Steric C–N bond activation on the dimeric macrocycle \[[\{P(\mu-\text{NR})_2(\mu-\text{NR})_2\}_2 \dagger\]

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Dimeric cyclophosphazanes \[[(\{P(\mu-\text{NR})_2(\mu-\text{NR})_2\}_2) \mid \text{R} = \text{^tBu} (1) and \text{^tPr} (3)\] were oxidized with elemental selenium. During these reactions an unexpected C–N bond cleavage and N–H bond formation occurred. Compound 1 produced \[P_2(\mu-\text{N}^t\text{Bu})_2(\mu-\text{NH})_2\text{Se}_4 (2)\] where three \(^t\text{Bu}\) groups were lost in the form of isobutylene. In contrast, during the oxidation of the less sterically hindered 3, the resulting product, \[P_2(\mu-\text{N}^t\text{Pr})_2(\mu-\text{NH})\text{Se}_4 (4)\], showed only one substituent loss. Theoretical studies confirmed the steric nature of the driving force underlying the different outcomes.

Cyclophosph\[(\mu/\mu)\text{azanes, with a variety of substituents, have been extensively studied over the past decades.}^1\] One of their main features is that they can be found in a myriad of topographical arrangements spanning from simple dimeric units to cage\(^2\) and macrocyclic arrangements.\(^3\)\(^-\)\(^5\) Moreover, the facile functionalization of dimeric rings of the type \[{\text{ClP(\mu-\text{NR})}_2}_{\text{(I)}}\] with a variety of nucleophiles\(^6\) lends these species unprecedented versatility as building blocks towards the synthesis of a wide range of multidentate ligands to transition metals.\(^7\)\(^-\)\(^8\) However, the intrinsic bond lability of these species renders them incapable of withstanding standard reaction conditions required for many simple organic transformations. Furthermore, an increased stability of the P–N framework upon oxidation of the phosphorus center has been reported.\(^9\) For example, Balakrishna et al. reported the synthesis of a series of cyclophosph(v/v)azanes showing enhanced air and moisture stability.\(^10\) These results prompted us to investigate the stability of cyclic cyclophosphazane cage frameworks of the type \[[\{P(\mu-\text{NR})_2(\mu-\text{NR})_2\}_2 › \text{R} = \text{^tBu} (1) and \text{^tPr} (3)\] upon oxidation. Herein, we report the unexpected C–N bond activation that occurs upon oxidation of the dimeric cyclophosphazanes \[[\{P(\mu-\text{N}^t\text{Bu})_2(\mu-\text{N}^t\text{Bu})_2\}_2 (1)\] with elemental Se. Compound 1, was previously synthesized via the condensation reaction between \[{\text{ClP(\mu-\text{N}^t\text{Bu})}_2}_{\text{(II)}}\] and \[{\text{LiN}^t\text{BuP(\mu-\text{N}^t\text{Bu})}_2}_{\text{(III)}}\], and was isolated as co-crystals of \[{\text{LiN}^t\text{BuP(\mu-\text{N}^t\text{Bu})}_2}_{\text{(I)}} \mid \text{R} = \text{^tBu} (1) : \text{^tPr} (3)\] \[= 4.4\]\footnote{Our initial work focused on the purification of 1, which was successfully crystallized in THF at room temperature.} Oxidation of 1 with elemental Se (molar ratio of 1 : 4.4) was performed in refluxing toluene for 72 hours. In situ \[\text{^31}P\{\text{^1H}\}^\text{NMR spectroscopic studies were used to monitor the reaction mixture. The expected fully oxidized and symmetrical product, \[[\{P(\text{Se})(\mu-\text{N}^t\text{Bu})_2(\mu-\text{N}^t\text{Bu})_2\}_2 (2')\] would display a single resonance within the range of \(\delta 0–50\) ppm (cf., 117.2 ppm in 1) consistent with previously reported cyclophosph(v/v)azane selenide derivatives (e.g., \[[P(\text{Se})(\mu-\text{N}^t\text{Bu})_2(1,3-\text{O}2-\text{C}_6\text{H}_4)_2]\] and cis-\[\text{Cy(H)NP(\text{Se})(\mu-\text{NCy})}_2\] \[\delta 32.7\] and \[\delta 35.4\] ppm, respectively.} To our surprise, after 36 hours of reflux, the in situ \[\text{^31}P\{\text{^1H}\}^\text{NMR spectrum showed a neat incipient set of signals composed of two groups of multiple resonances at \(\delta ~24\) and \(\sim 22\) ppm, respectively. In situ \[\text{^31}P\{\text{^1H}\}^\text{NMR spectrum taken after refluxing for a further 36 hours confirmed reaction completion. The reaction mixture was filtered through Celite, and the filtrate dried off under vacuum. Crystallization of the residue in toluene afforded crystals of compound 2 suitable for X-ray crystallographic studies (Scheme 1).}

The X-ray solid state structure of \[P_2(\mu-\text{N}^t\text{Bu})_2(\mu-\text{NH})\text{Se}_4 (2)\] shows a tetrathiolene-oxidized product where three \(^t\text{Bu}\) groups, two on the ring-bridging nitrogen atoms and one on the \(P_2\text{N}_2\) ring, were cleaved off as is shown in Fig. 1.

\text{Scheme 1} Synthesis of compound 2.
The cleavage of N-\textsuperscript{t}Bu bonds resulted in the formation of three new N-H bonds. The \textsuperscript{31}P\textsuperscript{[\textsuperscript{1}H]} NMR spectrum of 2 shows two second order multiplets centred at 1.64 and 2.08 ppm, respectively, corresponding to the two distinct phosphorus environments present. The three remaining \textsuperscript{t}Bu groups in 2 can be observed in the \textsuperscript{1}H NMR spectrum as two singlet resonances with an intensity ratio of 1:2 at 6 1.31 and 1.62 ppm (cf. 1.57 and 1.53 ppm in 1). Furthermore, the \textsuperscript{1}H NMR spectrum displays signals resulting from the newly formed N-H bonds in the form of two broad singlets at 6 5.64 and 4.28 ppm, with an intensity ratio of 2:1, which correspond to the protons on the ring-bridging nitrogen and the P2N2 ring, respectively. This observation is also consistent with typical NH absorptions bands observed at 3192 cm\textsuperscript{-1} in the IR spectrum.

The mean P-Se bond distance in 2, 2.0897(16) Å, is comparable with previously reported analogues (cf. 2.0913 in cis-[\textsuperscript{\textmu}\textsuperscript{N}-\textsuperscript{t}Bu\textsubscript{4}P(SeNC\textsubscript{2}H\textsubscript{4}NMe\textsubscript{2})]). The P-N average bond distances within the P\textsubscript{2}N\textsubscript{2} ring and the P-N bridging groups are 1.700(5) and 1.604(5) Å (cf. 1.717(5), N(3)–P(4) 1.675(5), N(5)–P(2) 1.604(5), N(5)–P(3) 1.613(5), Se(1)–P(1) 2.085(5), Se(2)–P(2) 2.102(6), N(1)–P(1)–Se(1) 121.8(18), N(2)–P(2)–Se(1) 118.6(17), N(6)–P(1)–Se(1) 109.3(17), N(1)–P(1)–Se(2) 115.1(17), N(2)–P(2)–Se(2) 116.3(18), N(5)–P(3)–Se(3) 109.3(18), N(6)–P(4)–Se(4) 107.5(17). Ellipsoids are drawn at 50% probability. H atoms have been omitted for clarity.

In order to gain insights into the reaction mechanism, the reaction was performed in a sealed Young’s tap NMR tube. The in situ \textsuperscript{31}P\textsuperscript{[\textsuperscript{1}H]} NMR spectra show the initial formation of a series of partially oxidized intermediates before the formation of compound 2. During the course of the reaction, the \textsuperscript{1}H NMR spectra show the appearance of two new signals with intensity ratio 1:3 at 6 4.76 and 1.64 ppm respectively, as the only detectable by-product. These two resonances were found to be in accordance with \textsuperscript{1}H NMR spectrum of isobutylene (Fig. 2, compound a).\textsuperscript{14}

Several unsuccessful attempts were made to isolate the partially oxidized transient species (i.e., 1a, 1b and 1c in Scheme 2, see ESI†); however a singlet at ~25 ppm observed during the course of the reaction can be attributed to the fully-oxidized non-cleaved compound 2'. We observed that only once 2' is detected in the \textsuperscript{31}P\textsuperscript{[\textsuperscript{1}H]} NMR spectra the signal set corresponding to isobutylene commence to appear in the \textsuperscript{1}H NMR spectra (see ESI†). To further evaluate the influence of stoichiometry and reaction conditions upon the obtainment of 2 and/or observed intermediates in isolatable form, 1 was oxidized in a 1:2.2 ratio under identical experimental conditions. Further to this, the use of alternative solvents such as THF and ACN was explored under reflux and at room temperature in both 1:4.4 and 1:2.2 molar ratios. In all cases a mixture of compound 2 and the di-oxidized derivative (1b) were observed in the in situ \textsuperscript{31}P\textsuperscript{[\textsuperscript{1}H]} NMR spectrum after prolonged reaction times, suggesting that the cleaved product 2 is the most thermodynamically stable fully-oxidized product.

Since the P\textsubscript{n}N\textsubscript{6} framework contracts upon oxidation (vide supra), we presume that steric strain release is the driving force for the observed C–N bond cleavage. In order to validate this observation, the less sterically hindered dimeric phosphazane \[(\textsuperscript{\textmu}\textsuperscript{N}-\textsuperscript{t}Bu)\textsubscript{2}\textsuperscript{[\textsuperscript{\textmu}\textsuperscript{N}-\textsuperscript{t}Pr]}\] was synthesized. In this context, it is important to note that frameworks of the type \[(\textsuperscript{\textmu}\textsuperscript{N}-\textsuperscript{NR})\textsubscript{2}\textsuperscript{[\textsuperscript{\textmu}\textsuperscript{N}-\textsuperscript{NR}]}\] are thermodynamically unstable with respect of their adamantoyl isomeric forms \[(\textsuperscript{\textmu}\textsuperscript{N}-\textsuperscript{NR})\]. In the case of compound 1, isomerization to an adamantoid structure is prevented by its highly sterically encumbered nature.\textsuperscript{15} Since compound 3 is less thermodynamically

![Fig. 1](https://example.com/f1.png)

**Fig. 1** ORTEP diagram for 2. Selected bond lengths [Å] and angles [deg]: N(1)–P(2) 1.717(5), N(1)–P(1) 1.682(5), N(2)–P(2) 1.723(5), N(2)–P(1) 1.700(5), N(3)–P(3) 1.716(4), N(3)–P(4) 1.675(5), N(5)–P(2) 1.604(5), N(5)–P(3) 1.613(5), Se(1)–P(1) 2.085(5), Se(2)–P(2) 2.102(6), N(1)–P(1)–Se(1) 121.8(18), N(2)–P(2)–Se(1) 118.6(17), N(6)–P(1)–Se(1) 109.3(17), N(1)–P(1)–Se(2) 115.1(17), N(2)–P(2)–Se(2) 116.3(18), N(5)–P(3)–Se(3) 109.3(18), N(6)–P(4)–Se(4) 107.5(17).

![Fig. 2](https://example.com/f2.png)

**Fig. 2** \textsuperscript{1}H NMR spectrum of the reaction of \textsuperscript{\textmu}\textsuperscript{N}-\textsuperscript{t}Bu\textsubscript{4}P\textsubscript{[\textsuperscript{\textmu}\textsuperscript{N}-\textsuperscript{t}Pr]}\ (1) with elemental Se in a sealed NMR tube under reflux in toluene-d\textsubscript{6}.
Furthermore, despite several attempts, it was not possible to decomposes upon isolation prohibiting any further characterization. This was further corroborated by single crystal X-ray diffraction studies that show that one iPr group is almost planar (puckered by 4.7°) and 1H NMR spectra show complex signals at 55.25 and 30.65 ppm, respectively (Fig. 3). Similarly to 2, the mean P–N and P–Se bond distances in 4 are 1.700(10) and 2.075(4) Å, respectively. The symmetric P2N2 rings are almost planar (puckered by 4.7°) and are virtually perpendicular to the macrocyclic plane (ca. 89°). Unfortunately, compound 4 readily decomposes upon isolation prohibiting any further characterization. Furthermore, despite several attempts, it was not possible to achieve the synthesis of analogues to 1 and 3 containing less bulky substituents (e.g., °Pr, Et or Me). However, to rationalize the energetics of the reaction and further elucidate the observed C–N bond cleavage, DFT calculations were performed. An assessment of the relative thermodynamic stabilities the compounds can be obtained by directly comparing the enthalpies of reactants and products that share the same degree of oxidation (i.e., 2 and 4, 3b and 3c). Our studies indicate that in the case of °Bu substituents, the loss of three substituents is largely favoured (−82.5 kcal mol−1) followed by the instance in which two groups are cleaved (−81.4 kcal mol−1). In contrast, for the less bulky °Pr substituent only the C–N bond is energetically favoured (−0.8 kcal mol−1). To further illustrate the steric nature of the observed C–N bond activation we performed parallel studies for the hypothetical °Pr substituted analogue. In this case, the loss of any number of substituents would result in an overall energy cost making it the most thermodynamically unfavourable of modelled systems. Considering solely thermo-
dynamic control over the reaction products we can establish that the C–N bond cleavage is due to steric factors and is generated to ease tensions within the ring upon oxidation of the phosphorus centres. The DFT calculated trends are in agreement with the experimental data obtained for compounds 2 and 4 which underwent C–N cleavage of three and one substituents respectively (see ESI†).

Furthermore, geometry optimizations of the fully oxidized uncleaved product 2′ and 4′, reveal noticeable differences between the C–N bond distances for internal P2N2 ring and bridging nitrogen atoms (1.53 and 1.60 Å for 2′; 1.52 and 1.50 Å for 2″ respectively). Whilst bond distances within 1 and 3 follow the same trend, there is a greater discrepancy between P2N2 ring and bridging C–N bond lengths for the oxidized products than their unoxidized counterparts (0.07 Å in 2 versus 0.04 Å in 1). Calculation of the energy differences associated with the cleavage of the nitrogen substituent at each of these two distinct positions showed that, as expected, the loss of substituents in bridging positions is more favourable than those bonded directly to the P2N2 rings (see ESI†). Our theoretical studies only account for relative stabilities of compounds 2 and 4 (and their subsequent cleavages) when compared with their corresponding uncleaved species 2′ and 4’. We acknowledge that this approach does not account for other important factors involved in this process such as entropy or thermal conditions of the reaction, Gibbs energies, etc. Such studies are currently underway. However, for the purpose of this work, our approach provided results that were consistent with our experimental observations with a feasible computational cost.

In summary, we have shown that the oxidation reaction of [(P(n-R)2)2(P(n-R))]2, R = °Bu and °Pr, with elemental Se prompts the activation of C–N bonds due to ring contraction resulting in their cleavage under mild experimental conditions. Our experimental and theoretical studies demonstrate that steric factors, most probably steric strain release, play a crucial role in the activation of the C–N bonds. The work reported here highlights the potential of steric bond activation and its implications with a broader scope for application to main group frameworks in general. The rational design of synthetic approaches, which fully exploit this approach in a wider range of main group frameworks, is an exciting challenge in the area of metal-free bond activation.

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