

## NEW DINUCLEAR CYANIDO COMPLEXES WITH AMINE ALCOHOL LIGAND: SYNTHESIS, CHARACTERIZATION AND BIOTECHNOLOGICAL APPLICATION POTENTIAL

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*In this study, the cyanido complexes given by the formula  $[Ni(Abut)Ni(CN)_4] \cdot 8H_2O$  (C1),  $[Cu(Abut)_2Ni(CN)_4] \cdot 7H_2O$  (C2),  $[Zn(Abut)Ni(CN)_4] \cdot 8H_2O$  (C3) and  $[Cd(Abut)Ni(CN)_4] \cdot 7H_2O$  (C4) were obtained by microwave synthesis method. The powder forms of the complexes were characterized by elemental, FT-IR spectroscopy, and thermal analysis. And also antibacterial, antibiofilm and anticancer activities were investigated. The splitting stretching bands of cyanido groups in the FT-IR spectra of C1-C4 indicated the assets of terminal and end cyanido groups. The antibacterial activities of C1-C4 were tested with nine Gram negative and six Gram positive bacteria. The most efficient antibacterial activity of complexes was observed at  $1000 \mu g/ml^{-1}$  concentration. Anticancer activity was tested using HeLa cell line and MTT test. The studied cyanide complexes have been shown to decrease the viability of HeLa cells with  $IC_{50}$  values 14.86, 6.5, 7.2 and  $19.2 \mu g/ml$  for C1, C2, C3 and C4 complex, respectively.*

*Key words:* cyanido complexes, antimicrobial, anticancer activity, HeLa cells.

The cyanido complexes, which are still up to date in coordination chemistry and organo-metallic chemistry, are among the first coordination compounds that have been synthesized, characterized, and used [1,2]. These complexes exhibit different structures due to the ability of cyanido ligand to bind the transition metal atoms in different ways. Cyanido is one of the strong ligands after carbonyl in the spectrochemical series used in the synthesis of 1D, 2D and 3D cyanido complexes. The four-electron donor cyanido is a bidentate ligand with potentially reactive nitrogen and carbon atoms and exhibit  $\sigma$ -donor ( $C \rightarrow M$ ,  $N \rightarrow M'$ ) and  $\beta$ -acceptor

( $M \rightarrow C$ ,  $M' \rightarrow N$ ) features. The cyanido complexes have different structural properties, allowing for a wide use in the field of chemistry, biology, and material chemistry [2]. The applications of these complexes have been focused on catalysis [3], nonlinear optics, ion exchange materials, gas storage [4], molecular magnets [5-9], molecular sieves [10], enzymatic [11], antibacterial [13] and anticancer activities [5, 12, 13] and thermally stable materials [14-17]. Homoleptic tetracyanidonickelate(II) building block has been extensively used for the synthesis and design of different dimensional networks [18-21]. Cyanido complexes exhibit different structures due to

the ability of the cyanido ligand to bind differently to the second metal atoms such as Hg<sup>II</sup> [22], Mn<sup>II</sup> [23], Co<sup>II</sup> [24], Cd<sup>II</sup> [25,26], Cu<sup>II</sup> [27-29] and Zn<sup>II</sup> [30-33].

The 2-amino-1-butanol (*Abut*) ligand is a bidentate neutral ligand that can coordinate with a metal atom with nitrogen and oxygen and can form different complex structures originating from its geometry. Although many examples of cyanido complexes were found in the literature review, complexes using this ligand could not be found. This research aimed at synthesizing four cyanido complexes having general formulas ( $[M(\text{Abut})_n\text{Ni}(\text{CN})_4]$ , (M=Ni<sup>II</sup>, Cu<sup>II</sup>, Zn<sup>II</sup>, Cd<sup>II</sup>; Abut: 2-amino-1-butanol;  $n = 1$  or 2) and their spectroscopic (IR) and thermal properties, antibacterial properties on 16 different human disease-causing bacteria and antibiofilm activity.

### Materials and Methods

**Materials and instrumentation.** In this study, all chemicals NiCl<sub>2</sub>·6H<sub>2</sub>O, CuCl<sub>2</sub>·2H<sub>2</sub>O, ZnCl<sub>2</sub>, CdSO<sub>4</sub>·8/3H<sub>2</sub>O and 2-amino-1-butanol (*Abut*, C<sub>4</sub>H<sub>11</sub>NO) were obtained commercially and used without further purification. The IR spectra of **C1-C4** were recorded at room temperature in powder form between 400-4000 cm<sup>-1</sup> using a FT-IR spectrophotometer (Jasco Spectra Manager V.2). Thermal analysis of C1-C4 containing tetracyanonickelate were done. The TG, DTG and DTA (DSC) curves were recorded simultaneously in a Pt crucible using 5-10 mg of sample, in a nitrogen atmosphere, at a heating rate of 10°C/min in the temperature area 25-860°C on a Hitachi / STA 7300 thermal analyzer.

**Synthesis of complexes.** **[Ni(C<sub>4</sub>H<sub>11</sub>NO)Ni(CN)<sub>4</sub>]·8H<sub>2</sub>O (C1):** At room temperature, NiCl<sub>2</sub>·6H<sub>2</sub>O (0.30 g; 1.41 mmol) was dissolved in 20 cm<sup>3</sup> of water-alcohol solution (1:1; v/v) and KCN (0.19 g; 1.45 mmol) was added to this solution, forming a cloudy light blue one. The second metal salt (NiCl<sub>2</sub>·6H<sub>2</sub>O, 0.35 g; 1.47 mmol) was added to this solution to form a light blue solution. Abut ligand (0.22 g; 1.48 mmol), which was prepared by dissolving in 15 cm<sup>3</sup> of ethanol, was added to the resulting solution to give a clear lilac solution. The result solution was transferred to microwave synthesizer (MW) and the reaction medium was applied at 100°C, 500W and 25 min synthesis program. The cloudy light green solution was filtered after MW synthesis program and allowed to stand at room temperature and undergo slow evaporation whereupon powder product was obtained.

The procedure for the synthesis of **C1** was similar to that of **C2**, **C3** and **C4** complexes using CuCl<sub>2</sub>·2H<sub>2</sub>O (0.18 g; 1.06 mmol), ZnCl<sub>2</sub> (0.35 g; 2.56 mmol) and CdSO<sub>4</sub>·8/3H<sub>2</sub>O (0.45 g; 1.7 mmol) instead of second metal salt NiCl<sub>2</sub>·6H<sub>2</sub>O, respectively.

**[Ni(C<sub>4</sub>H<sub>11</sub>NO)Ni(CN)<sub>4</sub>]·8H<sub>2</sub>O (C1):** calcd. %: C 21.13; H 5.98; N 15.40 %; found %: C 21.22; H 5.01; N 15.11 %. MW:454.71g/mol. Colour: Cream. Yield: 61%. IR spectrum (cm<sup>-1</sup>): 3601ν<sub>(O-H)</sub>, 3337, 3270 ν<sub>(N-H)</sub>, 2964, 2932, 2881 ν<sub>(C-H)</sub>, 2168, 2129ν<sub>(C≡N)</sub>, 1615 δ<sub>(N-H)</sub>, 1460 δ<sub>(CH<sub>2</sub>)</sub>, 1114 ν<sub>(C-N)</sub>, 1031 ν<sub>(C-O)</sub>.

**[Cu(C<sub>4</sub>H<sub>11</sub>NO)<sub>2</sub>Ni(CN)<sub>4</sub>]·7H<sub>2</sub>O (C2):** calcd. %: C 27.16; H 6.84; N 15.84 %; found %: C 27.39; H 5.99; N 15.94 %. MW:530.69 g/mol. Colour: Navy blue. Yield: 32%. IR spectrum (cm<sup>-1</sup>):3600 ν<sub>(O-H)</sub>, 3314, 3123 ν<sub>(N-H)</sub>, 2968, 2920, 2877, 2853 ν<sub>(C-H)</sub>, 2170, 2115 ν<sub>(C≡N)</sub>, 1615, 1591 δ<sub>(N-H)</sub>, 1460 δ<sub>(CH<sub>2</sub>)</sub>, 1118 ν<sub>(C-N)</sub>, 1055 ν<sub>(C-O)</sub>.

**[Zn(C<sub>4</sub>H<sub>11</sub>NO)Ni(CN)<sub>4</sub>]·8H<sub>2</sub>O (C3):** calcd. %: C 20.82; H 5.90; N 15.18 %; found %: C 19.80; H 4.82; N 15.99 %. MW:461.43 g/mol. Colour: Blue. Yield: 59%. IR spectrum (cm<sup>-1</sup>): 3596ν<sub>(O-H)</sub>, 33213250 ν<sub>(N-H)</sub>, 2964, 2936, 2877 ν<sub>(C-H)</sub>, 2190, 2130 ν<sub>(C≡N)</sub>, 1511 δ<sub>(N-H)</sub>, 1456 δ<sub>(CH<sub>2</sub>)</sub>, 1118 ν<sub>(C-N)</sub>, 1031 ν<sub>(C-O)</sub>.

**[Cd(C<sub>4</sub>H<sub>11</sub>NO)Ni(CN)<sub>4</sub>]·7H<sub>2</sub>O (C4):** calcd. %: C 19.59; H 5.14; N 14.28 %; found %: C 18.96; H 5.02; N 14.88 %. MW:491.01 g/mol. Colour: Blue. Yield: 61%. IR spectrum (cm<sup>-1</sup>): 3603 ν<sub>(O-H)</sub>, 3373, 3155 ν<sub>(N-H)</sub>, 2928, 2881, 2849ν<sub>(C-H)</sub>, 2170 ν<sub>(C≡N)</sub>, 1621 δ<sub>(N-H)</sub>, 1456 δ<sub>(CH<sub>2</sub>)</sub>, 1114 ν<sub>(C-N)</sub>, 1012 ν<sub>(C-O)</sub>.

**Antibacterial activity.** To determine antibacterial characteristic of complexes (**C1**, **C2**, **C3** and **C4**), nine Gram (-) (*E. aerogenes* ATCC 13048, *S. infantis*, *K. pneumoniae*, *P. aeruginosa* DSMZ 50071, *S. kentucky*, *S. enteritidis* ATCC 13075, *S. typhimurium*, *E. coli* ATCC 25922, *S. marrescens* ATCC 13048) six Gram (+) (*E. faecalis* ATCC 29212, *L. innocua*, *E. durans*, *S. aureus* ATCC 25923, *S. epidermidis* DSMZ 20044, *B. subtilis* DSMZ 1971) were analyzed. All bacteria strains were supplied by Department of Molecular Biology and Genetic at Bartın University (Bartın, Turkey). The minimum inhibition concentration (MIC) of the complex were identified based on method applied by Andrews [34]. All bacteria were inoculated into Luria Bertani broth (LB) and bacterial growth were standardized to 0.5 Mc Farland turbidity (1.5×10<sup>8</sup> CFU/ml). The concentration of complexes was serially diluted, and the mixture was incubated at 37°C during overnight. By UV-visible spectrum at 600 nm, MIC value was

designated regarding bacterial growth turbidity. The suspected wells were inoculated into petri dishes and were incubated at 37°C. The minimum bactericidal concentration (MBC) was detected the lowest concentration that killed microorganisms.

**Antibiofilm activity.** Antibiofilm activity against six bacteria (*E. coli* ATCC 25922, *E. aerogenes* ATCC 13048, *S. infantis*, *B. subtilis* DSMZ 1971, *S. aureus* ATCC 25923, *S. epidermidis* DSMZ 20044) was examined by using complexes (**C1**, **C2**, **C3** and **C4**). The mixture complexes and bacteria culture and were incubated within a 96-well microplate at 37°C for 48 h. After incubation, all wells were rinsed and desiccated at room temperature. Methanol (95%) was added for fixation for about fifteen minutes and all wells were stained with 0.1% crystal violet. Gram (+) and (-) respectively were treated 33% glacial acetic acid and 95% ethyl alcohol and then the wells were measured by UV spectrophotometer. Eventually, the percentage of antibiofilm activity was calculated by formula:

$$\% \text{Inhibition} = (\text{Control OD}_{600} - \text{Sample OD}_{600}) / \text{Control OD}_{600} \times 100 \text{ [35].}$$

**Anticancer studies.** The anti-cancer activity of complexes (**C1**, **C2**, **C3** and **C4**) was tested using human cervical cancer cell line HeLa obtained from the American Type Culture Collection (ATCC). Cells were cultured in DMEM containing 10% bovine fetal serum (FBS-12A, Capricorn, South America), 1% penicillin/ streptomycin and 1% glutamine (15140148, Thermo) in a 75 cm<sup>2</sup> culture flask at 5% CO<sub>2</sub> in incubator at 37°C. The cytotoxicity of Ag-NWs was evaluated by MTT assay using HeLa cell lines. At 5 mg/mL concentration, MTT (3-(4,5-dimethylthiazol-2-yl)-2,5-diphenyltetrazolium bromide) was prepared. 100 µL of MTT was added and incubated for 4 h. The crystals of purple colored formazone were formed and then dissolved with 200 µL of di-methyl sulphoxide (DMSO). Complexes (**C1**, **C2**, **C3** and **C4**) (0.5 to 64 µg/ml) including DMEM solution were added into 96-well plates and 2×10<sup>4</sup> cells and were inoculated into wells. MTT assays were applied and OD values were measured using a microplate reader (SPECTROstar Nano, BMG) at 570 nm after 24 h after incubation.

## Results and Discussion

**IR spectra.** The characteristic peaks of Abut ligand were observed with the shifts in their positions in the IR spectra of complexes **C1-C4** (Fig. 1). The stretching vibrations of the OH-groups in alcohol-

derived compounds were observed in the IR spectrum in the range of 3700-3500 cm<sup>-1</sup> [36, 37]. Therefore, the stretching vibration of OH-group of **C1**, **C2**, **C3** and **C4** was observed at 3601, 3600, 3596 and 3603 cm<sup>-1</sup>, respectively. The NH<sub>2</sub> group of the abut ligand was observed in the range of 3155-3373 cm<sup>-1</sup> with wide weak interpreted as the result of overlapping of the OH-groups of lattice water in their structures. The CH<sub>2</sub> groups of the ligand that has different sphere were observed as a multi peaks in the range of 2849-2968 cm<sup>-1</sup>.

The most characteristic and distinctive peak in cyanido complexes was the sharp cyanido peak observed in the 2200-2000 cm<sup>-1</sup> range. The M-C≡N-M' bridged type cyanido group stretching vibration shifts to a higher frequency than the end cyanido group. Therefore, the presence of both bridges and end cyanido groups in the structure could be observed by splitting of cyanido peaks [38]. Based on this information, the peaks observed at 2168, 2129 cm<sup>-1</sup> (**C1**), 2170, 2115 cm<sup>-1</sup> (**C2**) and 2190, 2130 cm<sup>-1</sup> (**C3**) indicated the presence of bridged and end cyanido groups in the structures. The single peak observed at 2170 cm<sup>-1</sup> (**C4**) was attributed to all cyanido groups bridged in the complex structure or end cyanido groups involved in weak HB interactions. In the fingerprint region of the spectrum the δ<sub>(N-H)</sub>, δ<sub>(CH<sub>2</sub>)</sub>, ν<sub>(C-N)</sub> and ν<sub>(C-O)</sub> were observed in the range of 1621-1012 cm<sup>-1</sup>. In addition, it was estimated that the small intense peaks in the 600-400 cm<sup>-1</sup> range were due to the bonds of Ni-C, Ni-N and Ni-O.

**Thermal analyses.** To assess the thermal stability of **C1-C4**, thermal gravimetric experiments were carried out. The TGA results indicate that the **C1-C4** complexes were stable at room temperature. The first weight loss of 30.74% (calc: 31.66%) in the range 89-250°C was corresponded to the removal of eight water molecule (Fig. 2). The second weight loss in the range of 250-346°C was shown a loss of 20.86%, indicating the removal of neutral Abut ligand (calc: 19.60%). The continuous and last weight loss in the range 480-840°C was shown a mass loss of 22.07%, indicating the removal of four cyanido ligands (calc: 22.87%), which was followed by decomposition of the framework. The remaining weight percentage of 26.33% (calc: 25.87%) at 840°C is in agreement with the theoretical value of a mixture of Ni+Ni. Thermal decomposition of **C2** consists of three degradation stages from 80°C to 211°C, 211°C to 410°C and 410°C to 865°C corresponding to release of seven water molecule, two Abut ligand and four cyanido,



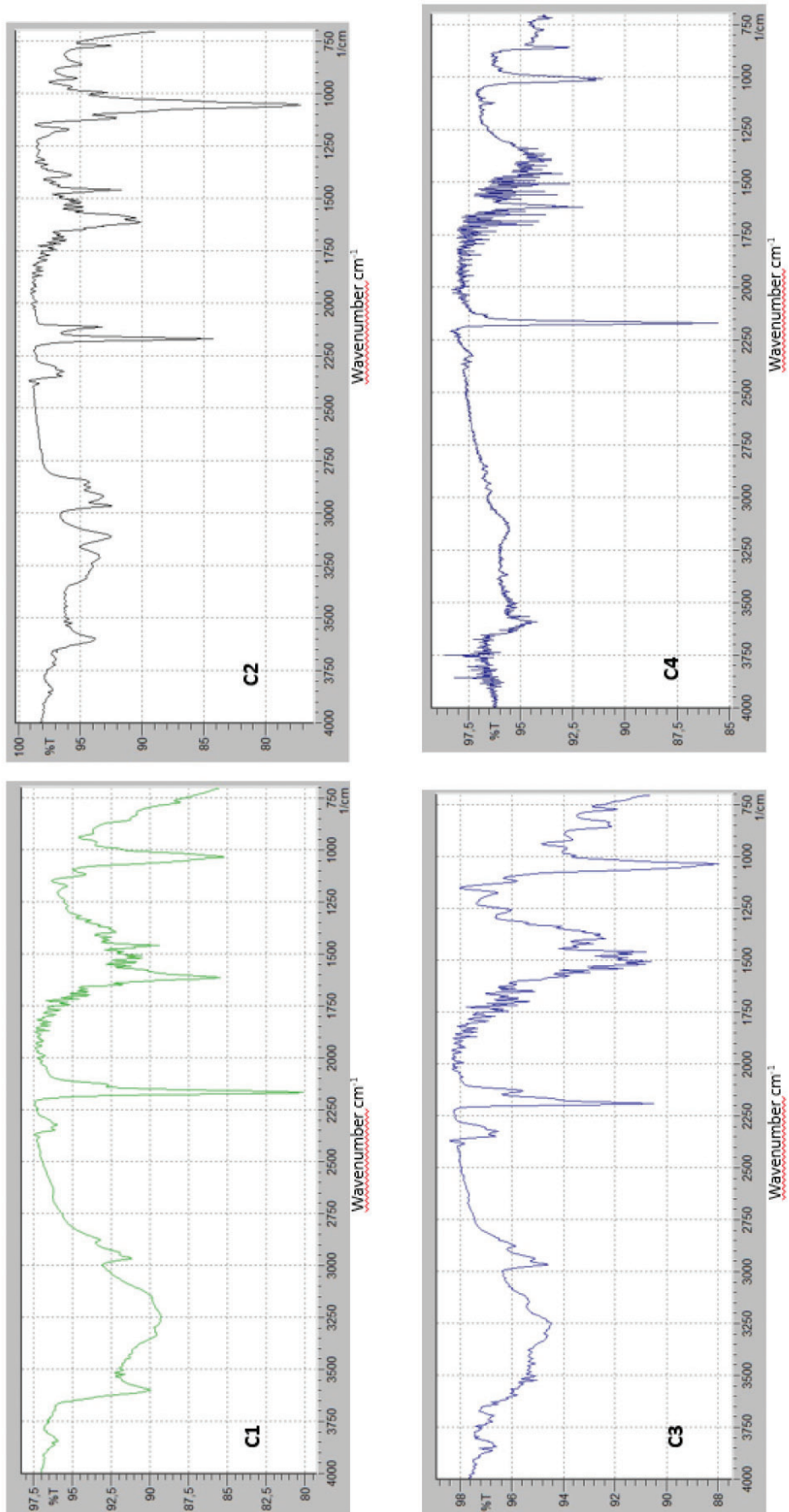


Fig. 1. IR spectra of C1-C4

respectively. The remaining mass loss of **C2** was pointed to Ni+Cu as end product (found: 24.09%; calc: 23.08%). The **C3** and **C4** decomposed similarly at 75-840°C (**C3**) and 80-190°C (**C4**) in two stages with *Abut* and four cyanido groups were liberated at 410-840°C for **C3** and 190-840 for **C4**. Thermal

analyses data for **C1-C4** complexes were given in Table 1. The TG and DTA curves of **C1-C4** complexes were shown in Fig. 2.

*Antibacterial activity.* Almost all complex synthesis whose antibacterial activities were studied indicated an antibacterial effect. However, no bacte-

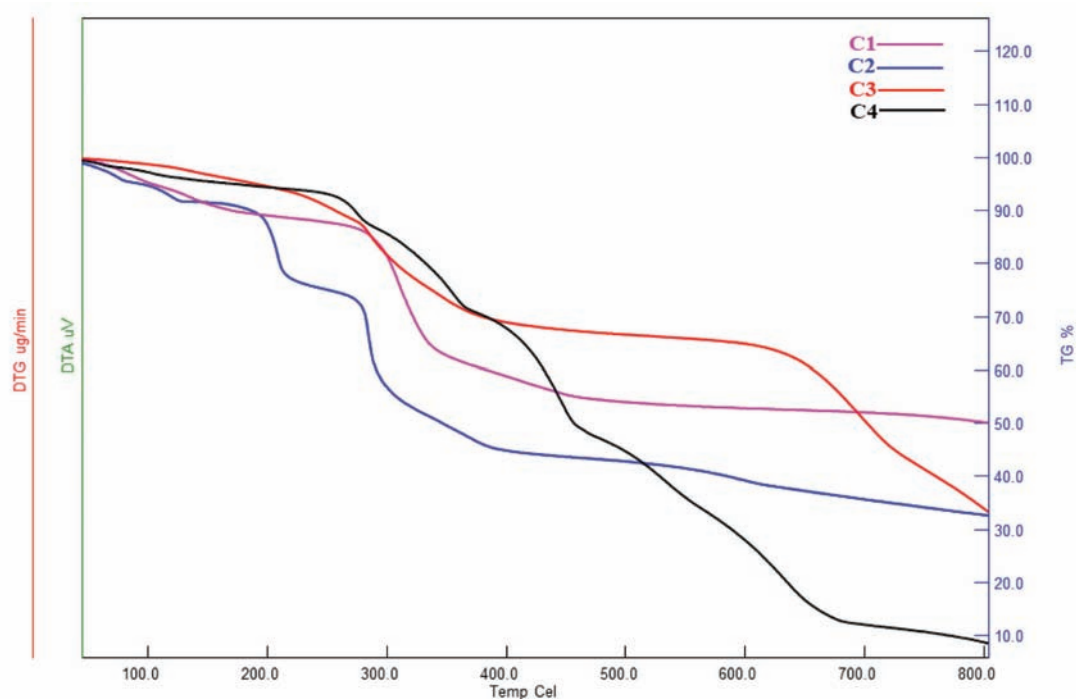


Fig. 2. The TG curves of **C1-C4**

Table 1. The thermal analysis data of **C1-C4** complexes

Complex	Step	Temperature range (°C)	Mass loss, $\Delta m$ (%)		Total mass loss, $\Delta m$ (%)		Leaving group
			Exp.	Calc.	Exp.	Calc.	
[Ni( <i>Abut</i> )Ni(CN) <sub>4</sub> ] $\cdot$ 8H <sub>2</sub> O MA:454.71 C <sub>8</sub> H <sub>27</sub> N <sub>5</sub> O <sub>9</sub> Ni <sub>2</sub> ( <b>C1</b> )	1	89-250	30.74	31.66	30.74	31.66	8H <sub>2</sub> O
	2	250-346	20.86	19.60	51.60	51.26	<i>Abut</i>
	3	480-840	22.07	22.87	73.67	74.13	4CN
[Cu( <i>Abut</i> ) <sub>2</sub> Ni(CN) <sub>4</sub> ] $\cdot$ 7H <sub>2</sub> O MA:530.69 C <sub>12</sub> H <sub>36</sub> N <sub>6</sub> O <sub>9</sub> NiCu ( <b>C2</b> )	1	80-211	23.01	23.74	23.01	23.74	7H <sub>2</sub> O
	2	211-410	32.89	33.59	55.90	57.33	2 <i>Abut</i>
	3	410-865	20.01	19.59	75.91	76.92	4CN
[Zn( <i>Abut</i> )Ni(CN) <sub>4</sub> ] $\cdot$ 8H <sub>2</sub> O MA:461.40 C <sub>8</sub> H <sub>27</sub> N <sub>5</sub> O <sub>9</sub> NiZn ( <b>C3</b> )	1	75-410	31.00	31.20	31.00	31.20	8H <sub>2</sub> O
	2	410-840	40.71	41.85	50.56	50.51	<i>Abut</i> +4CN
[Cd( <i>Abut</i> )Ni(CN) <sub>4</sub> ] $\cdot$ 7H <sub>2</sub> O MA:491.01 C <sub>8</sub> H <sub>25</sub> N <sub>5</sub> O <sub>8</sub> NiCd ( <b>C4</b> )	1	80-190	24.91	25.66	24.91	25.66	7H <sub>2</sub> O
	2	190-840	38.75	39.81	64.72	64.41	<i>Abut</i> +4CN

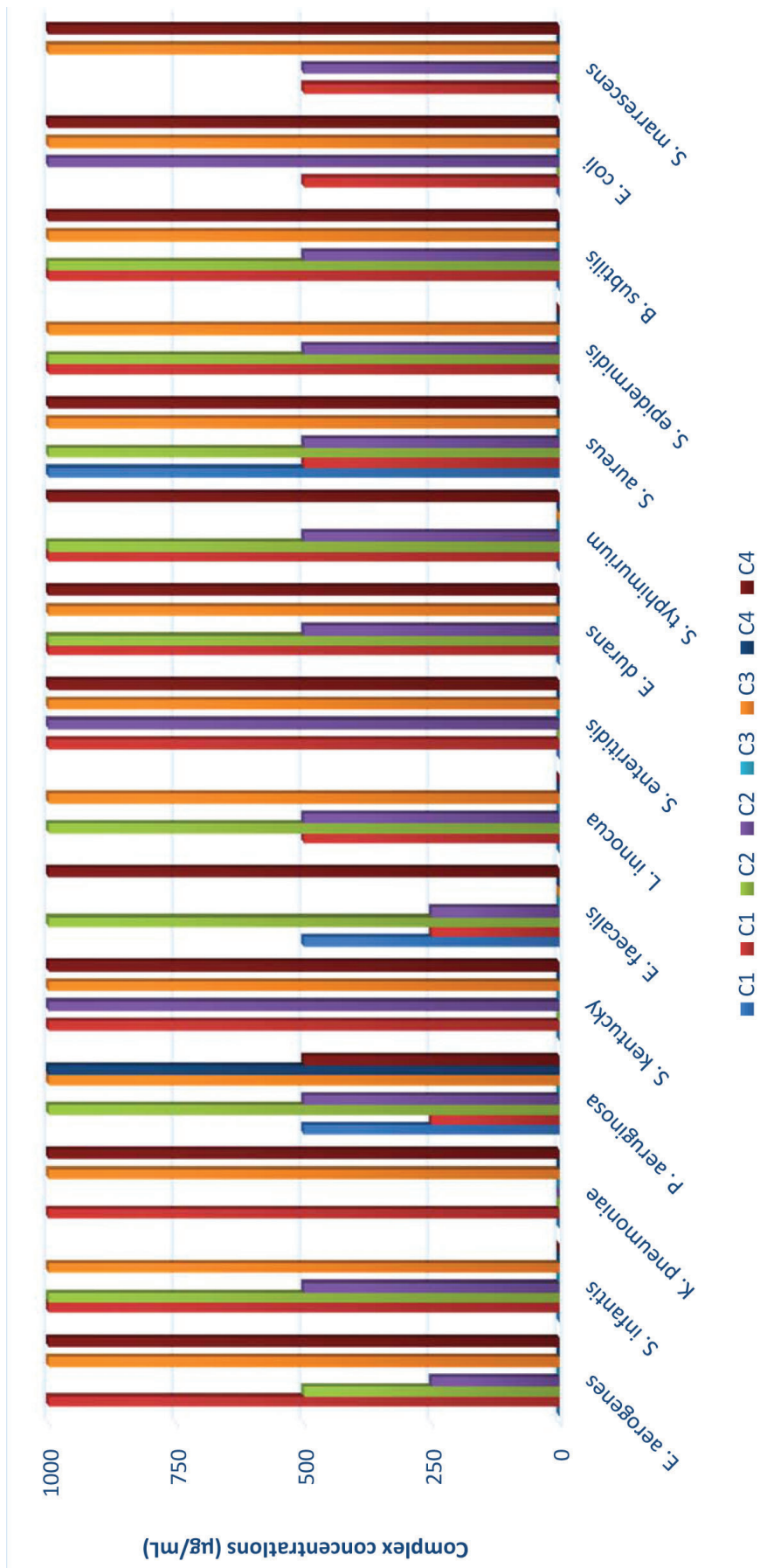


Fig. 3. Antibacterial activity of C1-C4 complexes concentrations (mg/ml)

ricidal effect was observed against *K. pneumoniae*, *S. kentucky*, *S. enteritidis*, *E. coli* and *S. marrescens*. According to the results (Fig. 3), it was observed that **C2** complex both inhibits bacterial growth and showed a bactericidal effect even at low concentrations. Metal ions act as chelators and due to some important factors, such as ion coordination area, hydrophilicity and binuclear structure, bimetallic complexes have critical effects on the antibacterial activity on bacteria. Metal ions bind to enzymes, proteins, and DNA, and investigate antibacterial properties on bacteria [39]. The presence of metal ions also changes the biological activity because of changing the solubility and conductivity of substances. In addition, metal ions affect the permeability

of cell membranes. By connecting metal ions to cell membranes, they can damage bacterial membranes or by binding to lipids on the membrane, increasing membrane permeability. As a result, it alters the amount of complex-synthesis up taken into the cell, leading to increased intracellular toxicity and bacterial death or inhibition of bacterial growth [12, 40-42]. (**C1**) analyzed the antibacterial effects of their complex-synthesis and used *B. subtilis*, *S. aureus*, *E. coli*, *E. aerogenes*, *P. aureginosa* and *S. enteritidis* bacteria strains was used in present study. Compared to the complex syntheses in this study with **C1** and **C4** complex-synthesis containing Ni-CN bridge synthesized by both, they were shown antibacterial effect against tested bacteria.

Table 2. Inhibition percentages of the formation of complex-syntheses biofilm layer

Con., µg/ml	Bacteria						
	<i>E. coli</i> ATCC 25922	<i>E. aerogenes</i> ATCC 13048	<i>S. infantis</i>	<i>B. subtilis</i> DSMZ 1971	<i>S. aureus</i> ATCC 25923	<i>S. epidermidis</i> DSMZ 20044	
<b>C1</b>	1000	57	55	70	53	67	42
	500	11	17	1	0	47	3
	250	0	0	0	0	0	8
	125	0	0	5	0	0	0
	62.5	0	4	4	0	0	9
	31.25	0	3	0	0	0	0
<b>C2</b>	1000	58	55	66	32	63	22
	500	29	3	1	7	57	0
	250	7,5	0	0	0	0	8
	125	7	0	0	0	3	0
	62.5	4	0	1	6	1	0
	31.25	5	0	2	6	0	0
<b>C3</b>	1000	44.3	46.8	71.2	57	57.5	28.8
	500	40.1	17.1	60.8	38.2	36	5
	250	8.7	0	1	0	18.4	0
	125	6.5	6.9	3.1	0	15.5	7.5
	62.5	8.7	0	10.4	17.4	20.8	0
	31.25	6	3.1	13.2	28.5	16.8	0
<b>C4</b>	1000	24.3	26.6	30.8	13.3	53.2	0
	500	0	0	0	0	0	0
	250	1.3	0	0	0	0	0
	125	1.9	8.2	0	0	8.3	0
	62.5	8.4	0	0.5	7	6.9	0
	31.25	4.6	5.4	8.2	0	6.5	0

**Antibiofilm activity.** The biofilm layer is a collection of cells that are attached to a living or inanimate surface of bacteria, embedded in the extracellular and self-produced polymer matrix. Biofilm layer can occur on different surfaces such as glass, plastic, wood, metal, soil, medical implants, food product and tissue [43, 44]. The most efficient biofilm inhibition activity for all complexes was observed at  $1000 \mu\text{g}\cdot\text{ml}^{-1}$ , the highest concentration. Considering the results obtained in the study, it was noticed that the highest percentage of biofilm inhibition was obtained by **C3** complex-synthesis with a concentration of  $1000 \mu\text{g}\cdot\text{ml}^{-1}$ , with a 71.2% against *S. infantis*. Only **C4** complex synthesis did not act an antibiofilm effect against the *S. epidermis* strain at all concentrations. The cell membranes of Gram (-) bacteria contain a thinner polysaccharide layer, and the polysaccharide layer of Gram (+) bacteria is thicker and contains teichoic acid. According to the results, it was determined that the percentage of Gram (-) bacteria biofilm inhibition was higher than other.

**Anticancer activity.** **C1-C4** complexes were investigated for anticancer activity on HeLa cell lines. After 24 h incubation with different doses of prepared complexes **C1-C4** (0.5, 1, 2, 4, 8, 16, 32, 64  $\mu\text{g}/\text{ml}$ ), they showed cytotoxic activity at high doses HeLa cell lines. The increasing concentration of complexes **C1-C4** decreased cell viability, as expected. Therefore, as shown in Figure 4 the anticancer activity of the complexes **C1-C4** was dose-dependent in confluent cultures, and the  $\text{IC}_{50}$  values were 14.856  $\mu\text{g}/\text{ml}$ , 6.527  $\mu\text{g}/\text{ml}$ , 7.218  $\mu\text{g}/\text{ml}$  and 19.250  $\mu\text{g}/\text{ml}$  for HeLa cell line (Fig. 4).

**Conclusions.** In this paper, we synthesized four cyanido complexes by using tetracyanonickelate anion and Abut ligand. All of them were in powder form and characterized by IR spectroscopy, elemental, and thermal analysis. The characteristic peak of cyanido ligand was seen in the  $2115\text{-}2190 \text{ cm}^{-1}$  range as expected. As a general trend from the thermal analysis curves of complexes, first hydrate water, then neutral Abut ligand and finally anionic cyanido ligands were away from the structure. At the end of

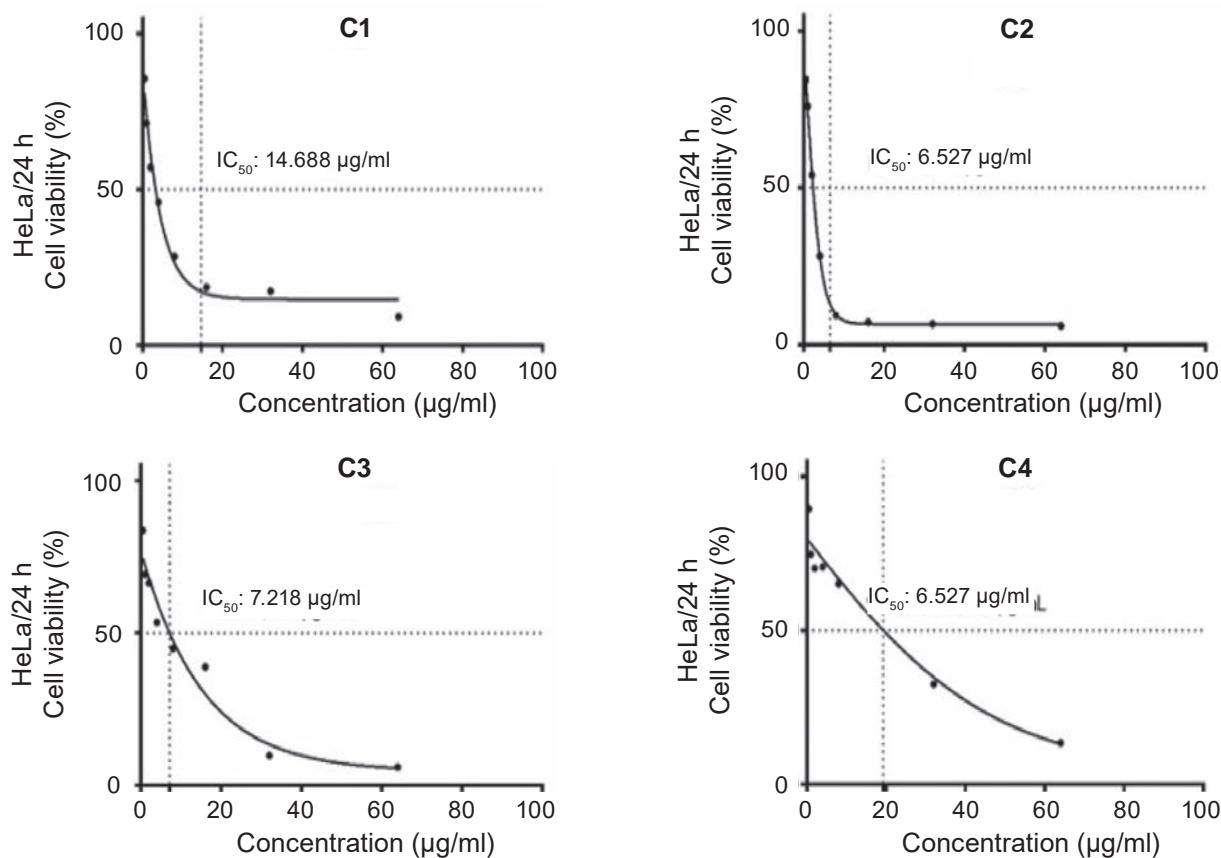


Fig. 4. Anti-cancer assay of complexes (**C1-C4**)



thermal decomposition, Ni+Ni, Ni+Cu, Ni+Zn and Ni+Cd metal mixtures were obtained. The complex syntheses analyzed in current study were found to demonstrate antibacterial properties at low concentrations. In literature, cyanido-bridged bimetallic complex-syntheses have been studied in many areas such as antibacterial, antifungal, anticancer. The type of biofilm inhibition test applied in this study had not been performed in the earlier studies and it was observed that the results provided inhibition of biofilm layer at very low concentrations. The cyanido complexes **C1-C4** had anticancer activity against service cancer cell line HeLa. These cyanido complexes showed activity in the concentrations of 0.5 to 64 µg/ml.

**Conflict of interest.** Authors have completed the Unified Conflicts of Interest form at [http://ukr-biochemjournal.org/wp-content/uploads/2018/12/coi\\_disclosure.pdf](http://ukr-biochemjournal.org/wp-content/uploads/2018/12/coi_disclosure.pdf) and declare no conflict of interest.

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## НОВІ ДВОЯДЕРНІ КОМПЛЕКСИ ЦІАНІДІВ ІЗ ГИДРОКСАМІНОВИМ ЛІГАНДОМ: СИНТЕЗ, ХАРАКТЕРИСТИКА ТА МОЖЛИВОСТІ ЗАСТОСУВАННЯ

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Метою цієї роботи було синтезувати комплекси ціанідів [Ni(*Abut*)Ni(CN)<sub>4</sub>]·8H<sub>2</sub>O (**C1**), [Cu(*Abut*)<sub>2</sub>Ni(CN)<sub>4</sub>]·7H<sub>2</sub>O (**C2**), [Zn(*Abut*)Ni(CN)<sub>4</sub>]·8H<sub>2</sub>O (**C3**) та [Cd(*Abut*)Ni(CN)<sub>4</sub>]·7H<sub>2</sub>O (**C4**) та дослідити їх антибактеріальну та цитотоксичну дію. Одержані комплекси вивчали методом мікрохвильового синтезу з використанням

ІЧ-Фур'є спектроскопії та термічного аналізу. Досліджено антибактеріальну та цитотоксичну активність синтезованих комплексів. Встановлено, що розщеплення смуг коливання ціанідогруп в ІЧ-Фур'є спектрах **C1-C4** вказує на наявність кінцевих ціаногруп. Антибактеріальну активність **C1-C4** вивчали на дев'яти видах грам-негативних та шести грам-позитивних бактеріях. Найбільшу антибактеріальну активність комплексів спостерігали за концентрації 1000 мкг/мл. Цитотоксичну активність оцінювали з використанням клітинної лінії HeLa та МТТ-тесту. Виявлено, що досліджувані ціанідні комплекси знижують життєздатність клітин HeLa зі значеннями IC<sub>50</sub> 14,86; 6,5; 7,2 та 19,2 мкг/мл для комплексів **C1**, **C2**, **C3** та **C4**, відповідно.

**Ключові слова:** комплекси ціанідів, антибактеріальна, цитотоксична активність, клітини HeLa.

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