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ARTICLE

A Short, Versatile Route Towards Benzothiadiazinyl Radicals

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A family of substituted 1,2,4-benzothiadiazine 1-chlorides have been prepared by treatment of *N*-arylamidines in neat thionyl chloride at reflux. The S(IV) 1-chlorides are readily reduced under mild conditions to persistent 1,2,4-benzothiadiazinyl radicals which have been characterised by EPR spectroscopy and cyclic voltammetry. Crystallographic studies on isolated radicals indicate that the radicals dimerise *via* pancake bonding in the solid-state, resulting in spin-pairing and net diamagnetism.

Introduction

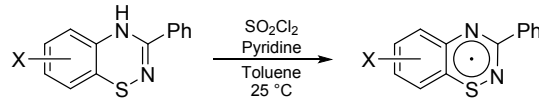
Stable molecular radical species have long been targeted as potential molecular magnets for future magnetic storage applications, and as test-beds for fundamental studies into magnetic communication.¹ Several metal-free radical families² have been explored which exhibit divergent properties from metal centred species. These are of particular interest because, in addition to their magnetic properties, many exhibit a unique form of bonding wherein they dimerise *via* π - π interactions (termed “pancake bonding”) rather than formation of simple $2c2e^-$ bonds.³ These interactions are key to dictating both magnetic and electronic properties of organic systems, and represent a versatile and important motif in supramolecular recognition.⁴

This pancake bonding mode is accessible because many of the “organic” radical families known utilise π -delocalisation, typically onto electronegative elements or extended systems, to achieve radical stability, as exemplified by the verdazyl⁵ (N centred radicals), nitroxyl⁶ (N-O centred radicals), and dithiazolyl, and dithiadiazolyl (S-N centred radicals) systems.⁷ The five-membered ring S-N radicals provide a particularly rich vein of potential radical systems due to their amenability to isoelectronic substitutions and modular synthesis. For example, the 1,2,3,5-dithiadiazolyl class incorporates a substituent at the 4-position which is not directly conjugated into the radical heterocycle and whose influence on properties is therefore limited to steric and inductive effects.⁸ However, comparatively few S-N radicals based on six-membered rings are known, and studies on the benzo-1,2,4-thiadiazinyl class are sparse, with

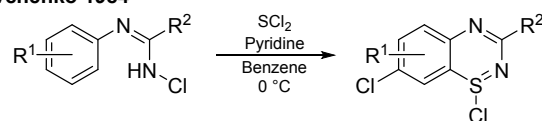
only two examples crystallised to date;⁹ some mesophases containing these ring systems have also been characterised.¹⁰

Scheme 1. Reported synthetic routes towards 1,2,4-benzothiadiazines and related systems.

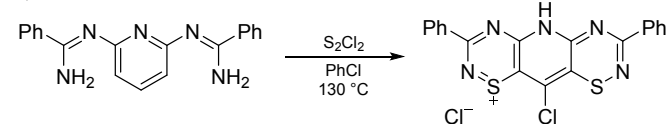
a) Kaszynski 2004



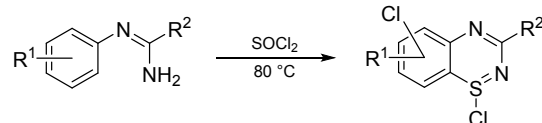
b) Levchenko 1984



c) Oakley 2005



d) This work



The benzo-1,2,4-thiadiazine heterocyclic system is accessible in a range of oxidation states from S(II) to S(VI), and incorporation of 2-pyridyl moieties at the pendant 3 position gives rise to chelating ligands for which first row transition metal¹¹ and Ir(III)¹² complexes are known. The S(III) radicals (see **Scheme 1a**) are accessible through one-electron oxidation of the S(II) systems,⁹ or more conveniently by one-electron reduction of S(IV) precursors.¹³ Work by Oakley has allowed the isolation of a range of mixed valent S(II)/S(III) thiadiazine based systems; in these cases, an extended π -system containing an additional S-N heterocycle is used to provide stability.^{14,15} Overall, these systems are comparatively unexplored and offer great potential as molecular radical building blocks as the substituents at the 3 position are not directly conjugated into the singly occupied

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Electronic Supplementary Information (ESI) available: [details of any supplementary information available should be included here]. See DOI: 10.1039/x0xx00000x



molecular orbital (SOMO), analogous to the dithiadiazoly radicals. The radical properties can therefore in principle be tuned as desired by chemical modification of both the benzo-fused ring, affecting the radical electronic structure, and the 3-substituent whose modification is expected to affect steric demand and packing interactions predominantly.

The expansion of this class of radicals is hampered by the lack of general synthetic methodologies available. The radicals themselves may be formed by either $1e^-$ oxidation¹⁶ or reduction of appropriate closed shell precursors,¹⁴ but few synthetic routes towards these precursors are known and these are not particularly general, (see **Scheme 1**) often involving long, multi-step syntheses. Indeed, for the five radicals reported by Kazsynski in 2004, five bespoke synthetic routes were required.⁹ The three points in common across these synthetic

routes are the presence of an amidine intermediate to provide the NCN fragment of the heterocyclic ring, ring-closure effected by nucleophilic attack on an S(IV) species which is in turn generated by *in situ* oxidation of a S(II) precursor, and halogenation of the benzo-fused ring to stabilise the resultant radicals and enable their isolation. This last point introduces an additional synthetic consideration, necessitating pre-halogenated precursors, or exhaustive chlorination after ring-closure using Cl_2 . In Levchenko's original 1984 work,¹⁷ SCl_2 acted simultaneously as the sulphur source, oxidising agent, and chlorinating agent to give benzothiadiazine 1-chlorides with varying degrees of chlorination about the benzo-fused ring, followed by direct chlorination with Cl_2 , whilst Oakley's mixed S(II)/(IV) parent species use S_2Cl_2 in the same role.¹⁴

Entry	N-Aryl-Amidine	Yield (%)	Benzo-1,2,4-thiadiazine 1-chloride	Yield (%)	Entry	N-Aryl-Amidine	Yield (%)	Benzo-1,2,4-thiadiazine 1-chloride	Yield (%)
a		90		71	i		77		81
b		71		61	j		74		63
c		87		76	k		88		^a
d		79		31	l		85		87
e		78		40	m		71		0 ^b
f		66		46	n		72		0 ^b
g		63		46	o		68		0 ^b
h		66		53	p		41		0 ^b

Table 1. Substituted amidines (**1**) and corresponding 1,2,4-benzothiadiazine 1-chlorides (**2**) after refluxing with SOCl_2 . ^a Mixture of species. ^b Target product not obtained.



Recognising this, we sought a simple, direct route to the required heterocyclic ring starting from the easily handled and cheap S(IV) source, thionyl chloride. We report here the development of a SOCl₂-based route towards 1,2,4-benzothiadiazine 1-chlorides and studies on the derived S(III) radicals.

Results and Discussion

Synthesis

Initial studies on the reaction of the prototypical *N*-phenylbenzamidine, **1a**, with stoichiometric SOCl₂ were unsuccessful when performed in DCM, THF, and toluene, in each case producing a copious quantity of **1a**·HCl as a colourless solid. Addition of stoichiometric base (pyridine, Et₃N and ⁱPr₂EtN) did not improve matters and, for the trialkylamines, led to a violent side reaction which is likely due to oxidation of the trialkylamine by SOCl₂.¹⁸ It was reasoned that a polar, high-boiling solvent would (i) retain the protonated amidine in solution and (ii) permit the HCl formed to be driven off at elevated temperatures, driving the desired cyclisation process. As thionyl chloride itself fulfils these criteria ($\mu = 1.44$ D, bp = 74.6 °C), the reaction of **1a** in neat SOCl₂ was attempted. After reflux overnight, the reaction mixture was deep orange, characteristic of benzothiadiazine 1-chlorides. Layering the reaction mixture with ⁿhexane, followed by slow diffusion, led to isolation of analytically pure, crystalline **2a** in 71% yield. Reaction of the related amidines **1b-j** in refluxing SOCl₂ similarly led to the isolation of benzothiadiazine 1-chlorides **2b-j** in moderate to good yields (31 – 81%) as yellow to red crystalline solids (Table 1). Notably, partial chlorination of both the benzo ring and any methyl substituents on the benzo ring occurs during all these reactions. However, the presence of excess SOCl₂ led to a single product in nearly all cases. For **1d**, the chloride salt **2d** selectively precipitated from solution in 31% yield, but slow cooling of the mother liquor afforded orange needles which comprised a mixture of the isomeric compound **2s'** and the dichloromethyl derivative **2s''** (Figure 1b) in a ca. 44:3 ratio based on crystallographic refinement, suggesting the potential for further chlorination when using extended reaction times. Attempts to prepare **2k** yielded a yellow solid that was practically insoluble in most organic solvents and only poorly soluble in SOCl₂. ¹H NMR analysis was consistent with the formation of the expected heterocycle and partial chlorination at the 6 position (H:Cl = 3:2). Prolonged reflux of **2k** in the presence of excess SO₂Cl₂ and larger volumes of SOCl₂ only slightly increased the degree of chlorination (H:Cl = 2:3), possibly due to the poor solubility of the salt, whilst shorter reflux times afforded other unidentified species. Attempts to deprotonate compound **2k** with basic amines (pyridine, Et₃N, DABCO) resulted in the immediate formation of a dark blue, EPR active solution from which poor quality crystals of the neutral radical (H:Cl = 4:1) could be grown (see ESI for details), indicating that these species are highly oxidising (*vide infra*). Selective chlorination of the benzo-fused ring is reminiscent of Levchenko's reaction of *N*-chloroamidines with SCl₂¹⁷ in which

the position *para* to the amidine nitrogen was always chlorinated. Similarly 1,2,3-dithiazolium salts prepared by Herz from 2-mercaptoanilines and SOCl₂ also undergo chlorination *para* to the amino group.^{19,20} For these benzo-fused 1,2,4-thiadiazolium salts, chlorination of the benzo ring occurred at all positions except the 8-position and chlorination of methyl groups on the benzo ring typically afforded selectively the chloromethyl derivatives unless the methyl group was at the 8 position where no chlorination was observed. The selectivity leading to retention of the C-H at C(8) is attributed to the combination of successive deactivation by chlorination elsewhere and conjugation into the larger heterocycle, lowering the HOMO energies, directed by the location of C(8) *meta* to the *o,p*-directing amidine group and *ortho* to the deactivating S(IV) functionality. This is consistent with the exhaustive chlorination reported by Levchenko and Kaszynski which required direct treatment with excess Cl₂.^{9,17} In contrast to the benzo substituents, phenyl and tolyl groups at the 3-position generally appear resistant to chlorination under these conditions. However, activated aryl groups appear to promote chlorination of the 3-aryl group. For example, while **2i** formed cleanly, reaction of **1j** with SOCl₂ formed **2j** selectively under short reaction times but a mixture comprising **2j** and up to ~45% of **2q** (Figure 1a) was observed as a by-product under extended reaction times. Despite this, attempts to produce pure **2q** under extended reaction times up to 72 hours was unsuccessful. Attempted purification of **2q** by recrystallisation from thionyl chloride containing SO₂Cl₂ as stabiliser instead allowed the isolation of a few crystals of **2r** with additional chlorination of the 6-chloromethyl group. Nevertheless, **2q** was fully characterised by ¹H and ¹³C{¹H} NMR spectroscopy. In the case of **1l**, the basic nature of the 3-(4'-pyridyl) group led to isolation of the hydrochloride salt [**2l**·H][HCl₂]. No crystalline products could be isolated for the reactions of **1m-n** with SOCl₂. Since chlorination occurs *para* to the amidine functionality prior to, or at the same time as, ring-closure, it is believed that the presence of a methyl-group at this position inhibits the formation of the desired 1,2,4-benzothiadiazine 1-chloride. Attempts to prepare **2o** were unsuccessful; yielding a dark brown insoluble solid assigned as a mixture of partially chlorinated species by ¹H NMR spectroscopy. A low yield of pale peach solid was obtained during the attempted synthesis of **2p**; a few, low quality colourless crystals, grown by recrystallisation of the crude solid from boiling SOCl₂, were identified as doubly *ortho*-chlorinated **1p**·HCl by SCXRD but were only a component of a complex mixture as seen by ¹H NMR.

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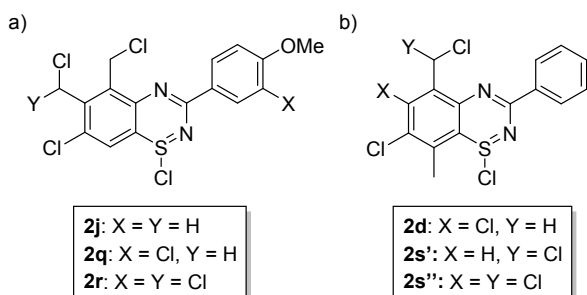
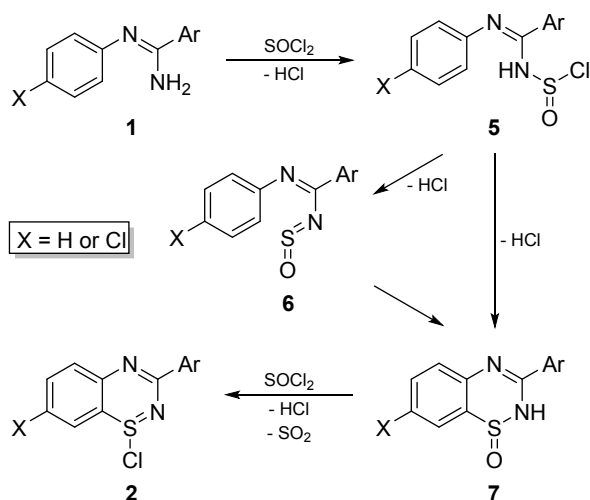


Figure 1. Partially chlorinated 1,2,4-benzothiadiazine 1-chlorides derived from **1j** and **1d**.

The patterns of partial chlorination observed imply that chlorination is rapid at the 5 and 7 positions, slow at the 6 position, and does not occur for the 8 position. The fact that no products could be isolated when substituents were present *para* to the amidine nitrogen (the pro-7 position) suggests that chlorination at this position is rapid, as seen for Herz reactions,^{19,20} producing highly reactive intermediates leading to uncontrolled further reactivity and decomposition. The isolation of partially chlorinated **1p**.HCl suggests that chlorination of the aryl ring is more rapid than ring-closure and that chlorination at both positions *ortho* to the amidine nitrogen inhibits the formation of the fused-ring. This would indicate that the rates of chlorination and ring-closure are competitive for these systems. Independent synthesis of **1t**, *N*-(4-chlorophenyl)-benzamidine, and reaction in SOCl_2 led cleanly to **2a**, supporting this species as an intermediate in ring closure. From these results, it is not possible to fully determine the mechanism, but it is plausible that it occurs by formation of an aminosulphonylchloride, **5**, (Scheme 2) and subsequent electrophilic aromatic substitution, either directly or *via* an intermediate *N*-sulfinylamine, **6**,²¹ followed by deoxygenation of an *S*(IV) 1-oxide, **7**, by reaction with a further equivalent of SOCl_2 to give the benzothiadiazine 1-chloride, **2**, which may then undergo additional chlorination at carbon, as shown in Scheme 2.



Scheme 2. Proposed cyclisation pathway.

Crystallographic Discussion

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Single crystals suitable for X-ray diffraction studies were obtained for all derivatives except **2e**, either by layering the reaction mixture with dry ⁿhexane or slow cooling of a saturated thionyl chloride solution. The benzothiadiazine 1-chlorides all adopt solid state structures distorted from planarity, exemplified by **2a** (Figure 2). The heterocyclic ring is not flat but instead bent along the 1,4 *S*...*N* axis, with angles between the two halves of the ring ranging from 7.11° to 15.10° (see Table 2). These nevertheless show on average less deviation from planarity than the related *S*(II) benzothiadiazines,²² likely due to the increased delocalisation in the formally 10 π benzothiadiazine 1-chlorides vs the 12 π anti-aromatic *S*(II) benzothiadiazines. The pendant aryl rings are not coplanar with the heterocycle, showing torsion angles in the range of 0.8(6) to 30.09(3)°; the torsion angles are largest for **2g** and **2h** with *ortho*-methyl substituents on the pendant ring. Examination of the *S*-Cl bonds is informative - the bond lengths range from 2.221(2) Å to 2.3412(7) Å, considerably longer than the 2.072 Å expected for a formal *S*-Cl single bond,²³ and comparable to the range seen for thiatriazines (2.283-2.357 Å).^{24,25} In addition to this, the transannular *N*-*S*-Cl angles are in a narrow range of 99.84(5)° to 106.80(5)°, consistent with a rigid and well defined bonding environment. These are therefore elongated and weakened covalent bonds, in contrast to the ionic character seen for dithiadiazolyl chlorides where the *S*-Cl contacts are much longer at 2.906-2.962 Å,²⁶ albeit still within the sum of the van der Waals radii, 3.55 Å.²⁷ This argues for a comparatively less stable cation following halide abstraction arising from poorer π -delocalisation about the heterocycle.

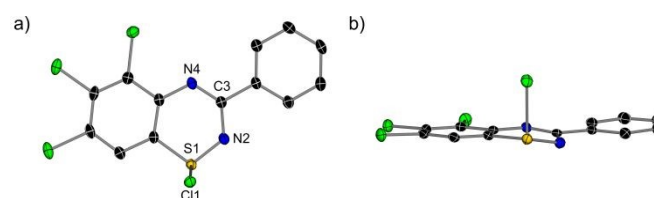
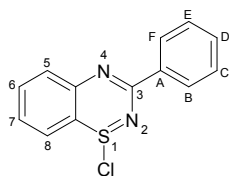


Figure 2. Molecular structure of **2a**. Thermal ellipsoids shown at 50% probability. Hydrogen atoms omitted for clarity. a) Top-down view; b) Side view.





	Deviation from Planarity / °	Torsion Angles / °		S-Cl Bond Length / Å	S-Cl Angle / °
		N ₂ -C ₃ -C ₄ -C ₅	N ₄ -C ₃ -C ₄ -C ₅		
2a	11.74	4.6(6)	0.8(6)	2.221(2)	100.79(8)
	8.17	13.7(6)	9.9(6)	2.306(1)	106.24(7)
2b	11.44	11.90(3)	9.05(3)	2.2811(8)	102.58(4)
2c	7.32	6.65(3)	4.18(3)	2.3412(7)	106.11(4)
2d-α	10.99	8.64(4)	6.16(4)	2.2974(9)	102.32(5)
2d-β	7.11	6.1(8)	1.7(8)	2.261(2)	107.0(1)
2f	9.61	3.83(2)	0.89(2)	2.2759(6)	104.06(3)
2g	8.26	19.80(1)	16.08(1)	2.309(2)	106.23(1)
	13.78	10.51(1)	9.96(1)	2.229(3)	99.97(1)
2h	15.10	30.90(3)	28.91(3)	2.2689(9)	99.84(5)
2i	7.20	9.59(3)	5.89(3)	2.3205(8)	106.80(5)
2j	8.25	8.25(5)	5.10(5)	2.279(1)	105.33(7)
[2l.H][HCl₂]	13.46	20.56(2)	18.15(2)	2.238(4)	100.25(2)
2r	9.46	9.98(1)	8.45(1)	2.264(4)	103.42(2)
2s	9.98	7.01(9)	4.03(9)	2.287(2)	104.89(1)

Table 2. Selected crystallographic parameters for the studied 1,2,4-benzothiadiazine 1-chlorides.

The benzothiadiazine 1-chlorides are chiral systems and, where partial chlorination of methyl groups occurs, the chloromethyl fragment may therefore lie either *syn* or *anti* to the S-Cl bond. In the case of **2c**, **2h**, **2j** and **2r**, the crystal structure shows both C-Cl bonds lying nearly co-parallel, *anti* to the S-Cl bond, whereas for **[2l.H][HCl₂]**, the two chloromethyl groups sit *anti* to one another. In contrast, **2d** is polymorphic with the α -phase crystallising with the E-Cl bonds *anti* to one another whilst the β -phase adopts a *syn* configuration. Although not possessing a CH₂Cl group, the OMe group of **2f** is disordered equally above and below the plane of the ring relative to the S-Cl bond. In all cases, no splitting of the methylene protons is evident in the ¹H NMR spectrum, consistent with rapid rotation and/or inversion about sulphur on the NMR timescale in solution.

Anion metathesis

Anion metathesis reactions are commonly employed to improve the solubility of S(IV) salts prior to reduction to form neutral radicals. With this in mind, some simple tests of the amenability of **2a** towards halide abstraction were performed. Reaction of **2a** with GaCl₃ in DCM resulted in the immediate formation of **[4a][GaCl₄]** as a dark purple solution whose identity was unambiguously confirmed by growth of single crystals by slow diffusion of ⁿhexane into the reaction mixture (**Figure 3**).

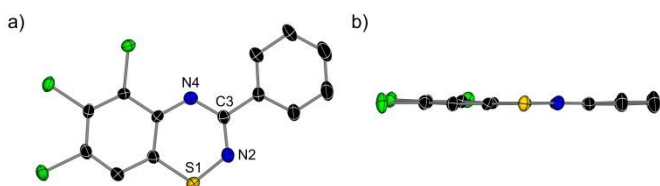


Figure 3. Molecular structure of **[4a][GaCl₄]**. Thermal ellipsoids shown at 50% probability. Hydrogen atoms and counter-anion omitted for clarity. a) Top-down view; b) Side view.

The abstraction results in an almost completely planar system forming, with minimal *trans*-annular bend (0.51°) and a torsion angle of only 2.57(6)° between the pendant phenyl ring and heterocycle. The S-C_{Ar} bond length in **[4a][GaCl₄]** is considerably shorter than that of **2a** (1.683(4) Å vs 1.733(2) Å), with minor alterations in other bond lengths, which coupled with the more planar structure argues for increased delocalization within the ring which is formally a 6π aromatic system (10π when considering the benzo-fused substituent). Two long cation-anion interactions of 3.341(1) Å and 3.463(1) Å (*cf.* 2.2572(7) Å for the S-Cl bond in **2a**) are observed between sulphur and the chlorine atoms of two adjacent GaCl₄⁻ anions, confirming the structure as ionic.

Treatment of **2a** with NaBAR^{Cl} in DCM resulted in a deep blue, EPR active solution consistent with the formation of the neutral S(III) radical, **3a**. Taken together, these results show that whilst benzothiadiazinyl cations are accessible, they are highly reactive and easily reduced to the radical such that judicious choice of weakly coordinating anions is required to allow their isolation.

1,2,4-Benzothiadiazinyl Radicals

Benzothiadiazinyl radicals have been accessed *via* one electron oxidation of benzothiadiazines,⁹ but this requires considerable optimisation for each derivative. An alternative approach is one electron reduction of the benzothiadiazine 1-chloride salts. In our hands, we found that the 1,2,4-benzothiadiazine 1-chlorides (**2a-j**) were easily reduced to the corresponding radicals with both Ph₃Sb and Fe(η⁵-C₅H₅)₂ but attempts to remove the by-products and isolate the pure radicals from these reaction mixtures failed. Preliminary small-scale studies utilising Ph₃P as a milder reducing agent in a variety of common organic solvents were promising, immediately yielding dark green, EPR active solutions, and clearly showing the formation of Ph₃PCl₂ (THF, toluene; δ -46 ppm) or [Ph₃PCl]Cl (DCM, MeCN; δ +60 ppm) by ³¹P NMR spectroscopy. Solutions of the radicals generated in DCM, THF, and toluene swiftly discoloured to give yellow-brown suspensions, whilst the radicals were only poorly soluble in MeCN. The green solutions of these benzothiadiazinyl radicals were observed to become a rich blue colour on cooling in liquid nitrogen which is believed to originate from a monomer-dimer equilibrium (*vide infra*), with the formation of closed-shell dimers being favoured at low temperatures.

EPR Spectroscopy

Although concentrated solutions of these radicals decolourise in minutes, dilute solutions proved sufficiently robust in dry, degassed toluene to allow EPR studies to be performed. This indicates that the short half-life in solution is a result of kinetic instability and subsequent reactivity rather than inherent thermodynamic instability. This is consistent with the thermal stability of the more highly chlorinated analogues reported by Kaszynski which were stable to sublimation at elevated temperature but decomposed over hours in solution.⁹ They are therefore considerably more reactive than their valence isoelectronic analogues, the Blatter radicals.²⁸ EPR spectra of radicals **3a-j**, generated *in situ* by reduction with Ph₃P, were



recorded in toluene at ambient temperature on a continuous wave X-band EPR spectrometer. The EPR spectra for the 1,2,4-benzothiadiazinyl radicals are all similar, exhibiting slightly asymmetric 1:2:3:2:1 quintets arising from coupling to two similar but non-identical nitrogen atoms within the heterocyclic ring. Significant line-broadening was observed in all cases due to chlorination of the fused ring; since both ^{35}Cl and ^{37}Cl are quadrupolar nuclei with $I = 3/2$, the spectra consist of multiple superimposed spectra for each isotopologue, resulting in net loss of resolution and increased line-broadening, meaning all features beyond nitrogen hyperfine coupling are lost. The EPR spectrum of **3a** is shown in **Figure 4** and the g -values and hyperfine coupling constants for radicals **3a-j** are summarized in **Table 3**.

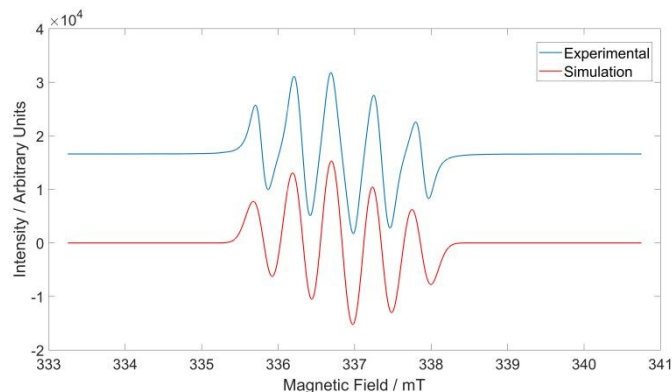


Figure 4. EPR spectra of **3a**.

	g -value	Line-Width / MHz	a_{N2} / MHz	a_{N4} / MHz
3a	2.0037	0.27	15.66	13.13
3b	2.0035	0.26	15.64	13.51
3c	2.0034	0.31	15.80	13.60
3d	2.0043	0.31	15.10	13.10
3e	2.0046	0.37	14.69	13.30
3f	2.0045	0.33	14.56	13.23
3g	2.0046	0.27	15.88	13.35
3h	2.0045	0.34	15.61	13.71
3i	2.0041	0.29	15.82	12.89
3j	2.0041	0.38	14.73	13.98

Table 3. EPR spectral parameters for radicals **3a-j**. Assignment of hyperfine coupling to N2 and N4 on the basis of DFT calculations; see ESI for details.

The SOMO geometry of **3a** (**Figure 5**), calculated at the UB3LYP/cc-pVDZ level using a UB3LYP/6-31G optimised geometry, reveals that it is largely independent of the substituents around the benzo-fused and pendant aryl rings. This is supported by the similarity of g -values and ^{14}N hyperfine coupling constants observed across the range of 1,2,4-benzothiadiazinyl radicals indicating that the electronic structure is only slightly perturbed by substitution at these positions.

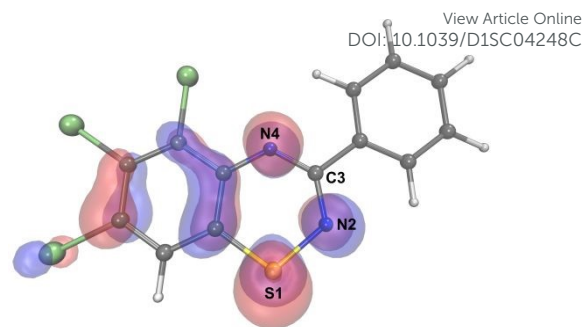
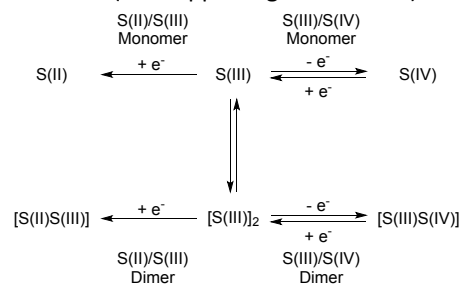


Figure 5. Calculated SOMO of **3a** (isovalue = 0.04).

Electrochemical Studies

Although all radicals could be generated *in situ*, pure samples could only be isolated for a minority of the radicals (*vide infra*). As a result of this and their poor long-term stability in solution, cyclic voltammetry studies were performed on the chlorides **2a-j** in dry, degassed DCM. The 1,2,4-benzothiadiazine 1-chlorides all displayed similar electrochemical behaviour, and the cyclic voltammogram of **2a** is presented as typical of this series (**Figure 6**). The two, often-overlapping, reversible processes at positive potentials (*vs* Fc/Fc^+) are assigned to S(III)/S(IV) couples, whilst the irreversible process at negative potential is attributed to the S(II)/S(III) couple. The two S(III)/S(IV) waves suggest the presence of radical monomers and dimers in solution, as shown in **Scheme 3**. Similar monomer-dimer equilibria have also been observed for 6-membered thiaziazines and independently confirmed by quantitative EPR measurements.²⁹ Further support for a monomer-dimer equilibrium was observed *via* variable concentration studies - at low concentrations the monomer is favoured whilst at high concentrations the cyclic voltammogram is dominated by signals for the dimer. These studies permit initial assignment of the two S(III)/S(IV) redox couples and show that the one electron reduction potential of the monomer is positive relative to the dimer.²⁹ In addition, the S(II)/S(III) reduction wave splits into two well resolved peaks at low concentrations (see Supporting Information).



Scheme 3. Proposed redox events.



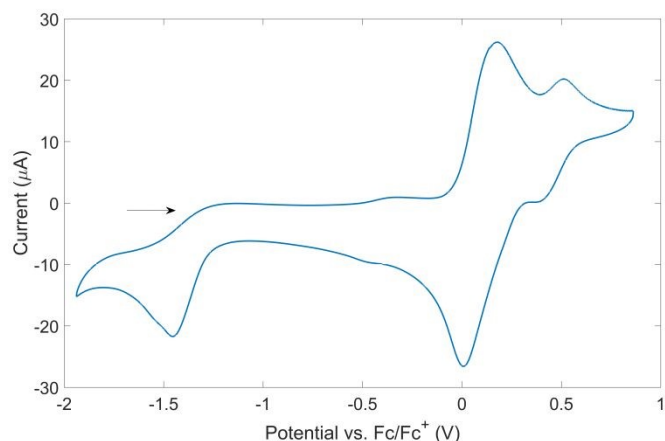


Figure 6. Cyclic voltammogram of **2a** in DCM with $[t\text{Bu}_4\text{N}][\text{PF}_6]$ supporting electrolyte.

The irreversibility of the redox event at negative potentials is attributed to a rapid E_rC (a reversible electron transfer followed by an irreversible chemical reaction) comproportionation reaction between the electro-generated S(II) anion and the bulk S(IV) species **2a**. This has been observed for several S-N systems.²⁹ No evidence of *quasi*-reversibility was observed, even at scan rates up to 2 Vs^{-1} , indicating that the comproportionation reaction is extremely rapid. The variable scan-rate data nonetheless show that the S(III)/S(IV) processes for both radical monomer and dimer are reversible electron transfer processes involving a freely diffusing redox species according to the Randles-Sevcik equation,³⁰ and confirm that the analyte is not adsorbed to the electrode surface (see Supporting Information).

The $E_{1/2}$ potentials for the S(III)/S(IV) couple follow the expected trend (see **Table 4**), with 1,2,4-benzothiadiazine 1-chlorides bearing electron-withdrawing groups such as **2a**, **2g** and **2i** being more easily reduced than those with electron-donating groups such as **2b-d**. The same trends were also observed for the S(II)/S(III) couple. The substituents on the benzo-fused ring were found to have a greater influence on the electrochemical behaviour and redox potentials compared to substituents on the pendant aryl ring. This is in good agreement with DFT studies which indicate that there is negligible delocalisation of the unpaired electron onto the pendant aryl ring but significant π -delocalisation across the benzo-fused and heterocyclic rings. There are, however, only minor changes observed across the $E_{1/2}$ potentials for substituted benzothiadiazine 1-chlorides.

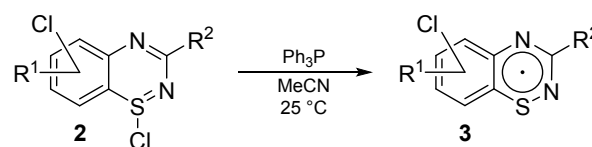
	S(II)/S(III)		S(III)/S(IV) (Dimer)		S(III)/S(IV) (Monomer)		
	E_{red}/V	E_{ox}/V	E_{red}/V	E_{ox}/V	E_{red}/V	E_{ox}/V	$E_{1/2}/V$
2a	-1.457	0.013	0.179	0.096	0.371	0.518	0.444
2b	-1.511	-0.112	0.084	-0.014	/	0.449	/
2c	-1.649	-0.094	0.054	-0.020	/	0.501	/
2d	-1.654	-0.084	0.036	-0.024	/	0.441	/
2e	-1.473	-0.027	0.142	0.057	0.336	0.469	0.403
2f	-1.557	-0.083	0.111	0.014	0.379	0.508	0.443
2g	-1.446	-0.011	0.226	0.108	0.400	0.481	0.440
2h	-1.599	-0.040	0.123	0.041	/	0.479	/
2i	-1.450	0.028	0.144	0.086	0.300	0.397	0.349
2j	-1.620	-0.071	0.035	-0.018	/	0.558	/

Table 4. Redox potentials of 1,2,4-benzothiadiazine 1-chlorides **2a-j**. Potentials are referenced against the Fc/Fc⁺ couple.

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Solid State Studies of 1,2,4-Benzothiadiazinyl Radicals

Isolated 1,2,4-benzothiadiazinyl radicals, **3a-j**, were prepared by treatment of the S(IV) 1-chlorides with half a molar equivalent of Ph_3P in thoroughly degassed and anhydrous MeCN (**Scheme 4**). Exploiting the poor solubility of radicals in MeCN, the radicals formed cleanly and rapidly, and precipitated as dark purple-green solids that were subsequently isolated by filtration and washed prior to drying under vacuum. The stability is significantly enhanced in the solid state, with samples giving clean elemental analysis and remaining spin active even after storage for several weeks at room temperature in an argon glove box. In contrast, the washings were found to swiftly discolour, even when stored at $-20 \text{ }^\circ\text{C}$. The growth of crystals suitable for SCXRD analysis was severely hampered by the poor stability of the 1,2,4-benzothiadiazinyl radicals in solution, with several approaches (slow cooling, vapour diffusion) and solvents (DCM, THF, toluene, pyridine, *n*-hexane, Et_2O) giving only rapid discolouration and formation of yellow-brown precipitates. Ultimately, single crystals suitable for SCXRD analysis of **3a** and **3c** were successfully grown by rapidly mixing the parent S(IV) 1-chloride and reducing agent in DCM and allowing the radical to slowly crystallise out of solution. This method, however, was found to be unreliable in forming X-ray quality crystals and often afforded just micro-crystalline material unsuitable for single crystal diffraction studies. Single crystals of **3e** grew readily from saturated toluene solutions. A few, poor quality crystals of **3k** could be isolated after attempting to dissolve **2k** in pyridine.



Scheme 4. Synthesis of 1,2,4-benzothiadiazinyl radicals **3a-j**.

The discussion herein focuses on the thiadiazinyl ring (see Supporting Information for full structures). The structurally characterised 1,2,4-benzothiadiazinyl radicals are notably closer to planarity than the S(IV) 1-chlorides; the deviation from planarity for the heterocyclic ring ranges from 1.22° to 5.78° (*cf.* 7.11 - 15.10° for **2a-s**), although the torsion angles for the pendant aryl ring are comparable $0.3(3)^\circ$ to $9.6(5)^\circ$ (*cf.* $0.89(2)$ - $11.90(3)^\circ$ for **2a-f**). Crystallisation of **3a** allows direct comparison of changes in heterocyclic ring structure between **2a**, **3a**, and **[4a]GaCl₄**, as shown in **Table 5**. For **3c**, the two CH_2Cl groups adopt an *anti* configuration, in contrast to the *syn* configuration observed in **2c**. Unlike Kaszynski's perfluoro- and perchloro-radicals which are monomeric in the solid state,⁹ radicals **3a**, **3c**, **3e**, and **3k** all crystallise as pancake dimers featuring short contacts between the heterocyclic rings.



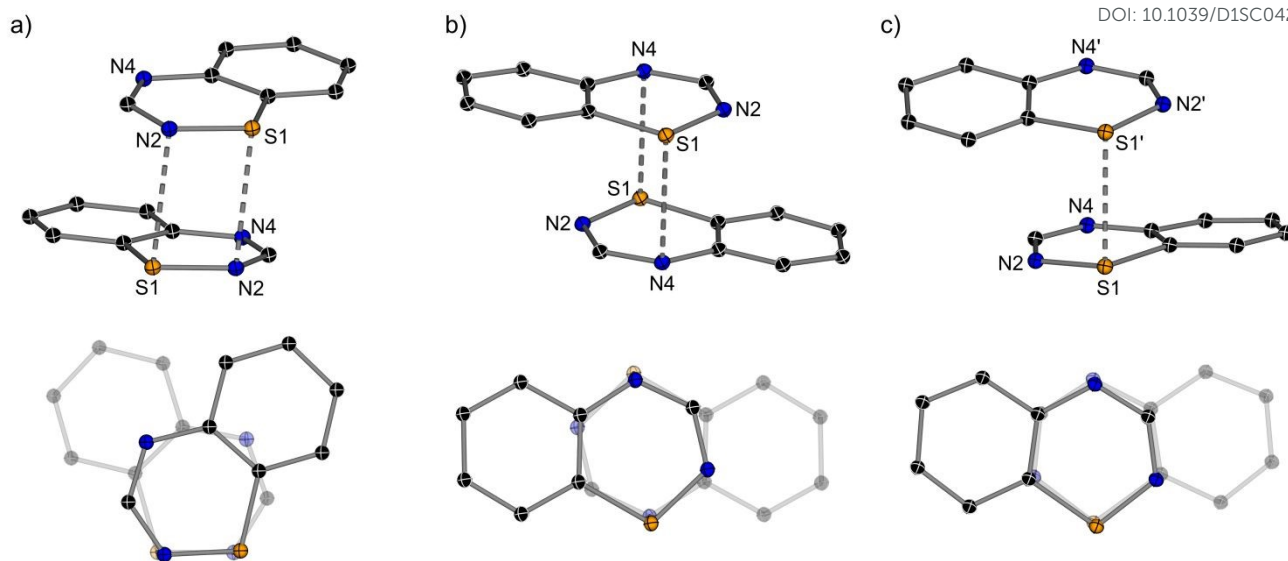


Figure 7. Dimerisation motifs, shown as side-on and top-down views, observed for 1,2,4-benzothiadiazinyl radicals: a) **3a** - suprafacial; b) **3c** and **3k** - *trans*-antarafacial; c) **3e** - *trans*-suprafacial. The pendant aryl ring and substituents on the benzo-fused ring have been omitted for clarity.

Bond	Bond Length / Å		
	2a	3a	[4a] [†]
S1-N2	1.564(2)	1.615(2)	1.574(4)
N2-C3	1.382(2)	1.349(5)	1.347(5)
C3-N4	1.307(2)	1.323(5)	1.339(5)
N4-CX	1.371(3)	1.377(5)	1.324(5)
CX-CY	1.403(3)	1.405(5)	1.442(5)
CY-S1	1.733(2)	1.735(4)	1.683(4)

Table 5. Comparison of heterocyclic bond parameters for **2a**, **3a** and [4a][†].

Three different configurations of dimer (**Figure 7**) were observed for the radicals. This is analogous to other sulphur-nitrogen radicals, especially those based on less delocalised five-membered ring systems such as dithiadiazolyls, for which multiple dimer motifs have been documented.⁸ Radical **3a** dimerises in a twisted, suprafacial motif (**Figure 7a**) with the two molecules related *via* a crystallographic 2-fold rotation axis. The two heterocyclic rings are not quite coplanar, exhibiting an interplanar angle of 7.82° and a pair of crystallographically equivalent S1...N2 contacts at 2.866(4) Å (*cf.* S...N contacts of 3.193(4) Å and 3.213(4) Å respectively for Kazsynski's monomeric perfluoro-analogue). Radicals **3c** and **3k** also adopt dimeric structures with molecules related *via* a crystallographic inversion centre. In this instance, the inversion centre generates a *trans*-antarafacial dimer motifs (**Figure 7b**) featuring two short identical S1...N4 contacts at 3.087(2) Å and 3.010(3) Å for **3c** and **3k** respectively with parallel heterocyclic rings, whilst **3e** dimerised in a *trans*-suprafacial motif (**Figure 7c**) with a single short S1...S1' contact of 2.846(1) Å and a non-parallel arrangement of the heterocyclic rings; the angle between the mean planes of 10.45°.

Computational investigation of these binding modes for the prototypical 3-phenyl benzothiadiazinyl radical found dimerization energies to be -14.3 and -14.8 kcal mol⁻¹ for the *trans*-antarafacial and *trans*-suprafacial motifs respectively, approximately twice those seen for the related dithiadiazolyl system³¹ (see Supporting Information for details). No energy minimum corresponding to a pancake dimer could be located for the suprafacial binding mode; instead this geometry was found to correspond to a transition state between S-N bonded σ -dimers. This discrepancy between gas-phase calculated geometries and solid-state structure for this orientation of interaction suggests that this mode of dimerisation depends on additional, second-sphere interactions in the solid state; this is unsurprising as pancake bonding is known to be sensitive to subtle changes in structure.³

Magnetism

Of the structurally characterised radicals, only **3c** and **3e** could be isolated in sufficient purity for magnetic studies, and their magnetic susceptibilities were measured over the temperature range 2-300 K. These confirm that the radicals are diamagnetic in the solid-state, as expected for spin-paired dimers, with no evidence of any magnetic phase transition at high temperature (see Supporting Information). Broken symmetry calculations^{32,33} have been successfully applied to a variety of heterocyclic sulphur-nitrogen radicals,³⁴⁻³⁶ and were performed to explore the nature of the magnetic properties in **3c** and **3e** (see Supporting Information for details). For both systems, the calculated exchange interaction between the radical pair is strongly anti-ferromagnetic ($J/k = -2629$ and -6795 K respectively), consistent with observed spin pairing, net diamagnetism, and the large calculated dimerisation energies. The interaction between neighbouring radicals along the π -



stack are also anti-ferromagnetic, albeit significantly weaker. This is in stark contrast to the behaviour seen for related benzothiadiazinyl radicals which retain their paramagnetism in the solid state^{9,16} and illustrates the sensitivity of these systems to substituents.

Conclusions

A new, efficient route to 1,2,4-benzothiadiazine 1-chlorides is reported wherein the use of neat SOCl₂ leads to reproducible patterns of chlorination on benzo and methyl substituents. This compact route proceeds directly from easily accessible precursors, with no need for pre-functionalisation of the amidines nor subsequent chlorination. Reduction of the benzothiadiazine 1-chlorides leads to the formation of persistent benzothiadiazinyl radicals which have been characterised *in situ* by EPR spectroscopy and cyclic voltammetry. The existence of solution-state monomer-dimer equilibria is supported by both thermochromism and voltametric measurements. Four of these radicals have been structurally characterized by X-ray diffraction and exhibit three differing modes of $\pi^*-\pi^*$ pancake bonding. Two of these radicals were of sufficient purity for magnetic measurements, which confirm that they are diamagnetic in the solid state between 2-300 K, with pancake bonding quenching the inherent magnetism of the radicals. This is markedly different from the behaviour seen for related perhalogenated systems and shows that there remains much to be explored for this tunable radical scaffold.

Author Contributions

This work was initially undertaken by E.R.C., A.A., and J.M.R. at the University of Cambridge and completed by A.M.B, P.J.S., and E.R.C. at the University of Kent. Project design and oversight was led by J.M.R. and E.R.C. Synthetic work and physical measurements were completed by E.R.C, A.M.B., and P.J.S. with additional crystallographic measurements by A.A. Initial manuscript drafting was completed by A. M. B. with all authors contributing to the final version of the manuscript.

Conflicts of interest

There are no conflicts to declare.

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