Tetragonal phosphorus(v) cations as tunable and robust catalytic Lewis acids†

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The synthesis and catalytic reactivity of a class of water-tolerant cationic phosphorus-based Lewis acids is reported. Corrole-based phosphorus(v) cations of the type [ArP(cor)][B(C6F5)4] (Ar = C6H5; C6F5, 3,5-(CF3)2C6H3; cor = 5,10,15-(C6H5)3corrolato3−, 5,10,15-(C6F5)3corrolato3−) were synthesized and characterized by NMR and X-ray diffraction. The visible electronic absorption spectra of these cationic phosphacorroles depend strongly on the coordination environment at phosphorus, and their Lewis acidities are quantified by spectrophotometric titrations. DFT analyses establish that the character of the P-acceptor orbital diminishes the propensity for phosphine oxide formation and thereby preserving the Lewis acidity at the cationic phosphorus.

We considered that an alternate approach to robust phosphorus-based Lewis acids might be accessible by deliberate alteration of the molecular geometry. Having previously demonstrated that molecular deformation of neutral phosphorus compounds allows for novel bond activation reactions and catalytic transformations, we reasoned that the undesirable hydrolysis pathway leading to inactive phosphine oxides for electrophilic phosphorus cations might be avoided by enforcing nontrigonal substitution at phosphorus. Specifically, we envisioned that a tetragonal substituent field would be less conducive to formation of a formal P=O multiple bond, diminishing the propensity for phosphate oxide formation and thereby preserving the Lewis acidity at the cationic phosphorus.

Introduction

Tetracoordinate phosphonium cations (R4P+) are electrophilic at the phosphorus center, a fact that can be leveraged in Lewis acid catalysis. Stephan reported a major innovation in this area by showing that fluorophosphonium cations (R,P−F) display an exceptionally low-lying σ*(P-F) orbital that is accessible to even weak Lewis bases. Tailoring of the fluorophosphonium core with electron-withdrawing substituents (i.e. (per)fluoroaranes, N-alklypyridiniums) accentuates the P-electrophilicity of the fluorophosphonium, permitting access to compounds with fluoride affinities exceeding that of SbF5, the benchmark Lewis acid. A family of compositionally diverse electrophilic phosphonium Lewis acids is now known, and work from Gabbaï has shown that the underlying design principle is portable to heavier group 15 Lewis acids.

Cofacing with their exceptional electrophilic character, fluorophosphonium cations display a marked sensitivity to water and hydroxyl functionality. Hydrolysis of the fluorophosphonium P-F bond results in decomposition by irreversible formation of a thermodynamically stable phosphate oxide (R,P=O) in which the P-centered electrophilicity is effectively quenched (Scheme 1, top). A reduced hydrolytic sensitivity of certain (trifluoromethyl)phosphonium Lewis acids has recently been achieved by Ingleson and Stephan. Additionally, Stephan has developed air-stable P(III) dicaticonic Lewis acids.

We considered that an alternate approach to robust phosphorus-based Lewis acids might be accessible by deliberate alteration of the molecular geometry. Having previously demonstrated that molecular deformation of neutral phosphorus compounds allows for novel bond activation reactions and catalytic transformations, we reasoned that the undesirable hydrolysis pathway leading to inactive phosphine oxides for electrophilic phosphorus cations might be avoided by enforcing nontrigonal substitution at phosphorus. Specifically, we envisioned that a tetragonal substituent field would be less conducive to formation of a formal P=O multiple bond, diminishing the propensity for phosphate oxide formation and thereby preserving the Lewis acidity at the cationic phosphorus.

Scheme 1 (top) Electrophilic phosphonium cations and their hydrolytic decomposition to phosphate oxides. (bottom) Notional tetragonal electrophilic phosphonium cations that enhance robustness but preserve Lewis acidity.

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† Electronic supplementary information (ESI) available. CCDC 1904833 and 1904834. For ESI and crystallographic data in CIF or other electronic format see DOI: 10.1039/c9sc02463h
center (Scheme 1, bottom). Indeed, Kadish and Vogel and Ravikanth have demonstrated that square pyramidal phosphine oxides embedded within a corrole binding pocket have a propensity to form hexacoordinate structures by association of an exogenous nucleophile. Moreover, it has been shown that phosphorus(v) corroles readily undergo apical halide/alkoxide exchange and are even stable in aqueous media.

We demonstrate here that square pyramidal corrole-based phosphorus(v) cations are robust, tunable, and catalytically-active Lewis acids. We find that the rigid tetragonal geometry imparts stability to water and alcohols while maintaining Lewis acidity, enabling these compounds to effect transformations which were previously inaccessible to phosphonium catalysts. This desirable superposition of properties is rationalized within an electronic structure argument that advances our ongoing program to establish new reactivity for p-block elements by imposition of underexplored molecular geometries.

Results and discussion

Synthesis of the target cations was achieved in two steps from the freebase corrole. First, treatment of 5,10,15-triphenylcorrole with 1 equiv. of phenyl tetrachlorophosphorane (PhPCl₄) in the presence of triethylamine furnished an unstable intermediate, which upon the addition of [Bu₄N][BH(OAc)₃] yielded hexacoordinate 1·H as a chromatographically stable green solid (Fig. 1a). The ³¹P{¹H} NMR spectrum of 1·H showed a resonance at high field (δ = 231.3 ppm), consistent with compositionally similar hexacoordinate phosphorus compounds reported previously. The proton-coupled ³¹P NMR resonance evolves into a doublet of triplets, with coupling constants evidencing a direct P–H bond (J_P–H = 928.0 Hz) as well as longer range coupling to the ortho protons of the apical P-aryl moiety (J_P–H = 25.1 Hz). In the ¹H NMR channel, the P–H unit was observed with complementary coupling (δ = 2.73 ppm, d, J_P–H = 928.8 Hz); the rather high-field chemical shift of this ¹H nucleus is attributed to shielding from the diamagnetic ring current of the corrole system, securing the assignment of the structure of 1·H as in Fig. 1.

The apical hydride of hexacoordinate compound 1·H is readily removed by treatment with abstraction reagents. Specifically, a green solution of 1·H in CH₂Cl₂ when treated with 1 equiv. of [Ph₃C][B(C₆F₅)₄] immediately produced a red solution, from which a new phosphorus-containing product
was obtained by precipitation via slow addition of pentane. A $^{31}$P NMR spectrum of the resulting maroon solid displayed a single new triplet resonance downfield of the starting compound ($\delta = 97.2$ ppm, $J_{P-11} = 20.6$ Hz). This chemical shift is indicative of a pentacoordinate phosphorus center shielded by diamagnetic ring current.$^{8,9,10,15}$ Loss of $J_{P-11}$ coupling and concomitant formation of triphenylmethane further evidence the formation of the hydride abstraction product $1^+$. Related phosphacorroles $2^+–4^+$ were synthesized analogously from the corresponding triarylcorrole and aryl phosphorane as depicted in Fig. 1.

The solid state structure of $1^+$ (as its triflate salt) was revealed by X-ray diffraction experiments (Fig. 1b). As expected, the rigid constraint imposed by the corrole ligand framework enforces a local geometry closely resembling a square pyramid ($\tau = 0.09$)$^{16}$ where the phosphorus center projects 0.373 Å out of the plane containing the four pyrrolic nitrogen atoms.$^{17}$ Fluorinated congener $4^+$ (Fig. 1c) similarly exhibits a near-perfect square pyramidal geometry ($\tau = 0.05$)$^{18}$ where the $P$ center protrudes less from the binding pocket relative to $1^+$ ($\Delta \delta = 0.023$ Å). In both instances, the overall geometry imposed by the corrole ligand may be viewed as a monovacant octahedron about phosphorus. In conjunction with structural data for known hexacoordinate phosphorus corrole compounds (where the phosphorus atom is essentially coplanar with the tetrapyrrolic nitrogens$^{8,11}$), association of Lewis bases to the apical site could be anticipated to proceed with a rather small energy penalty for structural reorganization.

The affinity of cationic phosphacorroles $1^+–4^+$ for Lewis bases was initially assayed by recording $^{31}$P NMR chemical shift differences ($\Delta \delta$) for a phosphine oxide ([n-octyl]$_3$P=O) probe upon binding according to a modification of the Gutmann–Beckett method.$^{18,19}$ In agreement with expectations based on inductive substituent effects, the $\Delta \delta$ values (Table 1) report a self-consistent picture of the increasing Lewis acidity ($1^+ < 2^+ < 3^+ < 4^+$) as a function of increasing modular fluorination. A direct comparison of the Lewis acidity of $1^+–4^+$ to other Lewis acids on the basis of these $\Delta \delta$ values is tempting, but we caution against such a potentially spurious interpretation in the present circumstance. In view of the diamagnetic ring current of the corrole moiety,$^{14}$ a phosphine oxide probe bound apically to the phosphacorrole cation would experience shielding effects that would tend to produce anomalously small $\Delta \delta$ values. Other NMR-based methods for the determination of Lewis acidity (Childs,$^{20}$ Hilt$^{21}$) would similarly be expected to show a systematic underestimation of Lewis acidity for porphyrinoid-based Lewis acids like $1^+–4^+$.$^{22}$

An unbiased quantification of Lewis acidity for $1^+–4^+$ is given by the binding dissociation constant ($K_d$). The marked difference in color between cationic five-coordinate phosphacorroles (red) and neutral six-coordinate congeners (green) provided a convenient colorimetric method for measuring equilibrium binding in $1^+–4^+$. The sensitivity of the color dependence to the concentration of an exogenous Lewis base was demonstrated by titrating cationic phosphacorrole $4^+$ with varying amounts of (n-octyl)P=O (Fig. 2, top). The presence of several isosbestic

Table 1 Summary of important metrics for phosphacorroles $1^+–4^+$

<table>
<thead>
<tr>
<th>Compound</th>
<th>$^{31}$P $\delta$ (ppm)$^a$</th>
<th>$\epsilon_{\text{LUMO}+N}$ (eV)$^b$</th>
<th>FIA (kJ mol$^{-1}$)$^c$</th>
<th>GEI (eV)$^d$</th>
<th>$\Delta \delta$ (ppm)$^{de}$</th>
<th>$K_d$ (μM)$^f$</th>
</tr>
</thead>
<tbody>
<tr>
<td>1$^+$</td>
<td>–97.1</td>
<td>–1.47</td>
<td>274</td>
<td>3.75</td>
<td>2.0</td>
<td>2660 ± 90</td>
</tr>
<tr>
<td>2$^+$</td>
<td>–102.2</td>
<td>–1.72</td>
<td>295</td>
<td>3.90</td>
<td>13.6</td>
<td>30.1 ± 0.7</td>
</tr>
<tr>
<td>3$^+$</td>
<td>–95.2</td>
<td>–1.85</td>
<td>298</td>
<td>4.23</td>
<td>15.3</td>
<td>26 ± 1</td>
</tr>
<tr>
<td>4$^+$</td>
<td>–100.3</td>
<td>–2.31$^e$</td>
<td>343$^e$</td>
<td>4.53$^e$</td>
<td>21.3</td>
<td>11.3 ± 0.4</td>
</tr>
</tbody>
</table>

$^a$ Chemical shift externally referenced to 85% H$_3$PO$_4$. Spectra recorded in CD$_2$Cl$_2$ at 293 K. $^b$ Computed at the B3LYP/def2-TZVP/CPCM(CH$_2$Cl$_2$)//B3LYP/def2-TZVP level. The orbital with appropriate symmetry was LUMO+N, where $N = 3$ (1$^+$), 4 (2$^+$), 2, (3$^+$), 3 (4$^+$). $^c$ Computed according to Christe’s pseudodesmesic method$^{28}$ at the B3LYP/def2-TZVP/CPCM(CH$_2$Cl$_2$)//B3LYP/def2-TZVP level. $^d$ Computed at the B3LYP/def2-TZVP/CPCM(CH$_2$Cl$_2$)//B3LYP/def2-TZVP level as described by Stephan et al.$^{11}$ $^e$ Change in $^{31}$P NMR chemical shift of (n-octyl)P=O. $^f$ Measurements are bracketed by one standard error. $^g$ Values obtained at the B3LYP/def2-TZVP/CPCM(CH$_2$Cl$_2$)//B3LYP/def2-SVP level of theory.
points (e.g. $\lambda = 566$ nm) confirms adduct formation free of decomposition or other deleterious reactivity.

The isomers obtained by monitoring absorption at 610 nm with increasing amounts of ($n$-octyl)$_3$P═O were fitted to the Hill equation\textsuperscript{21} to obtain equilibrium dissociation constants $K_d$ (Table 1, see ESI† for full details). The micromolar dissociation constants confirm the pronounced affinity of the cationic phosphacorroles for phosphine oxide Lewis bases and afford an intrinsic thermodynamic parameter of Lewis acidity for this chemotype.

To further understand the varying Lewis acidities of $1^+$–$4^+$, DFT calculations of the electronic structure of these cationic phosphacorroles were performed. The wavefunctions of $1^+$–$4^+$ were computed at the B3LYP/def2-TZVP/CPCM(CH$_2$Cl$_2$)//B3LYP/def2-TZVP level\textsuperscript{24} as implemented in the ORCA 4.0.0 software package,\textsuperscript{25} and are found to resemble experimental structures closely (see ESI† for full details). In excellent agreement with previous theoretical studies on isolobal Ga(III) corroles, the LUMO and LUMO+1 of $1^+$–$4^+$ correspond to the corrole $\pi$ manifold, and are apparently not responsible for the experimentally observed Lewis acidity of cationic phosphacorroles.

Another low-lying unoccupied orbital orbital (LUMO+3 for $4^+$, Fig. 3)\textsuperscript{27} is still quite low in energy (e.g. $-2.31$ eV for $4^+$, see Table 1) and projects into the apical space proximal to phosphorus, rendering it both energetically and sterically accessible for attack by exogenous nucleophiles. Indeed, the calculated fluoride ion affinities (FIAs) for $1^+$–$4^+$ correlate with the experimental dissociation constants $K_d$ for phosphine oxide binding, implying that Lewis acid/base interactions are hosted by this orbital (see ESI† for details). Most notably, the character of this P-acceptor orbital illustrates a fundamental distinction with respect to prior phosphonium Lewis acids that has important implications for their stability. Whereas the acceptor orbital for trigonal electrophilic phosphonium cations is $\sigma$-antibonding with respect to the trans-apical substituent in the developing Lewis acid/base adduct,\textsuperscript{3} the phosphorus-centered acceptor orbital in tetragonal phosphorus cations $1^+$–$4^+$ primarily constitutes basal P–N antibonding interactions. As a consequence, the mechanistic pathway initiated by water addition which leads to apical ejection of a P-substituent for trigonal phosphoniums\textsuperscript{29} is denied by the corrole chelate. On this basis, we anticipated that tetragonal cations $1^+$–$4^+$ might be relatively resistant to decomposition by water.

In order to probe this conjecture, we investigated the chemical stability of $1^+$–$4^+$ with respect to hydrolysis. In a representative experiment (Scheme 2), treatment of a red CD$_2$CN solution of $3^+$ with 10 equiv. of water resulted in a purple-green solution whose $^{31}$P{1H} NMR spectrum exhibited a broad resonance at $\delta = 175.4$ ppm. The chemical shift is consistent with a six-coordinate P atom, and the broadness of the peak suggests reversible binding to the Lewis acidic phosphorus. Importantly, there is no evidence of decomposition of this intermediate upon prolonged standing. Although this adduct has thus far eluded isolation, the addition of even a weak base (MgSO$_4$) cleanly gives $P$-hydroxide $3^+$ (Scheme 2). The same reasoning may be extended by analogy to $1^+$, $2^+$, and $4^+$.

The phosphacorrole cations $1^+$–$4^+$ are potent Lewis acid catalysts for a range of transformations.\textsuperscript{30} We benchmarked the catalytic activity of $1^+$ and $2^+$ with typical carbonyl reduction reactions, such as ketone hydrosilylation and deoxygenation using HSiEt$_3$ as a terminal reductant (see ESI† for full details). The high activity of $1^+$ and $2^+$ for these reductions encouraged us to attempt more challenging catalytic transformations. For instance, ring-forming C$_{sp3}$═H functionalization of substrate 5 is induced by catalytic $3^+$ (1 mol%) via $1,5$-

![Fig. 3 Kohn–Sham orbital LUMO+3 for $4^+$ (top: in perspective; bottom: side-on view) computed at the B3LYP/def2-TZVP//B3LYP/def2-SVP level.](image)

![Scheme 2 Reaction of water with $3^+$ and formation of putative water adduct $3^+$–OH$_2$. Deprotonation by MgSO$_4$ gives product $3$–OH, which is reactivated to $3^+$ by TMS–OTf.](image)
hydride shift from a N,N-dialkylanilnine donor to a malonate alkylidene acceptor followed by intramolecular cyclization to give 6 in good isolated yield (Scheme 3a). Furthermore, the robustness of the tetragonal cationic phosphorus corrole Lewis acids is exemplified by the observation that unprotected 13C6-glucose (7) is exhaustively deoxygenated by a catalytic amount of 4+ (5 mol%) in the presence of an excess of H2SiEt2 at room temperature to give a mixture of hexanes and hexenes (67% total yield, Scheme 3b). Notably, analysis after completion of the reaction shows that 4+ is not degraded; instead, an approximately equimolar amount of 4-H and 4+ were observed spectroscopically as the only 31P NMR resonances. The lack of reaction between 4+ and H2SiEt2 in a control experiment further indicated that 4+ is a true catalyst for this reaction. The persistence of 4+ with respect to a substrate that under these conditions presents a 100:1 ratio of free hydroxyl moieties to catalyst confirms the noteworthily chemical robustness inherent to tetragonal cationic phosphorus-based Lewis acids.

Conclusions
In summary, we have shown that cationic phosphacorroles are potent Lewis acids that exhibit marked tolerance toward hydroxyl functionality including water. We propose that this useful property arises from the tetragonal geometry of 1+–4+ as enforced by the tetraazamacrocyle, which produces an acceptor orbital that is distinct in character from prior trigonal electrophilic phosphonium cations and prohibits irreversible decomposition to phosphine oxides. The modularity of the corrole supporting framework allows the Lewis acidities of these electrophilic phosphorus species to be readily tuned. Together, these results establish within the family of designer main group Lewis acids a new structural type that extends the range of potential applications for this valuable class of compounds.

Conflicts of interest
There are no conflicts to declare.

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Notes and references
Chemical Science


13 A minor, inseparable impurity was also observed in the $^{31}$P and $^1$H NMR spectra of 1. We attribute this observation to the partial chlorination of the corrole β-pyrrolic carbons which presumably occurs by oxidation by Cl$_2$ generated in the refluxing reaction mixture. We were able to remove this species by hydrodechlorination with H$_2$ in the presence of catalytic amounts of Pd/C. See ESI for full details.


17 The counterion for both 1$^+$ and 4$^+$ resides outside the phosphorus coordination sphere, in agreement with solution phase NMR data.


19 We observed unacceptable run-to-run variability in replicates of the Guttmann–Beckett measurement with Et$_3$P=O, which we attribute to the hygroscopicity of this probe reagent (there may be competitive binding with water). We chose the hydrophobic tri-$n$-octylphosphine oxide (–octyl)$_3$P=O, which presents a similar steric and electronic profile to Et$_3$P=O as a rigorously anhydrous analog of Et$_3$P=O for studies measuring Lewis acidity.


27 For compounds 1$^+$, 2$^+$, and 4$^+$, the relevant orbitals were LUMO$^+$, LUMO$^+$, and LUMO$^+$ for LUMO$^+$+LUMO$^+$ in these instances correspond to π bonding orbitals within the apical π-aromatic substituent.


