A comparative study of the coordination of saccharinate (sac), thiosaccharinate (tsac) and benzisothiozolinate (bit) ligands to *trans*-[$PdCl_2(H_2NBz)_2$]: Molecular structure of *cis*-[$Pd(bit)_2(H_2NBz)_2$]

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Abstract A comparative study of reactions of saccharinate (sac), thiosaccharinate (tsac) and benzisothiozolinate (bit) with *trans*-[$PdCl_2(H_2NBz)_2$] is reported. While in all cases substitution of both chlorides occurs, product types differ for the three closely related ligands. With sodium saccharinate, *trans*-[$Pd(N\text{-}sac)_2(H_2NBz)_2$] results in which the sac ligands are N-bound. A similar N-bound coordination is observed with sodium benzisothiazolinate, but a crystal structure shows that they adopt a mutual *cis*-arrangement in *cis*-[$Pd(N\text{-}bit)_2(H_2NBz)_2$]. In contrast, with sodium thiosaccharinate it is proposed that the new ligands adopt an S-bound coordination mode in *trans*-[$Pd(S\text{-}tsac)_2(H_2NBz)_2$].

Keywords: palladium; benzylamine; benzisothiazolinate; saccharinate; thiosaccharinate

Introduction

Saccharinate (sac) and thiosaccharinate (tsac) anions (Chart) are versatile poly-functional ligands, shown to adopt a variety of coordination modes, and consequently their coordination chemistry has been widely studied [**1**]. Palladium(II) and platinum(II) complexes of these ligands have been detailed [**2-5**] with some showing promising biological properties [**6**]. In

contrast, the coordination chemistry of the related benzisothiozolinate (bit) anion (Chart), resulting from deprotonation of the acidic imine hydrogen in benzoisothiazolinone, remains virtually unexplored; as far as we are aware there are only two literature reports concerning the coordination chemistry of this ligand [**7-8**]. Griffith and co-workers have reported the synthesis of *cis*-[Pd(N-bit)$_2$(κ^2-en)] (en = ethylenediamine) and [Pt(NH$_3$)$_2$(N-bit)$_2$], the former being characterized by single crystal X-ray crystallography [**7**], while we have recently detailed the synthesis of a number of square-planar palladium complexes, *trans*-[Pd(N-bit)$_2$L$_2$], with amine, amide and diphosphine co-ligands [**8**]. The latter can be formed *via* two synthetic routes, namely reaction of [Pd(bit)$_2$].H$_2$O with neutral ligands or *via* displacement of both chlorides in *trans*-[PdCl$_2$L$_2$]. Herein we develop further the coordination chemistry of the benzisothiozolinate anion in a comparative study of reactions of saccharinate (sac), thiosaccharinate (tsac) and benzisothiazolinate (bit) with *trans*-[PdCl$_2$(H$_2$NBz)$_2$]. The surprisingly outcome of this simple study was the isolation of different product types in each case.

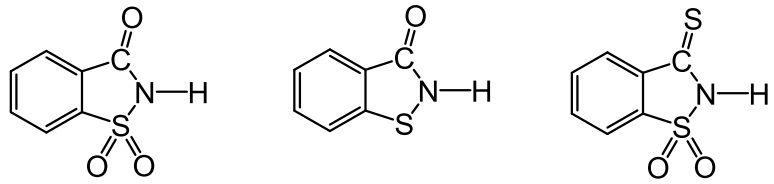


Chart. Saccharin (sacH), benzisothiazoline (bitH) and thiosaccharin (tsacH)

Experimental

General methods

^{1}H NMR spectra were recorded on Varian Unity spectrometer in CDCl$_3$ or d^6-dmso. IR spectra were recorded on Shimadzu FT-IR 8400 spectrophotometer in the 400-4,000 cm^{-1} range using KBr discs and in the 200-600 cm^{-1} using CsI discs Elemental analysis were carried out at Al Al-Bayt University, Jorden using a Euro-vector EURO EA 300 elemental analyzer. Melting points were measured on a Gallenkhamp melting point apparatus and are uncorrected. Conductivity measurements were carried out on 10^{-3} M solutions using a digital

conductivity meter. Na_2PdCl_4, benzoisothiazolinone (Hbit), benzylamine and sodium saccharinate were purchased and used as received. Thiosaccharin [**9**] and *trans*-$[PdCl_2(H_2NBz)_2]$ (**1**) [**10**] were prepared by literature methods.

Synthesis of 2

A solution of Nasac (0.285 g, 1.35 mmol) in MeOH (5 cm^3) was added to a solution of **1** (0.244 g, 0.62 mmol) in MeOH (10 cm^3). The mixture was stirred at room temperature for 3 h. The resulting yellow solid was collected by filtration, washed with MeOH and dried in vacuum. It was recrystallized from $CHCl_3$/MeOH to afford **2** as a yellow crystalline solid. Yield 0.341 g, 73%. *Ana*l. Calc. for $C_{28}H_{26}N_4O_6PdS_2$: C, 49.1, H, 3.8, N, 8.2. Found: C, 49.2, H, 3.7, N, 8.2. Molar conductivity (DMSO): 0.40 (Ω^{-1} mol^{-1} cm^{-1}). IR (KBr): 3265w, 3130w, 3029w, 1672s, 1593w, 1451w, 1290s, 1155m, 563m cm^{-1}. ^{1}H NMR ($CDCl_3$): δ 7.94-7.92 (m, 2H, sac), 7.87-7.85 (m, 2H, sac), 7.75-7.72 (m, 4H, sac), 7.30-7.22 (m, 10H, Ph), 4.34 (bs, 4H, $2NH_2$), 3.97-3.93 (m, 4H, $2CH_2$) ppm. Mp: 224-226 ℃.

Synthesis of 3

A solution of tsac (0.051 g, 0.26 mmol) in MeOH (5 cm^3) was added to a solution of **1** (0.051 g, 0.13 mmol) in MeOH (10 cm^3). The mixture was stirred at 30 ºC for 2 h. The yellow-orange solid formed was collected by filtration and dried under vacuum. Yield 0.068 g, 75%. *Anal.* Calc. for $C_{28}H_{26}N_4O_4PdS_4$: C, 46.9, H, 3.7, N, 7.8. Found, C, 46.9, H, 3.8, N, 8.0. Molar conductivity (DMSO): 0.40 (Ω^{-1} mol^{-1} cm^{-1}). IR (KBr): 3425sb, 3051w, 2922w, 1541m, 1463m, 1384s, 1163s, 1004m, 806m, 370s cm^{-1}. ^{1}H NMR (DMSO-d^6): δ 7.89 (dd, J 8.0, J 3.2, 4H, tsac), 7.71 (t, J 8.0, 2H, tsac), 7.58 (t, J 8.0, 2H, tsac), 7.29 (s, 10H, Ph), 4.58 (bs, 4H, $2NH_2$), 3.69 (s, 4H, $2CH_2$) ppm.

Synthesis of 4

A solution of Nabit (0.048 g, 0.28 mmol) in MeOH (5 cm^3) was added to a solution of **1** (0.055 g, 0.14 mmol) in MeOH (10 cm^3) and stirred for 3 h at room temperature to give a yellow-brown solution. The solution was filtered and left to evaporate to afford yellow crystals. These were collected by filtration, washed with water and dried in a vacuum oven. Yield 0.075g, 87%. *Anal*. Calc. for $C_{28}H_{26}N_4O_2PdS_2$: C, 53.3, H, 4.1, N, 9.2. Found: C, 53.4,

H, 4.4, N, 9.5. Molar conductivity (DMSO): 0.80 (Ω^{-1} mol^{-1} cm^{-1}). IR (KBr): 3195w, 3112w, 2927w, 1650s, 1539s, 1450m, 1290m, 1155m, 459w, 342m cm^{-1}. ^{1}H NMR (DMSO-d^6): δ 7.76 ppm (d, J 7.7, 2H, bit), 7.66 (d, J 7.7, 2H, bit), 7.57-7.23 (m, 14H, Ph+bit), 5.56 (s, 4H, $2NH_2$), 3.56 (s, 4H, $2CH_2$) ppm. Mp: 208-210℃.

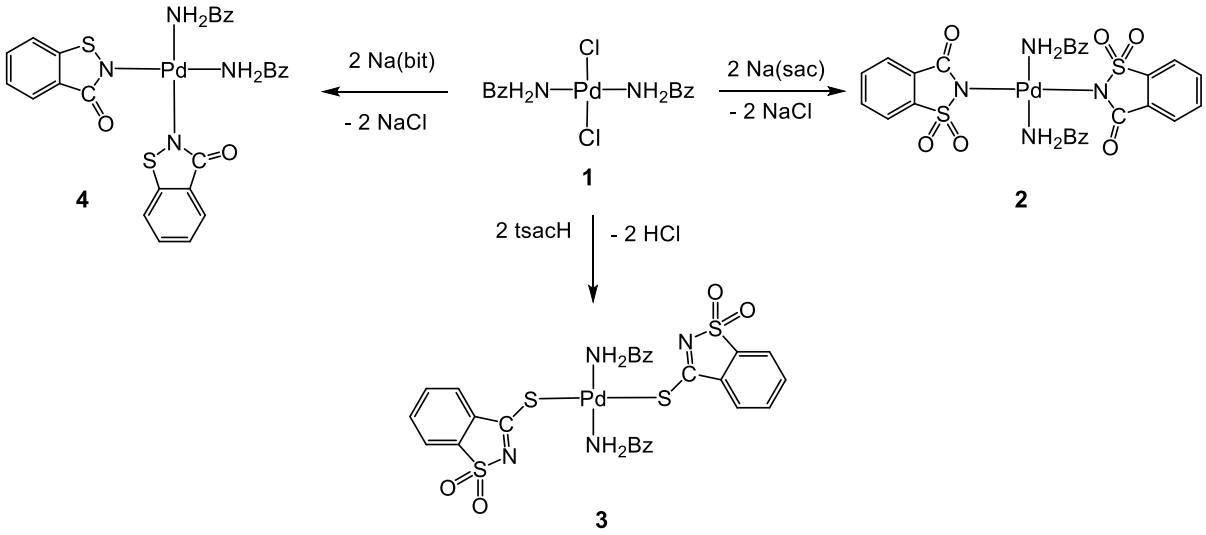
X-ray crystallography

Crystals of *cis*-[Pd(bit)$_2$(H_2NBz)$_2$] (**4**) suitable for X-ray crystallography were produced by slow evaporation of a methanol solution. A yellow crystal with approximate dimensions 0.10 x 0.10 x 0.10 mm^3 was mounted on a glass fiber and all geometric and intensity data were taken from this sample using a STOE-IPDS diffractometer with Mo-Kα radiation (λ = 0.7103 Å, graphite monochromator). Absorption corrections were made using the IPDS software package [**11**]. All structures were solved by direct methods and refined using full-matrix least-square routines against F^2 with SHELXL-97 [**12**]. Non-hydrogen atoms were refined with anisotropic displacement parameters. Hydrogen atoms were included in the models by calculating the positions (riding model) and refined with calculated isotropic displacement parameters. Illustrations were generated using DIAMOND 3.0 [**13**].

Results and discussion

Addition of two equivalents of sodium saccharinate to a methanol solution of *trans*-[$PdCl_2$(H_2NBz)$_2$] (**1**) resulted in the slow formation of *trans*-[Pd(N-sac)$_2$(H_2NBz)$_2$] (**2**) isolated in 73% yield as a yellow solid (Scheme). Elemental analysis supports the substitution of both halides in **1**, as does the symmetrical nature of the ^{1}H NMR spectrum. This simple substitution and formation of the *trans*-saccharinate complexes mirrors behavior previously noted by us [**5c-d**] and others [**4e**]. Reaction of **1** with thiosaccharin in methanol at 30 ^{o}C resulted in formation of *trans*-[Pd(S-tsac)$_2$(H_2NBz)$_2$] (**3**) as a yellow-orange solid in 75% yield (Scheme). Elemental analysis was indicative of the substitution of both chlorides and this is consistent with the ^{1}H NMR spectrum. On the basis of the observation of an IR band at 1004 cm^{-1}, which is attributed to the C-S vibration and is shifted some 35 cm^{-1} from the corresponding vibration in thiosaccharin, we propose that binding of the tsac ligands occurs through sulfur. This is not unexpected and is in accord with the established chalcogenophilic

nature of Pd(II) and also with previous work from our laboratory [**5d**]. While we have been unable to crystallographically characterise **2** and **3** we strongly believe that the *trans* arrangement confirmed in **1** is maintained upon chloride substitution. The basis of this is the relatively simple nature of their IR spectra and the aromatic region of the ^{1}H NMR spectra, both being consistent with retention of the (approximate) D_{2h} symmetry. This assignment is also made on the basis of the chemical shifts of the amine protons at δ 4.34 and 4.58 respectively (see below).



Scheme. Reactions of *trans*-[$PdCl_2(H_2NBz)_2$] (**1**) with two equivalents of Na(sac), tsacH and Na(bit)

Reaction of two equivalents of sodium benzisothiozolinate with **1** in methanol gave a yellow-brown solution and, unlike previous reactions with sodium saccharinate and thiosaccharin, no solids initially precipitated from the solution. However, after filtration and upon standing for a few days, slow evaporation of the methanol led to the growth of yellow crystals identified as *cis*-[$Pd(N\text{-}bit)_2(H_2NBz)_2$] (**4**) in 87% yield. The ^{1}H NMR spectrum was significantly different to those of **1**-**3**, being more complicated with overlapping signals in the aromatic region (indicative of a lowering of the D_{2h} symmetry), while the amine protons appeared at δ 5.56. We have recently reported [$Pd(H_2NBz)_3Cl$][Cl] and note that its ^{1}H NMR spectrum shows two amine resonances in an approximate 2:1 ratio at δ 4.70 (4H) and 5.26 (2H) [**15**] assigned to the mutual *trans* amines and that lying *trans* to the chloride respectively. This

suggested to us that the amines in **4** adopted a relative *cis* orientation. A single crystal analysis was carried out in order to determine the coordination mode of the bdt ligands and relative arrangement of amines. The results of this are shown in Figure 1 and its caption.

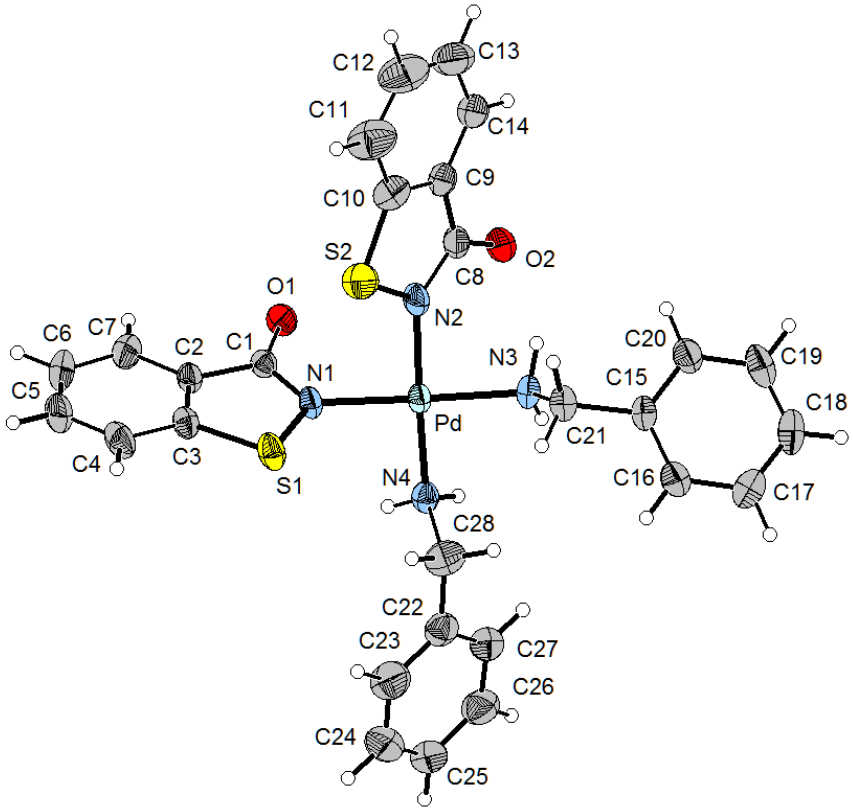


Figure 1. The molecular structure of *cis*-[Pd(N-bit)$_2$(H$_2$NBz)$_2$] (**4**) with selected bond lengths (Å) and angles (°): Pd-N(1) 2.022(2), Pd-N(2) 2.015(3), Pd-N(3) 2.045(2), Pd-N(4) 2.056(3), N(1)-Pd-N(2) 90.3(1), N(3)-Pd-N(4) 90.1(1), N(1)-Pd-N(3) 178.5(1), N(2)-Pd-N(4) 177.8(1).

The structure confirms that the two bit ligands bind in a monodentate fashion through nitrogen, but the main surprise was their relative *cis*-arrangement. All four palladium-nitrogen bond lengths are similar, although those to the benzisothiazolinate ligands [Pd-N(1) 2.022(2), Pd-N(2) 2.015(3) Å] are slightly shorter than to the benzylamine groups [Pd-N(3) 2.045(2), Pd-N(4) 2.056(3) Å]. The latter compare well with the related bonds in *trans*-[PdCl$_2$(H$_2$NBz)$_2$] [Pd-N 2.050(4) and 2.046(2) Å] [**14**] and [PdCl(H$_2$NBz)$_3$]Cl.H$_2$O [Pd(1)-N(1) 2.061(2), Pd(1)-N(2) 2.053(2), Pd(1)-N(3) 2.063(2) Å] [**15**]. Both Pd-N(bit) bond lengths in **4** are significantly shorter than those in [Pd(N-bit)$_2$(κ^2-Ph$_2$PCH$_2$CH$_2$PPh$_2$)] **[**Pd-N 2.070(3) & 2.100(3) Å] [**8**], being closer to in [Pd(N-bit)$_2$(κ^2-H$_2$NCH$_2$CH$_2$NH$_2$)] [Pd-N 2.029(2) & 2.031(2) Å] [**7**], suggesting that they may be sensitive to a *trans*-influence.

Complex **4** is the third example of a palladium-bis(benzisothiozolinate) complex and like the diphosphine and diamine derivatives, it also contains a *cis* arrangement of benzisothiazolinate ligands. Thus, it may be that these ligands inherently prefer to adopt a relative *cis* orientation, although in **4** this is the first example where the arrangement is not imposed by a chelating co-ligand. A possible explanation for the *cis* geometry in **4** comes from inspection of the intermolecular packing of the individual molecules. Thus as shown in Figure 2, pairs of molecules are strongly associated by hydrogen bonds between the amine protons and the oxygen atoms of the benzisothiazolinate ligands. This arrangement brings the two palladium atoms in close proximity [Pd….Pd 3.839 Å].

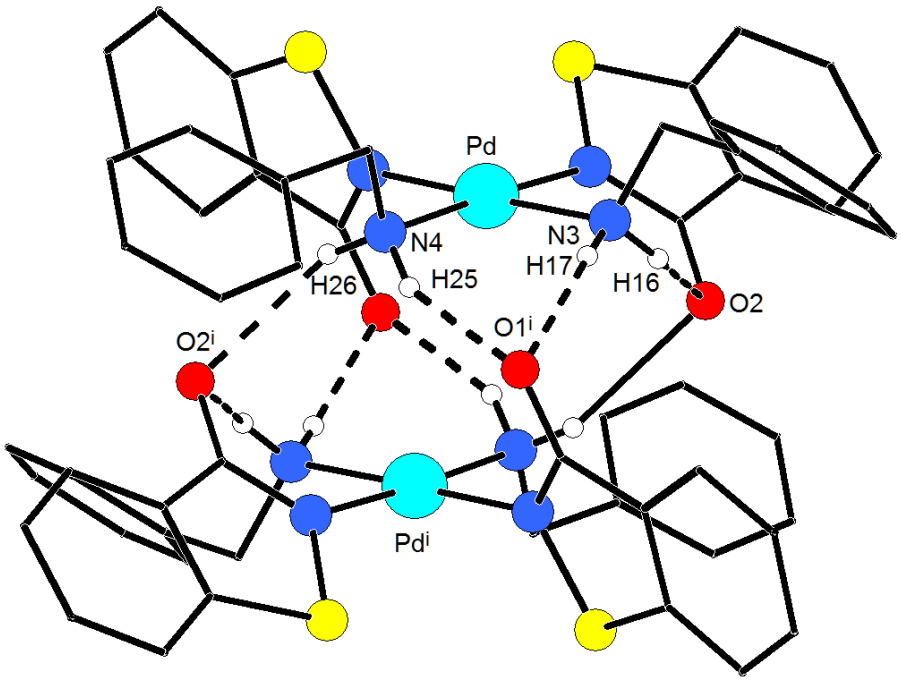


Figure 2. Packing of two molecules of **4** with intermolecular bond lengths (Å)

	D-H /Å	H...A /Å	D...A /Å	<(DHA) /°
N3-H16···O2	0.97(3)	1.95(3)	2.826(4)	150(2)
N3-H17···O1^{i}	0.82(3)	2.02(3)	2.813(3)	162(3)
N4-H25···O1^{i}	0.78(4)	2.18(4)	2.882(3)	151(3)
N4-H26···O2^{i}	0.76(4)	2.45(4)	3.102(3)	145(4)

Symmetry operator: i: -x+2, -y, -z. We located the protons on nitrogen from Fourier difference maps and acknowledge that this leads to abnormally short N-H distances but we favour this approach over that of using computationally generated positions.

Conclusions

In this contribution we have shown that, while simple exchange of both chlorides in *trans*-[$PdCl_2(H_2NBz)_2$] (**1**) for the related mono-anionic (X) N-heterocyclic saccharinate, thiosaccharinate and benzisothiozolinate ligands in all cases affords the expected palladium(II) complexes [$PdX_2(H_2NBz)_2$]. The molecular structure of the product is, however, sensitive to the nature of the incoming ligand with products *trans*-[Pd(N-sac)$_2$($H_2NBz)_2$] (**2**), *trans*-[Pd(S-tsac)$_2$($H_2NBz)_2$] (**3**) and *cis*-[Pd(N-bit)$_2$($H_2NBz)_2$] (**4**) resulting respectively. Formation of N-coordinated saccharinate and S-bound thiosaccharinate ligands to the same metal fragments has been previously noted [**5d**] and likely results from a preference of Pd(II) to bind to a soft sulfur centre when available. Palladium(II) bis(benzisothiozolinate) complexes are far less common [**7,8**] but the three crystallographically characterised examples all contain a *cis* arrangement of benzisothiozolinate ligands. In [Pd(N-bit)$_2$($H_2NBz)_2$] (**4**) this is the first time that this *cis* arrangement has not been imposed by the presence of a chelating co-ligand and the preferential precipitation of *cis*-**4** over its *trans* isomer (which may be initially formed) may result from the ability of the *cis* complex to form strong intermolecular hydrogen bonds with a neighbor thus favoring crystallization of this isomer.

Supplementary information

CCDC 1503153 contains the supplementary crystallographic data for **4**. These data can be obtained free of charge from The Cambridge Crystallographic Data Center via www.ccdc.cam.ac.uk/datarequest/cif

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Table 1. Crystallographic data for *cis*-$[Pd(bit)_2(H_2NBz)_2]$ (**4**)	
Empirical formula	$C_{28}\ H_{26}\ N_4\ O_2\ Pd\ S_2$
Formula weight	621.05
Temperature	200(2) K
Wavelength	0.71073 Å
Crystal system, space group	monoclinic, $P2_1/c$
Unit cell dimensions	a = 9.8581(4) Å, $\alpha = 90^o$
	b = 23.7295(8) Å, $\beta = 102.856(3)^o$
	c = 11.6318(5) Å, $\gamma = 90^o$
Volume	2652.79(18) $Å^3$
Z, Calculated density	4, 1.555 mg/m^3
Absorption coefficient	0.890 mm^{-1}
F(000)	1264
Crystal size	0.10 x 0.10 x 0.10 mm
Theta range for data collection	1.72 to 29.30°
Limiting indices	-13<=h<=13, -32<=k<=32, -15<=l<=14
Reflections collected / unique	19921 / 7115 [R(int) = 0.0597]
Completeness to theta = 25.00	97.9 %
Max. and min. transmission	0.9162 and 0.9162
Refinement method	Full-matrix least-squares on F^2
Data / restraints / parameters	7115 / 0 / 350
Goodness-of-fit on F^2	1.052
Final R indices [I>2sigma(I)]	$R_1 = 0.0377$, $wR_2 = 0.0732$
R indices (all data)	$R_1 = 0.0696$, $wR_2 = 0.0851$
Largest diff. peak and hole	0.605 and -0.628 $e.A^{-3}$