

Transition Metal Complexes of Derivatized Chiral Dihydro-1,2,4-triazin-6-ones. Template Synthesis of Nickel(II) Tetraaza-(4N-M) Complexes Incorporating the Triazinone Moiety

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Abstract: Template reactions of (L)-3-acetyl-5-methyl-1-phenyl-4,5-dihydro-1,2,4-triazin-6-one (8a) or (L)-3-acetyl-5-benzyl-1-phenyl-4,5-dihydro-1,2,4-triazin-6-one (8b) with nickel(II) acetate in the presence of a suitable diamine such as: 1,3-diaminopropane, o-phenylenediamine, or ethylenediamine afforded square planar tetraaza nickel(II) complexes of the type 4N-M (Scheme 4). Single crystal X-ray structural analysis of one of the complexes (9b) revealed square planar geometry around the nickel ion and symmetrical coordination of the triazinone moieties. The complexes were characterized by elemental analyses, magnetic measurements, and electrical conductivities.

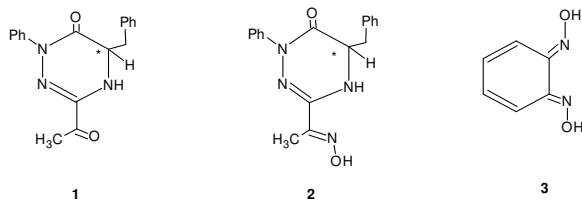
Key words: Triazinones, tetraaza complexes, template effect synthesis, nickel(II) complexes

INTRODUCTION

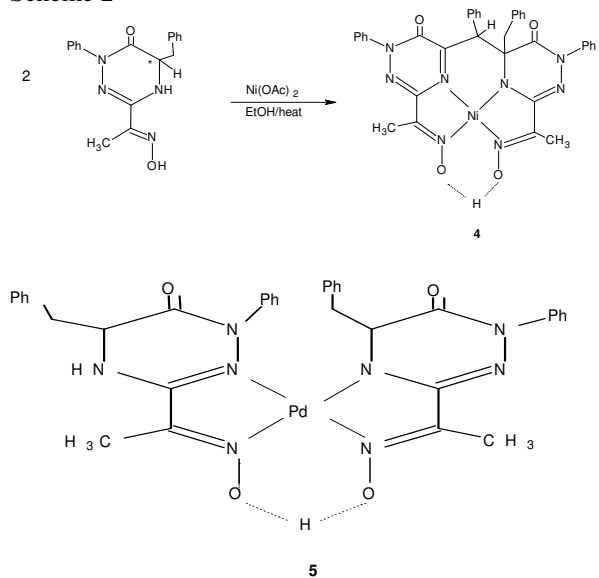
The chiral 4,5-dihydro-1,2,4-triazin-6-ones (e.g. compound 1, Scheme 1) were prepared by reacting α -amino esters with hydrazonoyl chlorides (precursors of nitrile imines, the reactive 1,3-dipolar species) in a one-pot synthesis^[1]. The respective 4,5-dihydrotriazinone oximes (e.g. compound 2, Scheme 1) were shown to be readily accessible via oximation of the C3-acetyl group using hydroxylamine. The oxime compounds(2) are capable of functioning as bidentate ligand systems, very much similar to 3,5-cyclohexadiene-1,2-dione dioxime (3) or dimethylglyoxime (DMG)^[2,3]

It has been found that the oxime (2) coordinates to metal ions such as Ni^{2+} in a bidentate fashion utilizing the amidrazone nitrogen HN(4) and the oxime nitrogen as the donor atoms. This occurred in concomitant with a carbon-carbon coupling between the benzylic carbon of one oxime ligand and the chiral carbon center of the

Scheme 1



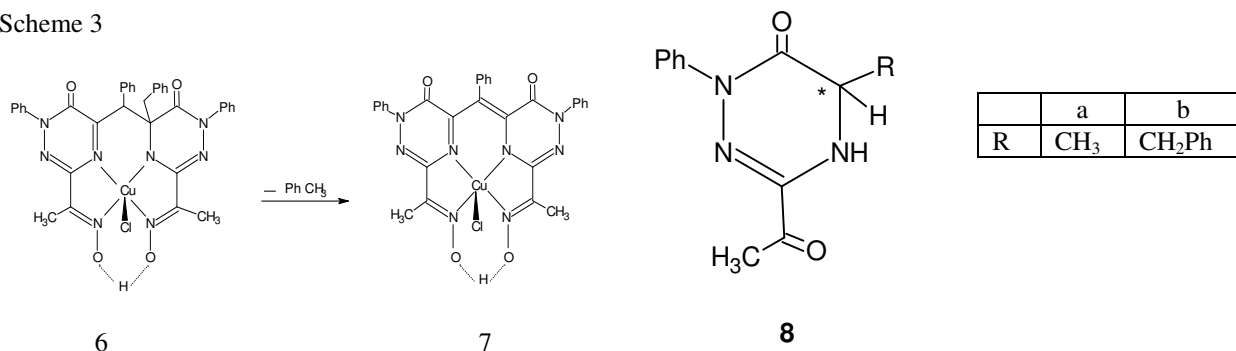
Scheme 2



other oxime ligand (Scheme 2), the resulting structures (4) are square-planar dioxime complexes^[4].

Attempts to introduce Pd^{2+} ion instead of Ni^{2+} in (4) resulted in a square-planar palladium(II) complex where the two oxime ligands (2) are coordinated unsymmetrically and thus not allowing the expected carbon-carbon coupling to take place (5)^[5]. This has occurred probably due to the larger size of the Pd^{2+} ion

Scheme 3



compared to Ni²⁺ and the pseudomacrocyclic cavity in (4) could not accommodate Pd²⁺.

In addition, the reaction of oxime (2) with copper(II) acetate gave complex (6) in minute quantities, apparently because toluene was eliminated from it giving rise to complex (7), (Scheme 3). The crystal structure of (7) was proved by X-ray crystallography and found to be slightly distorted square-pyramidal geometry^[6]. This geometry was surprisingly supported by the presence of a chlorine atom bonded to the copper center. The source of this chlorine atom is not definitely known but it could be from the chlorinated solvent used in the preparation and crystallization processes^[7].

Since the overall aim of this work is to incorporate the triazinone moiety into a real macrocyclic ring system, we attempted a new synthetic route and that is the reaction of the triazinone compounds with suitable diamines in the presence of the transition metal cation, i.e., template effect synthesis^[8].

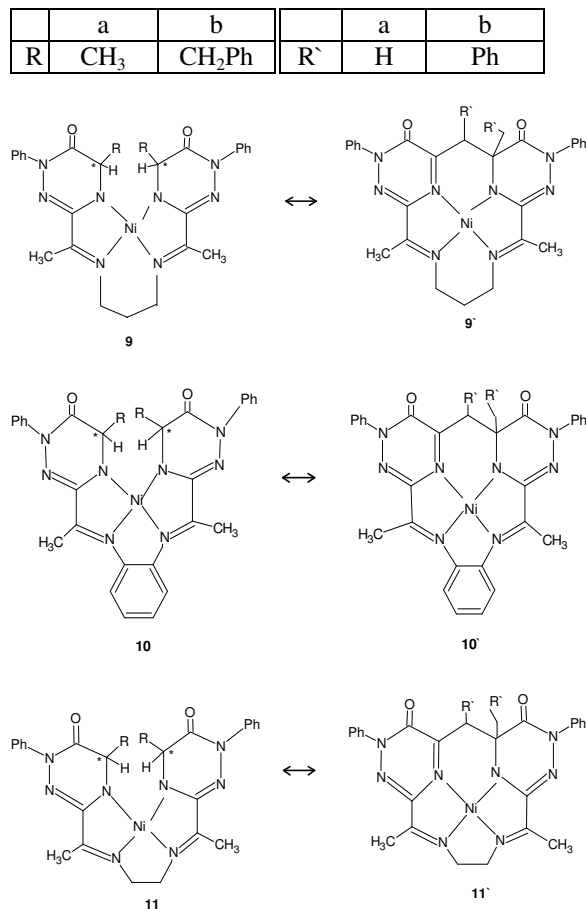
RESULTS AND DISCUSSION

It has been observed that in order for the carbon-carbon coupling between the benzylic carbon of one triazinone ligand and the chiral carbon center of the other to take place, the two oxime ligands must coordinate in a symmetrical way (e.g. compound 4). This has initiated the idea of linking the two triazinone moieties from the bottom, using a suitable diamine, assuming that the carbon-carbon coupling will link them from the top. As such this should produce a macrocyclic ring system of a tetraaza-type (4N-M). Thus, two types of the 3-acetyl triazinones (compounds 8) and three types of diamines were used, these are: 1,3-diaminopropane, o-phenylenediamine, and ethylenediamine.

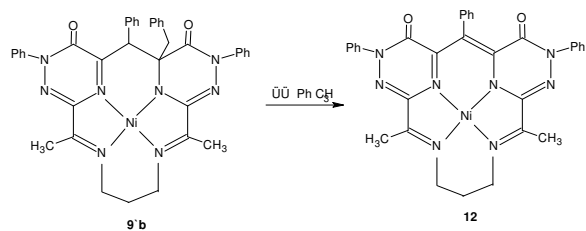
The reaction of two moles of the triazinone (8a) with one mole of 1,3-diaminopropane in the presence of one mole of nickel(II) acetate is expected to produce compound 9a or 9b (Scheme 4). The nature of the product depends on whether cyclization, via carbon-carbon coupling, has taken place or not. The resulting complex, however, is diamagnetic indicating square planar geometry for Ni²⁺. It is also neutral in 10⁻³M DMF solution. The IR spectral data are not diagnostic for such compounds. The elemental analyses indicate, most likely, an open-chain structure (9a). Similarly, two moles of the triazinone(8b) were reacted with one mole of 1,3-diaminopropane in the presence of one mole nickel(II) acetate. The produced oily/gummy product was purified until it was solid. The solid was recrystallized from hot aqueous methanol and the results of elemental analyses were consistent with structure 9b which is neutral in 10⁻³ M DMF solution. Magnetic measurements showed that 9b is diamagnetic. In an attempt to grow crystals of 9b, several organic solvents were tried. The crystals used in the X-ray crystallographic analysis for the determination of the structure depicted in Fig. 1 were grown from acetone. When a sample of 9b was recrystallized from dichloromethane, a much deeper color crystals were obtained. These deep color crystals are most likely compound 12 (Scheme 5), results of elemental analyses are consistent with structure 12 (for Ni C₃₂H₂₇N₈O₂ Calcd: C 62.57, H 4.40, N 18.25%; Found: C 62.26, H 4.56, N 18.05%). Compound 12 is also diamagnetic and neutral in DMF. Attempts to grow crystals of the compound are still going on to prove the structure further by X-ray work.

The same synthetic roots (with minor variations) were employed when o-phenylenediamine and ethylenediamine were used to prepare compounds 10a, 10b and 11a, 11b, respectively (Scheme 4). Apparently, macrocyclic closure was not observed in these

Scheme 4



Scheme 5



complexes, however, work is still going on to elucidate their structural features.

CRYSTAL STRUCTURE DETERMINATION OF 9B

Crystals suitable for X-ray crystallographic analysis were grown from acetone. The structure shown in Fig. 1 was determined at the X-Ray Crystallography

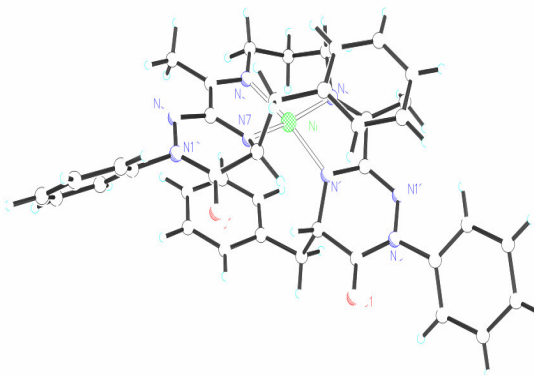


Fig. 1: Plot of the crystal structure of 9b

Laboratory, University of Kansas, USA. The absolute configuration that is shown was determined experimentally using anomalous dispersion of the X-rays.

EXPERIMENTAL

Melting points (uncorrected) were determined using a Gallenkamp melting point apparatus in one-end glass capillaries. The magnetic moments were measured for finely ground solid samples at ambient temperature using a Johnson Matthey Magnetic Susceptibility Balance. Elemental analyses were performed on an Euro EA3000 Elemental Analyzer (Euro Vector), at the Department of Chemistry, Hashemite University, Jordan.

Synthesis of 1-phenylhyrazono-1-chloroacetone (Ph-NH-N=C(Cl)-C(O)-CH₃) (Hydrazonoyl Chloride)

General procedure:

- **Preparation of benzenediazonium chloride PhN₂⁺Cl⁻ solution:** Aniline (0.1 mol) in 80 ml of 1:1 H₂O/conc. HCl was cooled to about 0°C in an ice bath with stirring. Then a solution of sodium nitrite (7.6 g/25 mL H₂O) was added dropwise. Stirring was continued for 30 min. This preparation was done according to reference 8 and the solution was used as it is in the next step.
- **Reaction of benzenediazonium chloride with 3-chloro-2,4-pentanedione:** To a cold stirred solution (-10°C) of 3-chloro-2,4-pentanedione (13.5g, 0.1 mol/80 mL H₂O), a cold solution (-5°C) of benzenediazonium chloride was added gradually. Stirring of the resulting mixture was

continued for 15 minutes. The crude solid product was collected by suction filtration and washed several times with cold water and then with little petroleum ether, dried and recrystallized from chloroform / petroleum ether. Yield = 15.1g (79%); mp 140-14°C ; literature mp. 141-142 °C [9].

Synthesis of: L-3-Acetyl-5-methyl-1-phenyl-4,5-dihydro^{-[1,2,4]}triazin-6-one(8a): This compound was prepared via direct interaction between 1-chloro-1-phenylhydrazono-2-propanone (hydrazonoyl chloride) and L-alanine methyl ester hydrochloride in the presence of triethylamine, according to the following general procedure:

To a stirred solution of the hydrazonoyl chloride (20 mmol) in 50 mL of CHCl₃, a solution of the appropriate amino acid ester hydrochloride (20 mmol) in 50mL of methanol was added. The mixture was cooled to about 0°C and triethylamine (18 mL, 0.17 mol) in 10 mL of methanol was added dropwise keeping the temperature not higher than 5°C . The reaction mixture was kept under continuous stirring at 0°C for 2-3 h. and then at ambient temperature for 24 hours. The solvent was then removed *in vacuo*, and the residue was washed with water. The resulting crude solid product was then collected and recrystallized from aqueous ethanol. The product was obtained in the form of prismatic crystals, 78% yield, mp 114-115°C.

Synthesis of: L-3 Acetyl-5-benzyl-1-phenyl-4,5-dihydro^{-[1,2,4]}triazin-6-one(8b): The same procedure for compound (8a) above except that L-phenylalanine methyl ester hydrochloride was used. Yield (83%), mp. 64-66°C.

N,N'-Bis [(1-phenyl-5-methyl-4,5-dihydro-6-oxo-1,2,4-triazin-3-yl)-ethylidene] propane-1,3-diamine nickel (II) , (9a): The acetyl compound (8a) (1.39g, 6 mmol) and nickel acetate tetrahydrate (0.75 g, 3 mmol) in 80 mL of absolute ethanol were stirred at room temperature for few minutes. 1,3-diaminopropane (0.24 g, 3.25 mmol) in absolute ethanol (20 mL) was added dropwise over a period of 10 min. The reaction mixture was brought to reflux under constant stirring for about 70 hours. During the first few hours the reaction mixture assumed dark green color which changed to red then to brown color. This color persisted until the end of the reflux time. The solvent was removed *in vacuo*, and a gummy residue was obtained. This was treated with hot aqueous ethanol and left aside to solidify. The

crude solid was recrystallized from hot aqueous methanol. Yield =1.15 g (70%), mp > 330°C.

Anal: Calcd for Ni C₂₇ H₃₀ N₈O₂ (556.7), C 58.20, H 5.39, N 20.12%

Found: C 57.07, H 5.99, N 20.07%

N,N'-Bis [(1-phenyl-5-benzyl-4,5-dihydro-6-oxo-1,2,4-triazin-3-yl)-ethylidene] propane-1,3-diamine nickel (II) , (9b): The acetyl compound (8b), (1.84g, 6 mmol) and nickel acetate tetrahydrate (0.75 g, 3 mmol) in 80 mL of absolute ethanol were stirred at room temperature for few minutes. 1,3-Diaminopropane (0.24 g, 3.25 mmol) in absolute ethanol (20 mL) was added dropwise over a period of 10 min. The reaction mixture was brought to reflux under constant stirring for about 70 h. During the first few hours the reaction mixture assumed dark green color which changed to red then to brown color. This color persisted until the end of the reflux time. The solvent was removed *in vacuo*, and a gummy residue was obtained. This was treated with hot aqueous ethanol and left aside to solidify. The solid was recrystallized from hot aqueous methanol. Yield = 1.26g (60%) , mp > 330°C.

Anal: Calcd for Ni C₃₉ H₃₈ N₈O₂ (708.7), C 66.04, H 5.36, N 15.80%

Found: C 67.07, H 5.56, N 15.64%

N,N'-Bis [(1-phenyl-5-methyl-4,5-dihydro-6-oxo-1,2,4-triazin-3-yl)-ethylidene] o-phenylenediamine nickel (II) , (10a): The acetyl compound (8a) (1.39g, 6 mmol) and nickel acetate tetrahydrate (0.75 g,3 mmol) in 60 mL of absolute ethanol were stirred at room temperature for few minutes. o-Phenylenediamine (0.35 g, 3.25 mmol) in absolute ethanol (20 mL) was added dropwise over a period of 10 min. The reaction mixture was brought to reflux under constant stirring for about 3 days. The reaction mixture assumed deep- red color which changed to brown then to black color. This color persisted until the end of the reflux time. The solvent was removed *in vacuo*, and the produced compound was recrystallised from dichloromethane. Fine powder was always obtained. Yield =0.97 g (55%) , mp > 330°C.

Anal: Calcd for Ni C₃₀ H₂₈ N₈O₂ (590.7), C 60.94, H 4.74, N 18.96%

Found: C 60.07, H 4.56, N 18.64%

N,N'-Bis [(1-phenyl-5-benzyl-4,5-dihydro-6-oxo-1,2,4-triazin-3-yl)-ethylidene] o-phenylenediamine nickel (II) , (10b): The acetyl compound (8b), (1.84g, 6 mmol) and nickel acetate tetrahydrate (0.75 g, 3

mmol) in 60 mL of absolute ethanol were stirred at room temperature for few minutes. *o*-Phenylenediamine (0.35 g, 3.25 mmol) in absolute ethanol (20 mL) was added dropwise over a period of 10 minutes. The reaction mixture was brought to reflux under constant stirring for about 7 days. The reaction mixture assumed green color which changed to brown. This color persisted until the end of the reflux time. The solvent was removed *in vacuo*, and the product obtained was recrystallised from dichloromethane and deep brown powder was obtained. Yield = 1.14g (50%), mp > 330°C. Anal: Calcd for Ni C₄₂H₃₆N₈O₂ (742.7), C 67.86, H 4.85, N 15.08%
Found: C 66.97, H 4.46, N 15.61%

N,N'-Bis [(1-phenyl-5-methyl-4,5-dihydro-6-oxo-1,2,4-triazin-3-yl)-ethylidene] ethane-1,2-diamine nickel (II), (11a): The acetyl compound (8a) (1.39g, 6 mmol) and nickel acetate tetrahydrate (0.75 g, 3 mmol) in 80 mL of absolute ethanol were stirred at room temperature for few minutes. Ethylenediamine (0.20 g, 3.33 mmol) in absolute ethanol (20 mL) was added dropwise over a period of 10 minutes. The reaction mixture was brought to reflux under constant stirring for about 3 days. During the first few hours the reaction mixture assumed red color and this color persisted until the end of the reflux time. The solvent was removed *in vacuo*, and an oily residue was obtained. This was treated with hot aqueous ethanol and left aside to solidify. The solid was recrystallized from acetone. Yield = 0.93g (55%), mp > 330°C
Anal: Calcd for Ni C₄₂H₃₆N₈O₂ (542.7), C 57.49, H 5.16, N 20.64%
Found: C 57.97, H 5.46, N 20.11%

N,N'-Bis [(1-phenyl-5-benzyl-4,5-dihydro-6-oxo-1,2,4-triazin-3-yl)-ethylidene] ethane-1,2-diamine nickel (II), (11b): The acetyl compound (8b), (1.84g, 6 mmol) and nickel acetate tetrahydrate (0.75 g, 3 mmol) in 80 mL of absolute ethanol were stirred at room temperature for few minutes. Ethylenediamine (0.20g, 3.33 mmol) in absolute ethanol (20 mL) was added dropwise over a period of 10 minutes. The reaction mixture was brought to reflux under constant stirring for about 3 days. During the first few hours the reaction mixture assumed red color and this color persisted until the end of the reflux time. The solvent was removed *in vacuo*, and an oily residue was obtained. This was treated with hot aqueous ethanol and left aside to solidify and the solid was recrystallized from acetone. Yield = 1.35g (65%), mp > 330°C

Anal: Calcd for Ni C₄₂H₃₆N₈O₂ (694.7), C 65.63, H 5.18, N 16.12%
Found: C 64.97, H 5.66, N 15.67%

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