Hydrogen Activation

A New Mode of Chemical Reactivity for Metal-Free Hydrogen Activation by Lewis Acidic Boranes

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Abstract: We herein explore whether tris(aryl)borane Lewis acids are capable of cleaving H₂ outside of the usual Lewis acid/base chemistry described by the concept of frustrated Lewis pairs (FLPs). Instead of a Lewis base we use a chemical reductant to generate stable radical anions of two highly hindered boranes: tris(3,5-dinitromesityl)borane and tris(mesityl)borane. NMR spectroscopic characterization reveals that the corresponding borane radical anions activate (cleave) dihydrogen, whilst EPR spectroscopic characterization, supported by computational analysis, reveals the intermediates along the hydrogen activation pathway. This radical-based, redox pathway involves the homolytic cleavage of H₂ in contrast to conventional models of FLP chemistry, which invoke a heterolytic cleavage pathway. This represents a new mode of chemical reactivity for hydrogen activation by borane Lewis acids.

The chemistry of Lewis acidic boranes reacting with H₂ is now almost exclusively described by the Lewis acid/base conceptual framework of frustrated Lewis pairs (FLPs), introduced by Douglas Stephan in 2006. While some precise mechanistic details are still debated, in general the ability of FLPs to cleave H₂ relies on the cooperative action of the two reactive centers that are sterically encumbered (“frustrated”) within an encounter complex of the Lewis acid–base pair. The Lewis acid, which is most often an organoborane, provides a vacant acceptor orbital, and the Lewis base, typically a phosphine or amine, provides a donor orbital with which to cleave the strong H–H bond. Activation of H₂ by borane-based FLPs is therefore widely thought to involve heterolytic bond cleavage, and to be controlled by the relative strengths of the Lewis acidic/Lewis basic components and the degree of steric encumbrance between them. This contrasts with the transition-metal-based complexes and biological systems that have dominated hydrogenation catalysis for the previous 150 years. In these complexes, the metal center provides both vacant and filled acceptor/donor orbitals at a single reactive site; the chemistry is, in a large extent, operating under redox control of the metal center, and homolytic H₂ bond cleavage is common.

The heterolytic mechanism proposed for FLP activation of H₂ is found generally to be in good agreement with observed trends in reactivity, and it has been supported by a number of computational studies. Nevertheless, definitive experimental proof has remained elusive (perhaps unavoidably so). As such it is interesting to consider that observed patterns of FLP reactivity could also be consistent with alternative H₂ activation pathways. These trends could also be consistent with plausible radical mechanisms, in which initial single-electron transfer (SET) from the Lewis base to the Lewis acid would transiently generate highly reactive radical pairs capable of activating H₂. For example, while the thermodynamic and kinetic ability of an FLP to activate H₂ is well known to correlate with the hydride-ion affinity of the Lewis acid (consistent with heterolytic bond cleavage), these parameters also correlate well with the one-electron reduction potential of the Lewis acid (consistent with SET). Indeed, recent studies have implied that for some families of borane Lewis acids, reduction potentials may even be a better indicator of reactivity towards H₂ than hydride-ion affinities.

There is also a growing body of evidence for the occurrence of radical mechanisms when small molecules, such as NO, Ph₃SnH, and peroxides, are used as the substrates of FLP reactions. To date, however, these frustrated radical pair (FRP) mechanisms have not been observed with H₂. Indeed, no FLP is known to cleave H₂ via a radical mechanism. Our previous work studying the electrochemistry of FLP components, together with the recent evidence for radical pathways in FLPs and FRPs reported by others, raises an obvious question that this article sets out to answer: can
boranes react with H$_2$ outside of an FLP chemical framework, if they can operate via a hitherto unknown redox controlled, radical reaction pathway instead?

To test our hypothesis, we carefully selected two boranes as models: tris(3,5-dinitromesityl)borane 1, and tris-(mesityl)borane 2 (Scheme 1). Both boranes have essentially identical steric shielding of the central boron atom by the six ortho methyl groups on the mesityl rings, leading to the formation of long-lived borane radical anions upon reduction.$^{[9,10]}$ Neither borane is currently known to be active for H$_2$ activation within an FLP. The addition of six electron-withdrawing nitro groups in 1 shifts the reduction potential in a positive direction to $-1.57$ V vs. Cp$_2$Fe$^{0+}$ (see the Supporting Information), making 1 as electrophilic and comparably facile to reduce as the archetypal electron-deficient borane B(C$_6$F$_5$)$_3$ used in FLP chemistry ($-1.52$ V vs. Cp$_2$Fe$^{0+}$)$^{[18]}$ and much easier to reduce than 2 (approximately $-2.8$ V vs. Cp$_2$Fe$^{0+}$)$^{[11]}$. The NO$_3$ groups in 1 also provide useful electron paramagnetic resonance spectroscopic markers for the characterization of reaction intermediates.

To examine whether the radical anions of Lewis acidic boranes are capable of cleaving hydrogen, a solution of 1 in either CD$_2$Cl$_2$ or [D$_2$]THF was chemically reduced using decamethylcobaltocene ([Cp$_5$Co]$^0$, $E^{0/1} = -1.94$ V vs. Cp$_2$Fe$^{0+}$)$^{[12]}$ heated in the presence of H$_2$, and the reaction periodically monitored using multinuclear NMR spectroscopy (see the Supporting Information for details). Figure 1a shows the resulting $^{13}$B NMR spectra. The formation of the borohydride product [Cp$_5$Co][I-H] is clearly evident by the observation of a characteristic doublet at $\delta = -13.6$ ppm ($J_{BH} = 82$ Hz) in the $^1$B NMR spectrum and the corresponding 1:1:1:1 quartet at $\delta = +3.8$ ppm ($J_{BH} = 82$ Hz) in the $^1$H NMR spectrum. The spectral assignment was further confirmed by comparison to an authentic sample of [Na][I-H] (Supporting Information, Figure S10). Control experiments using D$_2$ in proto-CH$_2$Cl$_2$ or proto-THF produced the analogous result, the generation of [Cp$_5$Co][1-D] (Supporting Information, Figures S11 and S12), observed as a partially resolved triplet at $\delta = -13.6$ ppm in the $^1$B NMR spectrum.

In these reactions, the cleavage of H$_2$/D$_2$ must be homolytic as there is no apparent plausible mechanism to allow for the formation of H$^+$ (no counter anion), which must be produced via heterolytic scission of H$_2$. Whilst very strong acids are known to protonate Cp$_5$Co$^{[13]}$ there is no observable evidence for the formation of this in these reactions. To examine the proposed radical homolytic dihydrogen cleavage mechanism, 1 was again reduced with Cp$_5$Co under H$_2$ but this time in the presence of 1 equiv of the radical spin-trap TEMPO ((2,2,6,6-tetramethylpiperidin-1-yl)oxy). TEMPO was selected because it does not coordinate to the bulky borane 1 and has a more negative reduction potential than Cp$_5$Co$^{[14]}$ thus precluding any possible redox inhibition to form [TEMPO$^+$]- and 1, which together could subsequently participate in FLP H$_2$ activation. In the presence of TEMPO, no H$_2$ cleavage was observed, consistent with inhibition of a radical reaction by the TEMPO spin-trap. Additional control experiments confirm that Cp$_5$Co alone does not activate H$_2$ under these conditions and that THF/I mixtures do not result in the observable formation of [I-H] via a solvent-FLP mechanism$^{[15]}$ in the absence of a reducing agent. Crucially, no evidence of reduction at the nitro groups is observed by NMR, EPR, nor IR spectroscopic characterization of the reaction products.

The very negative redox potential of 2 necessitates the use of a stronger reducing agent. When a solution of 2 in [D$_2$]THF is reduced over sodium metal$^{[16]}$ and heated in the presence of H$_2$, the appearance of a doublet in the $^1$B NMR spectrum at $\delta = -14.5$ ppm ($J_{BH} = 78$ Hz), and a corresponding 1:1:1:1 quartet in the $^1$H NMR spectrum at $\delta = 3.75$ ppm ($J_{BH} = 77$ Hz) is observed, characteristic of the formation of [Na]-[2-H] (Figure 1b).

Scheme 1. Reduction of tris(3,5-dinitromesityl)borane 1, and tris-(mesityl)borane 2, and subsequent reaction with H$_2$. 

Figure 1. Overlaid $^1$B NMR spectra expanded over the B–H bond region of interest, showing the progression of H$_2$ cleavage by chemical reduction of 1 in CD$_2$Cl$_2$ (a) and 2 in THF (b). Inset: The corresponding $^1$B NMR spectra recorded at the start and end of the experiments showing the conversion of the parent borane starting material to the borohydride product upon reduction and exposure to H$_2$. 

The experiments described above clearly indicate that the boron radical anions $1^-$ and $2^-$ can cleave H$_2$ in the absence of any exogenous Lewis base. These reactions are, however, slow in comparison to typical FLP H$_2$ activation reactions. In the case of the model borane $1$, this is advantageous, since it enables the reaction to be monitored in real time and reaction intermediates along the H$_2$ cleavage pathway to be observed using EPR spectroscopy.

Solutions of $1$ dissolved in either CD$_2$Cl$_2$ or [D$_2$]THF were chemically reduced using Cp*Co (see Supporting Information) and the EPR spectra resulting from exposure to H$_2$ were recorded (Figures 2a–d). Simulation of the EPR spectra yields the isotropic hyperfine coupling constants for the various $^1$H, $^{14}$N, and $^{11}$B nuclei, given in Table 1. These data, supported by DFT calculations (performed for the identifiable intermediates of both $1$ and $2$, and detailed in the Supporting Information), enable us to observe and characterize the structures of the intermediates and gain valuable insights into the reaction mechanism (given schematically in Figure 3) and the corresponding energetic profile by which organoborane radicals cleave H$_2$ homolytically (Figure 4).

Upon reduction of $1$ under N$_2$, the EPR spectrum shown in Figure 2a is observed, which is characteristic of $1^-$ with hyperfine coupling of the unpaired electron spin density to the boron nucleus as well as the methyl and nitro substituents on the aromatic rings (Table 1). $^{[11]}$ The initiation step is calculated to be exothermic for both compounds ($-56.7$ and $-11.7$ kcal mol$^{-1}$ for $1$ and $2$, respectively) and reflects the relative LUMO energy and reduction potential of each borane.

Figure 2b shows the resulting spectrum recorded upon first exposing the reaction to H$_2$ and before heating. An immediate change is evident with the appearance of a sharp $1:1:1$ three-line signal superimposed on the original signal of the $1^-$ parent. After heating the reaction for a further 10 minutes this three-line signal dominates the EPR spectral response (Figure 2c) for the next 48 hours. The only change to the system is the addition of H$_2$ and computational modelling of the possible interactions between $1^-$ and H$_2$ reveal two propagation pathways. Propagation 1a produces the diamagnetic borohydride product, and is endothermic ($+30.8$ and $+28.1$ kcal mol$^{-1}$ for $1$ and $2$), albeit to a lesser extent than homolytic H$_2$ splitting itself ($+107.1$ kcal mol$^{-1}$ at this level of theory). The alternative pathway, Propagation 1b avoids the

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Figure 3. The proposed radical chain-propagation mechanism for the homolytic cleavage of H$_2$ upon reduction of organoborane Lewis acids. Inset: the chemical structures corresponding to the $[\text{Ar}_2\text{B}(\text{H})-\text{Ar}(\text{H})]^-$ and $[\text{Ar}_2\text{B}+\text{Ar}(\text{H})]^-$ intermediates (substituents on the aryl rings have been omitted for clarity).
atom which is the most stable isomer in the case of 1 (whereas the \textit{meta} position is most stable in 2, see Table S1 in the Supporting Information). This is supported by what is observed experimentally during the EPR spectroscopic monitoring of hydrogen splitting by 1 where the magnitude of the resulting H\textsuperscript{−} atom hyperfine coupling fits well with coupling to spin density on the ring system in the \textit{para} position located between the two nitro groups (Figure 2d).

If the parent borane is present in excess of the radical anion (Propagation 2b), the hydrogen atom produced in step 1a (considered as [Ar,B(H)-Ar(H)]\textsuperscript{−}) may be transferred, and the borohydride product and the neutral [Ar,B-H] radical intermediate formed. Using the values calculated for propagation steps 1a and 2a, step 2b is energetically neutral. In the system reported herein, it is unlikely that the parent borane is present in excess of the radical anion initially, but as the reaction proceeds through step 3b and the consumption of the [Ar,B(H)-Ar(H)]\textsuperscript{−} progresses, this stabilization may become more relevant towards the end of the reaction. This situation may also have relevance to potential radical-FLP hydrogen cleavage mechanisms, where the parent borane is most likely present in excess of any potential radical anion intermediates throughout.

The final step in the reaction, which cannot be observed by EPR spectroscopy, is the formation of the diamagnetic [1-H\textsuperscript{−}] product, which is detected by \textsuperscript{11}B and \textsuperscript{1}H NMR spectroscopic analysis of the reaction mixture at the end of the experiment. Aside from the obvious recombination of 2H\textsuperscript{+} to form H\textsubscript{2} (the reverse of step 1), there are two termination pathways: Termination 3a (−39.5 and −49.6 kcal mol\textsuperscript{−1} for 1 and 2, respectively), and Termination-Propagation 3b (−39.3 and −48.2 kcal mol\textsuperscript{−1} for 1 and 2, respectively). Step 3a may also be written as [Ar,B]\textsuperscript{+} + H\textsuperscript{−}−[Ar,B-H\textsuperscript{−}] for consistency with the rest of the Scheme, or as a termolecular reaction: 2[Ar,B]\textsuperscript{+} + H\textsuperscript{−}−2[Ar,B-H\textsuperscript{−}] . Step 3b yields both the terminal borohydride product and regenerates the parent neutral borane for further reaction in propagation step 2a. Note that whilst it would appear from Figures 2c,d that the EPR spectra are dominated by the [1-H\textsubscript{−}]\textsuperscript{−} and [1-H\textsuperscript{−}]\textsuperscript{−} species, respectively, simulation of the spectral data reveals that these spectra are each superimposed over the parent \textsuperscript{1}H\textsuperscript{−} radical anion species. As the reaction proceeds with heating the weighting between the systems changes (\textsuperscript{1}H\textsuperscript{−} : [1-H\textsubscript{−}]\textsuperscript{−} = 98.5 : 1.5 in Figure 2b; 73.0 : 27.0 in Figure 2c, and \textsuperscript{1}H\textsuperscript{−} : [1-H\textsubscript{−}]\textsuperscript{−} = 63.0 : 37.0 in Figure 2d). The rate of consumption of \textsuperscript{1}H\textsuperscript{−} as measured by EPR (Figures 2a–d) correlates with the rate of conversion to borohydride as measured by NMR spectroscopy (Figure 1a).

In summary, using two model boranes, which produce stable radical anions upon one-electron reduction, we have successfully demonstrated homolytic dihydrogen cleavage in the absence of a Lewis base. This represents a new mode of chemical reactivity by Lewis acidic boranes towards H\textsubscript{2} that opens up new borane, and potentially other main group chemistries, beyond the framework of conventional FLPs. The reaction between the model borane radical anions and H\textsubscript{2} is slow, and the intermediates are sufficiently stabilized so that we can observe several distinct intermediates along the homolytic dihydrogen cleavage pathway using EPR spectroscopy.
copy and can model the energetics of the reaction pathway computationally. We are currently exploring the application of boryl radical $\text{H}_2$ activation as a convenient route to more active borane hydride species, which may have applications in catalysis and energy materials.

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**Conflict of interest**

The authors declare no conflict of interest.

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