Chem*, 2015; 7: 141-55.

### الاستشعار الكهروكيميائي لمادة عقار نالبوفين هيدروكلوريد في المختبر بواسطة تقنيات قياس الفولتميتر الدوري وقياس التوصيلية.

عمار عبدالله الفيومي<sup>1</sup> - عصام عرفة جمعة<sup>2</sup> -  
بدر الدين السيد عواد<sup>3</sup> ، مجدي محمد عواد<sup>3</sup>

- 1- طالب ماجستير، قسم الكيمياء، كلية العلوم، جامعة الأزهر، القاهرة، شركة المستقبل للصناعات الدوائية، بدر، القاهرة.  
2- قسم الكيمياء، كلية العلوم، جامعة المنصورة.  
3- قسم الكيمياء، كلية العلوم، جامعة الأزهر، القاهرة.

### الملخص العربي

تمت دراسة ذوبان كلوريد الكادميوم بمفرده أو في وجود عقار نالبوفين هيدروكلوريد بتقنية الفولتميتر الدوري في 0.05 مولر من كلوريد البوتاسيوم (KCl) كمحلول الكتروليتي داعم وتركيزات مختلفة من كلوريد الكادميوم عند 305.15 كلفن. تم إجراء تصوير الفولتموجرام الدوري بمعدلات مسح مختلفة 0.1 و 0.05 و 0.02 و 0.01 سم / ثانية. كما تم استخدام تراكيز مختلفة من عقار نالبوفين هيدروكلوريد لدراسة تأثيرها كمستشعرات كهروكيميائية على معاملات الذوبان والحركية لأيون الكادميوم. تم تحديد آلية الأكسدة والاختزال للنظام من البيانات الناتجة. علاوة على ذلك، تم تقييم طاقات جيبس الحرة لتكوين المعقد.

تم حساب ثوابت التكوين وطاقات جيبس الحرة من منحنيات معايرة التوصيلية. تم الحصول على النسب المولية للمعقد التي تشير إلى تكوين 1:1 و 1:2 (Cd: Nalbuphine) ثوابت تكوين للمعقدات المختلفة في (الماء: الميثانول) يتبع الترتيب:

$$(Cd: Nalbuphine) \downarrow (K_f (1: 2)) > K_f (1: 1)$$

كما تم تقدير ( $\Delta H$ ) و ( $\Delta G_s$ ) و ( $T\Delta S_f$ ) و ( $\Delta S_f$ ) لتشكيل المعقدات وربطها من قياسات الذوبان لنالبوفين هيدروكلوريد في درجات حرارة مختلفة.