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Synthesis, Spectroscopic, and Biological Assessments on Some New Rare Earth Metal Adrenaline Adducts

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Abstract: Adrenaline (Adr) reacts with chlorides of Y$^{3+}$, Ce$^{3+}$, Nd$^{3+}$ and Sm$^{3+}$ in methanol at 60 °C to yield metal ionic adducts of definite composition. These compounds are characterized by elemental analyses, molar conductivity, UV-Vis., 1H-NMR, Raman laser, scanning electron microscopy, energy-dispersive X-ray spectroscopy (EDX), and mid infrared spectral measurement investigations. The new compounds were tested against five gram-positive and two gram-negative bacteria, groups of the catechol moiety are linked to central metal ions based on the infrared and Raman laser spectra. The new compounds were tested against five gram-positive and two gram-negative bacteria, in addition to two Aspergillus strains. Metal adducts were shown to have stronger antibacterial and antifungal properties than free adrenaline compounds.

Keywords: adrenaline; rare earth metals; complexes; mid infrared; Raman laser; biological activity

1. Introduction

In addition to being a neurotransmitter, adrenaline (C$_9$H$_{13}$NO$_3$) is a hormone [1]. An increase in sympathetic nervous system activity is associated with increased heart rate, blood vessel constriction, and airway dilation [2]. Adrenaline was the first hormone discovered, and it was manufactured effectively in 1904. It belongs to the biogenic amines family, which contains serotonin and histamine. It belongs to the catecholamine family of compounds, which also contains norepinephrine and dopamine. Chronic stress is indicated by persistently elevated levels of catecholamines in the blood. The adrenal gland in the body of many animals produces the hormone adrenaline. It raises the heart rate, dilates blood vessels and air channels, and has a number of other modest effects when created in the body.

Adrenaline is a powerful vasoconstrictor and stimulant of the heart [3]. Adrenaline is a catecholamine that, like other catecholamines like dopamine and noradrenaline, plays a vital role in physiology as a neurotransmitter in the central nervous system. This makes adrenaline crucial for keeping homeostasis in the body and responding to acute and chronic stress by orchestrating cardiovascular, metabolic, and visceral functions [3]. Although...
adrenaline makes up just 5% to 10% of the total catecholamines in the central nervous system, it is thought to play a role in the central control of blood pressure [4], respiration [5], and pituitary hormone production [5]. In addition, adrenaline is the most common treatment for anaphylactic shock, and it should be given as soon as a person begins to have severe allergic responses.

The literature on adrenaline and its derivative adducts is extensive [6–15]. The complexes of the first-row transition-metal adrenaline complexes were formed, and the stabilities were discussed [6]. The chemical structures of the complexes of adrenaline and related compounds with Ni$^{2+}$, Cu$^{2+}$, Zn$^{2+}$, Cd$^{2+}$ and Pb$^{2+}$ were studied using spectroscopic and physicochemical analyses [7]. The magnesium, adenosine triphosphate, and catecholamine complexes have been synthesized and the application of new means of identifying solute-solute interactions of biological significance were achieved using NMR chemical shift data [8]. The role of metals in the enzymatic and nonenzymatic oxidation of epinephrine as well as the physics and chemistry of rare earths elements have been investigated [9,10]. The biological activity screening of metal copper(II) and nickel(II) tungstate complexes of the first-row transition-metal adrenaline complexes were formed, and the stability and luminescence properties of Tb(III) complexes with adrenaline [13] were assigned. A thermoanalytical study of unusual adrenaline coordination compounds with the divalent transition-metal ions Co(II), Ni(II) and Cu(II) was reported [14,15].

However, only a small amount of spectroscopic, mid infrared and Raman laser spectra data are provided, and without biotic evaluation directly connected to the organization of rare earth ions, as described in the literature. Adrenaline compounds containing rare earth metal ions, such as Y$^{3+}$, Ce$^{3+}$, Nd$^{3+}$ and Sm$^{3+}$, might be excellent models for learning about catecholamine adducting behavior and discovering antimicrobial characteristics of these compounds in solid form.

We evaluate the UV spectroscopic and biotic properties of adrenaline precipitated from CH$_3$OH solutions utilizing metal chlorides and adrenaline. With succeeding binding moiety particles being coupled to Y$^{3+}$ adrenaline adduct (1:1) and Ce$^{3+}$, Nd$^{3+}$, and Sm$^{3+}$ metal ions (1:2), the resulting compounds displayed molar proportions of 1:1 and 1:2. The characteristics of the obtained adducts were discovered by scanning electron microscopy, energy-dispersive X-ray spectroscopy (EDX), UV-Vis., $^1$H–NMR, Raman laser, conductivity measurements and mid infrared.

2. Experimental

2.1. Materials and Synthesis of Adrenaline Adduct

The compounds YCl$_3$·6H$_2$O, CeCl$_3$·7H$_2$O, NdCl$_3$·6H$_2$O, SmCl$_3$·6H$_2$O, and CH$_3$OH were bought from the Aldrich company (Laramie, WY, USA). Adrenaline was purchased from the Aldrich Company (Buchs, Switzerland). All compounds utilized in this investigation were pure and no more purification was undertaken. A 1:1 (M$^{3+}$)/Adr ratio was used to precipitate all adducts.

The [Y$_2$(Adr)$_2$(H$_2$O)$_6$]Cl$_3$·8H$_2$O, [Ce(Adr)$_2$(H$_2$O)$_2$]Cl$_3$·10H$_2$O, [Nd(Adr)$_2$(H$_2$O)$_2$]Cl$_3$·6H$_2$O and [Sm(Adr)$_2$(H$_2$O)$_2$]Cl$_3$·12H$_2$O adducts were obtained by heating (60 °C) and stirring a 25 mL 99.9% CH$_3$OH solution of YCl$_3$·6H$_2$O (0.303 g; 1 mmol), CeCl$_3$·7H$_2$O (0.373 g; 1 mmol), NdCl$_3$·6H$_2$O (0.359 g; 1 mmol), or SmCl$_3$·6H$_2$O (0.365 g; 1 mmol) rare earth metal chlorides to the added solution of adrenaline (0.183 g; 1 mmol) in 25 mL 99.9% CH$_3$OH. These mixtures were refluxed for 3 h, after which the produced white solid was washed with 99.9% CH$_3$OH, then dried in a vacuum over anhydrous CaCl$_2$.

2.2. Measurements

The atom investigations of C and H components were achieved via the microanalysis unit at Cairo University, Egypt, using a Perkin Elmer CHN 2400 instrument. The soluble adrenaline adducts were measured in freshly produced 1.0 × 10$^{-2}$ g/5 cm$^3$ dimethyl formamide (DMF) solutions using a Jenway 4510 conductivity meter (Jenway company,
The 4510 conductivity meter supplied with a glass conductivity probe with ATC ($K = 1$) (027 013), electrode holder and UK power supply, at a reference temperature of 25 °C and with the calibration at cell constant or standard solutions. On a Bruker FT-IR Spectrophotometer (4000–400 cm$^{-1}$), infrared spectra with KBr discs were recorded, and Raman laser spectra of materials were analyzed on a Bruker FT-Raman (Bruker company, Billerica, MA, USA), with laser of 50 mW. The $^1$H-NMR spectra were documented on Varian Mercury VX-300 NMR spectrometer (Bruker company, Billerica, MA, USA). The $^1$H spectra were performed by 300 MHz spectra in deuterated dimethylsulphoxide (DMSO-d$_6$). Chemical shifts are mentioned in δ and were correlated to that of the solvents. An UV2Unicam UV/Vis Spectrophotometer coupled with a quartz cell of 1.0 cm path length was used to measure the spectra of electron absorption of epinephrineadducts in DMSO solvent throughout the 900–200 nm region. Scanning electron microscopy (SEM) images and Energy Dispersive X-ray Detection (EDX) were recorded in Joel JSM-6390 equipment (Jeol Company, Peabody, MA, USA), with an accelerating voltage of 20 KV.

2.3. Antimicrobial Properties

A modified Kirby-Bauer disc diffusion technique [16] was used to evaluate the antimicrobial activity of the tested materials. In a nutshell, 100 µL of the finest bacteria/fungus were cultured in 10 mL of a new medium till a cell count of around 10^8 cells/mL for bacteria and 10^5 cells/mL for fungi [17]. Of this, 100 µL of the microbial suspension were distributed on agar plates that corresponded to the broth in which they were kept. Isolated colonies of each organism that might be detrimental were chosen from the main agar plates and their vulnerability was evaluated using the disc dispersion technique [18–28]. There are many research papers that deal with the disc dispersion technique in conducting the biological evaluation of the prepared chemical compounds, and the findings of these can be listed as follows: the use of Ruthenium complexes with 2-pyridin-2-yl-1H-benzimidazole as potential antimicrobial agents has been discussed according to the correlation between chemical properties and anti-biofilm effects [18]. The possibility of redesigning the nature of ruthenium flavonoid complexes with antitumor, antimicrobial and cardioprotective activities was studied [19]. The synthesis, spectral characterization and biological activities of Co(II) and Ni(II) mixed ligand complexes were investigated [20]. The antimicrobial efficiency against planktonic and adherent microbes and in vitro cytotoxicity features of the new cobalt (II) complexes with imidazole derivatives were screened [21]. Providing insight into their analytical applications and biological activity, the quinolone complexes with lanthanide ions [22], Cu(II), Zn(II) complexes with theophylline [23], Mn(II), Co(II), Ni(II), Cr(III) and Fe(III) melatonin drug complexes [24] were discussed. The antimicrobial and antifungal screening of the 3-benzenedicarbonyl-bis-(amino acid) and dipeptide candidates [25], as well as the pentafluorophenyl platinum(II) complexes of N-allyl and N-benzyl derivatives, were reported [26].

Mueller-Hinton agar is recommended by the National Committee for Clinical Laboratory Standards (NCCLS) because it has high batch-to-batch repeatability. The NCCLS [27–29] established an authorized standard technique M38-A for assessing the susceptibility of filament-like mold to antifungal drugs, which was used to evaluate the disc dispersion technique for filament-like fungi. The NCCLS [30] created a standard yeast diffusion technique (M44-P). Plates injected by filament-like fungi, Aspergillus niger, were maintained at an ambient temperature for two days; gram (+) bacteria, Bacillus subtilis, Streptococcus pneumonia, Staphylococcus aureus; gram (−) bacteria, Escherichia Coli, Pseudomonas aeruginosa, were kept at 35–37 °C for one to two days; similarly yeast Candida Albicans was kept at 30 °C for one to two days and the inhibition regions’ sizes were then evaluated in millimeters [16]. Positive controls for antimicrobial activity were Tetracycline (Antibacterial agent) and Amphotericin B (Antifungal agent), whereas a negative control was a filter disc coated with 10 µL of solvent distilled water, chloroform, DMSO.
Meuller-Hinton agar was utilized, which has been thoroughly evaluated for composition and pH. In the disc dispersion technique, the agar deepness in the plate is also a factor to consider. This approach is well-studied, and ordinary zones of inhibition for vulnerable values have been established. An amount of 10 µL of the measured concentration of the stock solutions were saturated onto blank paper discs Schleicher & Schuell (Sigma-Aldrich, Laramie, WY, USA), with a diameter of 8.0 mm. When a chemical is saturated, the filter paper disc is positioned on agar and the chemical disperses from the disc into the agar. As a result of this dispersion, the substance will only be available in the agar surrounding the disc. The substance’s solubility and size will define the substance’s penetration area surrounding the disc. If a microorganism is chemically sensitive, no growth will be observed surrounding the disc within the agar. A zone of inhibition or clear zone is the region around the disc where no development occurs. The zone diameters were evaluated using sliding calipers from the National for Clinical Laboratory Standards [27] for disc dispersion. Because agar-based techniques, such as Etest disc diffusion, are simpler and quicker than broth methods [31,32], they can be good alternatives.

3. Results and Discussion

The molar conductance measurements, elemental investigates and physical of Y3+, Ce3+, Nd3+, Sm3+, adrenaline adducts are provided in Table 1.

| Complexes | Mp, °C | M.Wt, g/mol | Color | Elemental Analysis (%) | Molar Conductance (Ω cm² mol⁻¹) | Yields, % |
|-----------|--------|-------------|-------|------------------------|-------------------------------|----------|
| [Y(Adr)₂(H₂O)₈]Cl₃·8H₂O | >250 | 938.579 | White | (23.01) 23.20 | 449 74 | |
| [Ce(Adr)₂(H₂O)₂]Cl₃·10H₂O | >250 | 828.887 | White | (26.06) 26.29 | 347 79 | |
| [Nd(Adr)₂(H₂O)₂]Cl₃·6H₂O | >250 | 761.007 | White | (28.38) 28.12 | 558 72 | |
| [Sm(Adr)₂(H₂O)₂]Cl₃·12H₂O | >250 | 875.127 | White | (24.68) 24.54 | 709 75 | |

The atomic investigation data of the produced adducts displaying (1:1) or (1:2) of the molar proportion (M: Adr), which are similar to the universal formulae [Y(Adr)₂(H₂O)₈]Cl₃·8H₂O, [Ce(Adr)₂(H₂O)₂]Cl₃·10H₂O, and [Nd(Adr)₂(H₂O)₂]Cl₃·6H₂O, [Sm(Adr)₂(H₂O)₂]Cl₃·12H₂O (Figure 1). The adrenaline compounds were produced at 60 °C, and the reactions may be described by the succeeding stoichiometric Equations (1)–(4):

\[
\text{YCl}_3 + \text{Adr} \rightarrow [\text{Y(Adr)}_2(\text{H}_2\text{O})_8]\text{Cl}_3\cdot8\text{H}_2\text{O} \quad (1)
\]

\[
\text{CeCl}_3 + \text{Adr} \rightarrow [\text{Ce(Adr)}_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot10\text{H}_2\text{O} \quad (2)
\]

\[
\text{NdCl}_3 + \text{Adr} \rightarrow [\text{Nd(Adr)}_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot6\text{H}_2\text{O} \quad (3)
\]

\[
\text{SmCl}_3 + \text{Adr} \rightarrow [\text{Sm(Adr)}_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot12\text{H}_2\text{O} \quad (4)
\]

The compounds are air-stable, moisture absorbing, have a high freezing-point, and are DMF and DMSO soluble. The 10–2 g/5 mL solutions of the synthesized compounds’ conductivities in DMF (Table 1) suggest that they are electrolytic with data disparity.
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Figure 1. (a): Suggested structures of [Y₂(Adr)₂(H₂O)₈]Cl₃·8H₂O adduct. (b): Suggested structures of [Ce(Adr)₂(H₂O)₂]Cl₃·10H₂O, [Nd(Adr)₂(H₂O)₂]Cl₃·6H₂O and [Sm(Adr)₂(H₂O)₂]Cl₃·12H₂O adducts.
3.1. Conductivity Measurements

In the DMF solvent (10–2 g/5 mL solution), the conductivity values for adrenaline compounds are 449, 347, 558, and 709 Ω\(^{-1}\) cm\(^2\) mol\(^{-1}\) for \([\text{Y}_2(\text{Adr})_2(\text{H}_2\text{O})_8]\text{Cl}_3\cdot8\text{H}_2\text{O}, [\text{Ce}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot10\text{H}_2\text{O, [Nd}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot6\text{H}_2\text{O, and [Sm}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot12\text{H}_2\text{O adducts.} Within the limitations of their solubility, conductive evaluations have regularly been utilized in the structure of metal adducts (coordination mode). They provide a way for determining an adduct’s ionization degree; the more liberation of ions in a molar adduct, the greater the molar conductivity of the adduct, and vice versa. The conductivity results clearly show that all the adducts are electrolytes. Furthermore, the molar conductance values show that all adrenaline adducts have anions present beyond the coordination sphere. The elemental analysis findings supported this conclusion. When the AgNO\(_3\) solution was added, however, Cl anions precipitated, indicating the existence of Cl anions. These findings demonstrate the adducts’ stoichiometry, which agrees with the general formulae proposed in Figure 1a,b.

3.2. Infrared and Raman Spectra

The adrenaline free binding chemical structure belongs to the \(\text{Cs}\) point group [33]. The N-methyl secondary amino group of adrenaline has been classified as point mass \(\text{CH}_2\text{NHCH}_3\). It contains 48 normal vibration modes, spread as \(\Gamma_{\text{vib}} = 33\Delta' + 15\Delta\) of the \(\text{Cs}\) point group. All modes are active in both “infrared and visible Raman laser spectra”. Figures 2 and 3 show the infrared and Raman spectra. Because of their poor symmetry and great complexity, the spectrum of the adrenaline molecule is difficult to understand, and because of our research goals, only 29 vibrations are investigated, which are dispersed as 19\(\Delta'\) vibrations and 10\(\Delta\) vibrations.
The vibrations allocated to the basic frequencies for adrenaline were established based on the size and comparative strengths of the observable bands in similarity with the associated substances [33] and are summarized in Table S1a,b.

The infrared spectra show two absorption bands for the aromatic C–C vibrations of the benzene ring at 1525 and 1496 cm$^{-1}$ and just one band at 1504 cm$^{-1}$. The infrared spectra of a substituted benzene ring show a stretching vibration motion of aromatic C–H vibrations at 3035 and 3023 cm$^{-1}$, while Raman spectra show it at 3024 and 3047 cm$^{-1}$. The absorption region for the alcohol C–O group, due to its extending vibration, is 1200–1000 cm$^{-1}$ which is absorbed in the infrared spectrum at 1157, 1175 and 1278 cm$^{-1}$, while in the Raman spectrum is absorbed at 1177 and 1288 cm$^{-1}$. The O–H vibrations infrared spectrum is at 3351, 3368 and 3386 cm$^{-1}$. For the aliphatic moiety, the C–H stretching vibrations occur in the IR spectrum at 3000 cm$^{-1}$ and in the Raman spectrum at 2995 cm$^{-1}$. The stretching vibrations (C–C) motion of ν(C–C) of the aliphatic moiety and CH–CH$_2$–NH–CH$_3$ absorbances are at 1205 and 1141 cm$^{-1}$, respectively. There are various bending deformation vibrations recorded at changed frequencies in the adrenaline-free moiety:

$\delta$(C–C–C): (IR = 612, 633, 649 and 689 cm$^{-1}$; Raman = 655 cm$^{-1}$)

$\delta$(C–O–H): (IR = 1029, 1061, 1082 and 1105 cm$^{-1}$; Raman = 1035, 1087 and 1109 cm$^{-1}$)

Using infrared and Raman laser spectra to compare free adrenaline ligand and its Y$^{3+}$, Ce$^{3+}$, Nd$^{3+}$, and Sm$^{3+}$ adrenaline adducts, we discovered that the distinctive bands ascribed to ν(O–H), ν(C–O) and δ(C–O–H) are shifted to lower wavenumbers, and their intensities

![Figure 3. Raman spectra of $[\text{Y}_2(\text{Adr})_2(\text{H}_2\text{O})_8]\text{Cl}_3\cdot8\text{H}_2\text{O}$, $[\text{Ce}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot10\text{H}_2\text{O}$, $[\text{Nd}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot6\text{H}_2\text{O}$, and $[\text{Sm}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot12\text{H}_2\text{O}$ adducts.](image-url)
are decreasing. These findings support the hypothesis that the adrenaline compounds of heavy metal ions are mediated by the two phenolic families of the monoamine moiety, as indicated in Figure 1a,b.

The Raman spectra of $\text{Y}^{3+}$, $\text{Ce}^{3+}$, $\text{Nd}^{3+}$, and $\text{Sm}^{3+}$ adrenaline adducts (Figure 3) show a spike enlargement with distortion in the extending vibration bands. This can be explained in light of the fact that the Raman analysis of fluorescent materials and structures is difficult to study due to the fluorescence intersection, which can overcome the integrally fragile Raman scattering simplification. The adrenaline bidentate ligand is thought to coordinate through the oxygens of phenolic groups, producing a steady compound [34,35].

3.3. Electronic Spectra

Two important bands of absorbances were detected at 299 and 376 nm and credited to the transitions $\pi \rightarrow \pi^*$, $n \rightarrow \pi^*$, correspondingly, for the benzene ring, phenolic, alcoholic, $\text{--NH}$, and methyl groups in the electronic spectra of adrenaline (Adr) free hormone moieties (Figure 4). These transitions were shifted either to a lower or higher wavenumber of the adrenaline adducts spectra of (Figure 4); $[\text{Y}_2(\text{Adr})_2(\text{H}_2\text{O})_8]\text{Cl}_3\cdot 8\text{H}_2\text{O}$ (297 and 308 nm), $[\text{Ce}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot 10\text{H}_2\text{O}$ (295 and 306 nm), $[\text{Nd}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot 6\text{H}_2\text{O}$ (295 and 308 nm) and $[\text{Sm}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot 12\text{H}_2\text{O}$ (297–308 nm).

![Figure 4](image-url). Electronic spectrum of adrenaline free and their complexes $[\text{Y}_2(\text{Adr})_2(\text{H}_2\text{O})_8]\text{Cl}_3\cdot 8\text{H}_2\text{O}$, and $[\text{Ce}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot 10\text{H}_2\text{O}$, $[\text{Nd}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot 6\text{H}_2\text{O}$, $\text{Sm}(\text{Adr})_2(\text{H}_2\text{O})_2]\text{Cl}_3\cdot 12\text{H}_2\text{O}$. 
The coordination of the adrenaline ligand with the trivalent rare earth metal ions has been confirmed. At 299 nm, the UV-visible peak matching to the ligand’s $\pi \rightarrow \pi^*$ transitions were detected [36]. As a result of coordination when binding with metal, the peaks of the $\pi \rightarrow \pi^*$ transitions are shifted to a lower wavelength, indicating the production of M–Adr metal adducts.

3.4. $^1$Hnmr Spectra

The typical $^1$H–NMR spectrum of adrenaline [37] consisted of 10 distinct signals resulting from ten different proton environments. The signals in the 7.0–8.0 ppm range and at 9.10, 9.04, 8.69, 6.804, 6.749, 6.619, 5.96, 4.775, 2.94, and 2.555 ppm are common for protons coupled to an aromatic or benzene ring. The H of the aromatic phenolic family emits signals at 9.10 and 8.69 ppm, the alcoholic –OH group emits signals at 5.96 ppm, and the H+ of the secondary amine –NH group emits signals at 9.04 ppm. The ligational behavior of the ligand was shown by comparing the $^1$H–NMR spectra of adrenaline free moiety with the $^1$H–NMR spectrum of the [Al(Adr)$_3$](NO$_3$)$_3$ adduct [38]. The presence of 2OH phenolic groups in the complexation behavior was suggested by the up–field of peaks at 9.10 and 8.69 ppm, which were attributed to H+s of the aromatic phenolic family to 8.363 and 8.913 ppm. The unshared secondary amine in the coordination process is ascribed to the downfield of –NH H+ from 9.04 ppm (Adr free ligand) to 9.264 ppm.

3.5. SEM and EDX Studies

SEM is a basic method that may be utilized to obtain a sense of the micro-features of the physical behavior of adrenaline adducts (Figure S1A–E). Even though this approach is not a reliable way to validate adduct formation, it may be used to check for the existence of a single component in synthetic adducts. The produced adducts exhibit a tiny particle dimension with nano-properties which result in the images. The generated adducts’ chemical examination by EDX shows a homogeneous distribution of the metal cation and adrenaline moiety. SEM studies of the shape of the surfaces of the adducts revealed tiny particles with a proclivity for forming aggregates of various forms in contrast to the starting materials. The peaks of these adducts’ EDX profiles (Figure 5A–E) relate to all components that make up the molecules of adrenaline adducts $[Y_2(Adr)_2(H_2O)_8]Cl_3\cdot8H_2O$, $[Ce(Adr)_2(H_2O)_2]Cl_3\cdot10H_2O$, $[Nd(Adr)_2(H_2O)_2]Cl_3\cdot6H_2O$, and $[Sm(Adr)_2(H_2O)_2]Cl_3\cdot12H_2O$ that were clearly identified, confirming the atoms ratio (molecular formulas).

![Figure 5. Cont.](image)
Figure 5. (A). EDX free adrenaline spectrum. (B). EDX spectrum of [Y\(_2\)(Adr)\(_2\)(H\(_2\)O)\(_8\)]Cl\(_3\)·8H\(_2\)O adduct. (C). EDX spectrum of [Ce(Adr)\(_2\)(H\(_2\)O)\(_2\)]Cl\(_3\)·10H\(_2\)O adduct. (D). EDX spectrum of [Nd(Adr)\(_2\)(H\(_2\)O)\(_2\)]Cl\(_3\)·6H\(_2\)O adduct. (E). EDX spectrum of [Sm(Adr)\(_2\)(H\(_2\)O)\(_2\)]Cl\(_3\)·12H\(_2\)O adduct.
3.6. Biological Evaluation

The antibacterial property of goal adducts against gram-positive bacteria, Bacillus subtilis, Streptococcus pneumonia, and Staphylococcus aureus and gram-negative bacteria, Escherichia coli and Pseudomonas aeruginosa, were tested in biological assessments. Additionally, two strains of fungus, Aspergillus niger and Penicillium, were also tested. Table 2 shows the results of the agar disc dispersion tests for the antibacterial activity of the target compounds, which are also shown in Figure S2. The diameters of the zone of inhibition (in mm) of the standard drug, tetracycline, against the gram-positive bacteria, B. subtilis and S. aureus, in addition to the gram-negative bacteria, E. coli and P. aeruginosa, were 36, 30, 31 and 35 mm, correspondingly, while the standard drug, amphotericin B, against Aspergillus flavus and Candida albicans were 18 and 19. Under matching circumstances, Table 2 displays that the adrenaline free binding moiety, \([Y \text{III}(\text{Adr})_2(H_2O)_8]\text{Cl}_3 \cdot 8\text{H}_2\text{O}, [Ce(Adr)_2(H_2O)_2]\text{Cl}_3 \cdot 10\text{H}_2\text{O}, [Nd(Adr)_2(H_2O)_2]\text{Cl}_3 \cdot 6\text{H}_2\text{O},\) and \([Sm(Adr)_2(H_2O)_2]\text{Cl}_3 \cdot 12\text{H}_2\text{O}\) adducts (10, 5, 5, 15, 5, 5 and 5), (15, 5, 0, 5, 10 and 10), (10, 5, 15, 10, 5, 15 and 10), respectively, relate to Bacillus subtilis, Streptococcus pneumonia, Staphylococcus aureus, Escherichia coli, Pseudomonas aeruginosa, Aspergillus niger and penicillium.

Table 2. Adrenaline and its adducts’ inhibition zone diameter (mm/mg sample).

| Sample | Inhibition Zone Diameter (mm/mg Sample) |
|--------|----------------------------------------|
|        | Bac. sub. (G+) | Srep. pen. (G+) | Stap. aure. (G+) | E. coli (G-) | Pseu. spp. (G+) | Asper. niger (Fungus) | Penici. spp. (Fungus) |
| Control: DMSO | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 |
| Standard | Tetracycline Antibacterial agent | 36 | 31 | - | 35 | 30 | - | - |
|          | Amphotericin B Antifungal agent | - | - | - | - | - | 18 | 19 |
|          | Adr | 8 | 6 | - | 10 | 5 | 0.0 | 0.0 |
|          | Y(III) | 10 | 5 | 5 | 15 | 5 | 5 | 5 |
|          | Ce(III) | 5 | 15 | 15 | 5 | 5 | 15 | 5 |
|          | Nd(III) | 15 | 5 | 5 | - | 5 | 10 | 10 |
|          | Sm(III) | 10 | 5 | 15 | 10 | 5 | 15 | 10 |

4. Conclusions

Four rare earth metal adrenaline complexes were formed from the reaction of adrenaline with \(Y^{3+}, Ce^{3+}, Nd^{3+}\) and \(Sm^{3+}\) ions in a methanolic solvent at 60 °C. The final reaction products have been isolated and characterized using elemental analyses (% of carbon, hydrogen and nitrogen), conductivity measurements, mid infrared, Raman laser, UV—Vis spectra, scanning electron microscopy and energy-dispersive X-ray spectroscopy (EDX). Upon the spectroscopic, conductivity and elemental analyses, the stoichiometric reactions indicated that the data obtained refer to 1:1 (M:L) for the \(Y^{3+}\), complex \([Y_2(\text{Adr})_2(H_2O)_8]\text{Cl}_3 \cdot 8\text{H}_2\text{O},\) while the molar ratio was 1:2 (M:L) for \(Ce^{3+}, Nd^{3+}\) and \(Sm^{3+}\) with formulas \([Ce(Adr)_2(H_2O)_2]\text{Cl}_3 \cdot 10\text{H}_2\text{O}, [Nd(Adr)_2(H_2O)_2]\text{Cl}_3 \cdot 6\text{H}_2\text{O},\) and \([Sm(Adr)_2(H_2O)_2]\text{Cl}_3 \cdot 12\text{H}_2\text{O}\). The infrared and Raman laser spectra interpreted the mode of interactions which associated through the two phenolic groups of the catechol moiety. The adrenaline compounds have been screened for their in vitro antibacterial activity against four bacteria, both gram-positive and gram-negative, and two strains of fungus. The metal complexes were shown to possess more antibacterial and antifungal activities than the free adrenaline compound.
**Supplementary Materials:** The following are available online at https://www.mdpi.com/article/10.3390/cryst11121536/s1, Figure S1A: SEM photo of adrenaline free binding moiety, Figure S1B: SEM photo of \( [Y_2(Adr)_2(H_2O)_{8}]Cl_3.8H_2O \) adduct, Figure S1C: SEM photo of \( [Nd(Adr)_2(H_2O)_2]Cl_3.6H_2O \) adduct, Figure S1D: SEM photo of \( [Ce(Adr)_2(H_2O)_2]Cl_3.10H_2O \) adduct, Figure S1E: SEM photo of \( [Sm(Adr)_2(H_2O)_2]Cl_3.12H_2O \) adduct, Figure S2: The width of the inhibition zone of adrenaline and its adducts against bacteria and fungus, Table S1a: Infrared frequency categorization (cm\(^{-1}\)) and preliminary designations of adrenaline and its adducts, Table S1b: Categorization Raman frequencies (cm\(^{-1}\)) and tentative adrenaline.

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