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# Untangling the catalytic importance of the Se oxidation state in organoselenium-mediated oxygen-transfer reactions: the conversion of aniline to nitrobenzene†

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Seleninic acids and their precursors are well-known oxygen-transfer agents that can catalyze several oxidations with H<sub>2</sub>O<sub>2</sub> as the final oxidant. Until very recently, the Se(IV) "peroxyseleninic" acid species has been considered the only plausible catalytic oxidant. Conversely, in 2020, the involvement of Se(VI) "peroxyselenonic" acid has been proposed for the selenium mediated epoxidation of alkenes. In this work, we theoretically probe different mechanisms of H<sub>2</sub>O<sub>2</sub> activation and of Se(IV) to Se(VI) interconversion. In addition, we investigate through a combined theoretical (DFT) and experimental approach the mechanistic steps leading to the oxidation of aniline to nitrobenzene, when Se(IV) seleninic acid or Se(VI) selenonic acids are used as catalysts and H<sub>2</sub>O<sub>2</sub> as the oxidant. This process encompasses three subsequent organoselenium mediated oxidations by H<sub>2</sub>O<sub>2</sub>. These results provide a mechanistic explanation of the advantages and disadvantages of both oxidation states (IV and VI) in the different stages of catalytic oxygen-transfer reactions: hydrogen peroxide activation and actual substrate oxidation. While the Se(VI) "peroxyselenonic" acid is found to be a better oxidant, the privileged role of "peroxyseleninic" acid as the main active species is assessed and its origin is identified in the lower catalyst-distortion that seleninic acid undergoes when activating H<sub>2</sub>O<sub>2</sub>. Conversely, the higher catalyst-distortion that characterizes the reaction of selenonic acid with H<sub>2</sub>O<sub>2</sub> supports an inactivating role of Se(IV) to Se(VI) interconversion.

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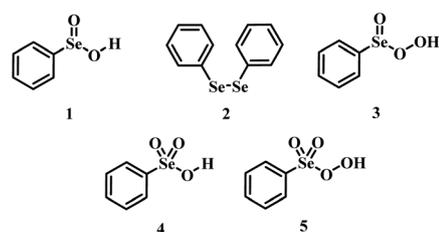
## Introduction

Oxidations are an important class of reactions allowing the manipulation of a variety of functional groups.<sup>1</sup> While many effective oxidants are available, the employment of green oxidants such as hydrogen peroxide for the practical oxidation of organic substrates usually requires the use of a catalyst to activate hydrogen peroxide into an oxygen-transfer species which is kinetically more reactive.<sup>2,3</sup>

Organoselenides are recognized as effective oxygen-transfer catalysts<sup>1</sup> in epoxidation/dihydroxylation reactions,<sup>4–7</sup>

sulfoxidation,<sup>8</sup> aldehyde<sup>9</sup> and amine oxidations,<sup>10,11</sup> with H<sub>2</sub>O<sub>2</sub> as the final oxidant. For reactions catalyzed by phenyl seleninic acid **1** or its respective precursor (e.g., diphenyl diselenide **2**), until recently, phenyl peroxyseleninic acid **3** was deemed the only plausible active catalytic intermediate (Scheme 1).

The first hypothetical participation of peroxyseleninic acids in oxidation reactions dates back to the early days of organoselenium chemistry, when the oxidation of seleninic to peroxyseleninic acid was postulated to rationalize the autocatalytic behavior of selenide oxidation.<sup>12</sup> Later on, their participation in



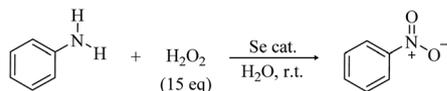
**Scheme 1** Organoselenides employed as catalysts or pre-catalysts for selenium mediated oxygen-transfer reactions and postulated plausible active oxidants.

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† Electronic supplementary information (ESI) available: Additional computational details, energetics and TOF calculations carried out with OPBE and OPBE0 density functionals, additional activation and reaction energies, Cartesian coordinates and electronic energies of all structures discussed in the article and DLPNO-CCSD(T) calculations. Synthetic procedures and experimental details on control experiments are also reported. See DOI: <https://doi.org/10.1039/d4sc03329a>





Scheme 2 On-water selenium catalyzed oxidation of aniline to nitrobenzene at room temperature (r.t.).

various organic substrate oxidations was proposed<sup>4,8,13</sup> and it consolidated in the past fifty years.<sup>1,14</sup> In 2020, Back and coworkers<sup>6</sup> reported for the first time evidence of the involvement of the high oxidation state intermediates seleninic **4** and peroxyseleninic acid **5** in the selenium-mediated epoxidation of cyclooctene. This observation, while thought-provoking, was suggested to be not completely general. Indeed, in 2021, evidence was reported for the involvement of the “conventional” peroxyseleninic acid **3** in another selenium catalyzed oxygen-transfer process, *i.e.*, the on-water oxidation of aniline to nitrobenzene<sup>11</sup> (Scheme 2).

In the work by Tanini *et al.*,<sup>11</sup> Se(vi) selenonic acid **4** was recovered in the water after complete oxidation of the substrate but was found inactive towards further catalytic activity if not reduced back to Se(IV) seleninic acid **1**. Conversely, in the selenium-mediated epoxidation of alkenes, selenonic acid was found to be more active than seleninic acid itself.<sup>6</sup> Puzzled by these apparently conflicting results, we chose to investigate accurately the mechanistic details of Se-promoted H<sub>2</sub>O<sub>2</sub> activation, which should be a key reactive step common to all reactions catalyzed by **1** or **2**.

Despite the long-standing experimental experience in the field, to the best of our knowledge, no detailed theoretical mechanistic investigation has ever been carried out on catalytic oxygen-transfer reactions mediated by organoselenides. Only a previous report discussed peroxyseleninic acid **3** formation starting from the parent diphenyl diselenide **2**, addressing the autocatalytic decomposition of H<sub>2</sub>O<sub>2</sub>.<sup>15</sup> Indeed, while much theoretical mechanistic effort was devoted in the past twenty years to the elucidation of glutathione peroxidase-like catalytic potential of organoselenides,<sup>16,17</sup> other catalytic reactions remained somewhat unexplored *in silico*. Among the different selenium catalyzed oxygen transfer processes,<sup>1</sup> we chose to focus on the aniline oxidation to nitrobenzene, which represents an intriguing case involving three consecutive catalytic oxygen-transfers. Indeed, according to previous studies,<sup>10,18</sup> the oxidation of aniline to nitrobenzene goes through hydroxyanilines and nitrosobenzene as intermediates, which are progressively oxidized (Scheme 3).

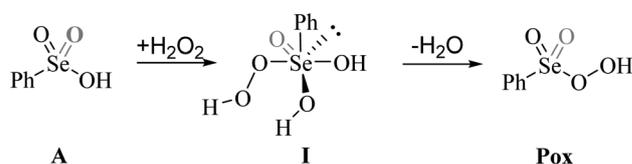
As previously mentioned, no theoretical mechanistic investigation has to date been reported for these and similar catalytic reactions.<sup>11</sup> Additionally, while experimental evidence supports a Se(IV) only catalysis, no theoretical explanation as to why only

one oxidation state of Se is active in the reaction is available until now. Thus, with the inactive nature of the Se(vi) selenonic acid as a catalyst in mind, we aim to provide a detailed mechanistic picture of the title reaction to understand the different properties of Se(IV) and Se(vi) acids in the three different progressive oxygen-transfers and the hydrogen peroxide activation step.

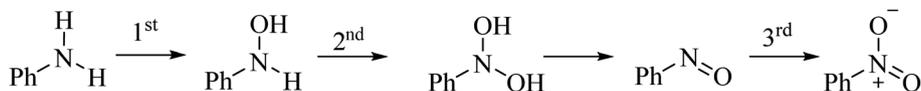
## Results and discussion

Previous experimental evidence showed that seleninic acid and selenonic acid alone do not oxidize aniline.<sup>10,11</sup> Thus, the catalyst must first react with hydrogen peroxide to produce the actual oxygen-transfer species. In oxygen-transfer processes catalyzed by Se(IV) or Se(vi), the activation of H<sub>2</sub>O<sub>2</sub> is thought to occur *via* the formation of the corresponding organoselenium peroxyacid, and this oxygen-transfer species is believed to be fundamental also for other organoselenium catalyzed oxidations.<sup>1,14</sup> Since this step is required to produce the active oxidant in the reaction, we first discuss the formation of these species starting from the corresponding acids and hydrogen peroxide. Different mechanisms for peroxyacid formation were investigated, both concerted and stepwise, and the most favorable energetics were obtained for a mechanistic hypothesis that closely matches the one made by Back *et al.*,<sup>6</sup> in which the H<sub>2</sub>O<sub>2</sub> activation step is independent of the nature of the substrate and proceeds in two steps. This stepwise process proceeds through the intermediate formation of a peroxy selenurane (I), which can then undergo a non-redox dehydration to the corresponding peroxyacid (Pox) (Scheme 4).

In the H<sub>2</sub>O<sub>2</sub> activation process (from A to Pox), Se(IV) seleninic acid **1** appears to be the best reactant. Indeed, not only the formation of the peroxyseleninic acid **3** is slightly more favored thermodynamically as compared to the same process for Se(vi), *i.e.* the conversion of **4** to **5**, ( $\Delta G_r$  of  $-4.7$  and  $-1.7$  kcal mol<sup>-1</sup> respectively), but the overall process occurs on a much lower energy surface. While the formation of the peroxy selenurane (I) is endergonic for both Se oxidation states, the dehydration then occurs with a relatively low activation energy. Both H<sub>2</sub>O<sub>2</sub>



Scheme 4 Hydrogen peroxide activation by organoselenium acids (A) in the form of peroxyacids (Pox). The grey double bond represents the additional bond present only in the Se(vi) selenonic acid and derivatives.



Scheme 3 Oxidation of aniline to nitrobenzene, with the three oxygen-transfer steps highlighted.



addition and the peroxy selenurane (I) dehydration are located much lower on the PES when seleninic acid is involved, as compared to selenonic acid. Particularly, the selenurane (I) itself is much more stable for Se(IV) than for Se(VI), being located on the PES respectively at 13.9 and 23.8 kcal mol<sup>-1</sup>. This energy difference is roughly conserved also when comparing TS1 and TS2, *i.e.*, in the transition states for the H<sub>2</sub>O<sub>2</sub> addition (from A to I) and dehydration processes (from I to Pox). Thus, overall, the peroxyacid Pox formation occurs with an activation energy 10 kcal mol<sup>-1</sup> lower for seleninic acid than for selenonic acid, making the former much more privileged energetically in H<sub>2</sub>O<sub>2</sub> activation ( $\Delta G^\ddagger$  of 22.1 and 31.7 kcal mol<sup>-1</sup> for H<sub>2</sub>O<sub>2</sub> addition, TS1, and of 24.0 and 33.9 kcal mol<sup>-1</sup> for the dehydration, TS2, respectively). The choice of the solvent did not affect these outcomes, even if highly non-polar solvents (*e.g.*, benzene) were found to moderately decrease the height of all the barriers as compared to polar solvents. (Table S1†) Along the manuscript, only data in water will be discussed for consistency with the previous experimental work by Tanini *et al.*,<sup>11</sup> which was carried out under on-water conditions (see also Computational details).

Conversely, two other possible H<sub>2</sub>O<sub>2</sub> activation mechanisms (all independent of the nature of the substrate) were found to be kinetically less favored by more than 10 kcal mol<sup>-1</sup>. Both investigated concerted processes (Table 1b and c) have a higher activation energy than the corresponding H<sub>2</sub>O<sub>2</sub> addition step (Table 1a) and higher than the overall stepwise H<sub>2</sub>O<sub>2</sub> activation process. Particularly, the previously proposed concerted mechanism,<sup>15</sup> in which one proton of H<sub>2</sub>O<sub>2</sub> is transferred to the -OH group of the acids **1** or **4** (Table 1b) has an activation energy already more than 10 kcal mol<sup>-1</sup> higher than H<sub>2</sub>O<sub>2</sub> addition, in which the proton of H<sub>2</sub>O<sub>2</sub> is transferred to the corresponding Se-O double bond moiety (Table 1a). A concerted mechanism

corresponding to an “O insertion” within the Se-OH bond of seleninic and selenonic acids, thus turning the -OH function into the -OOH function (Table 1c), displays an even higher, unfeasible activation energy. These results suggest that the formation of the peroxy selenurane intermediate (I) is pivotal in the actual conversion between the acid and the peroxyacid species. Interestingly, also for the two concerted (unlikely) mechanisms, Se(IV) appears to be consistently more reactive, a result further corroborating our previous conclusions about H<sub>2</sub>O<sub>2</sub> activation by seleninic and selenonic acids, and suggesting that the relative inertia of Se(VI) in activating H<sub>2</sub>O<sub>2</sub> might be intrinsic in its high oxidation state.

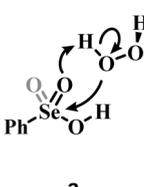
To provide a quantitative discussion on the effect of the oxidation state on the H<sub>2</sub>O<sub>2</sub> activation, ASA and EDA were performed. (see Computational methods). Particularly, the addition of H<sub>2</sub>O<sub>2</sub> to seleninic and selenonic acid was compared to provide a detailed insight into the chalcogen different oxidation state effect in the most favored pathway, *i.e.*, the stepwise activation (Scheme 3).

The ASA for the H<sub>2</sub>O<sub>2</sub> addition step is represented in Fig. 1a. All energies were plotted at consistent values of O-H bond breaking. This reaction coordinate (*r.c.*) undergoes a well-defined variation along the reaction since the proton is transferred from H<sub>2</sub>O<sub>2</sub> to the Se=O bond of seleninic and selenonic acid. Analogous conclusions can be drawn by analysis of the complementary *r.c.* *i.e.*, the H-O bond formation in the peroxy selenurane (I). (Fig. S1†) It can be seen how, from the beginning of the reaction to the TS, which occurs roughly at the same value of *r.c.* for the two reactions (at *ca.* 0.35 Å) the  $\Delta E_{\text{int}}$  for both OS is superimposed, suggesting that both seleninic and selenonic acids interact similarly with H<sub>2</sub>O<sub>2</sub>. Conversely, seleninic acid has a consistently lower  $\Delta E_{\text{str}}$ , which appears to be the main player in its reduced activation energy in H<sub>2</sub>O<sub>2</sub> addition.

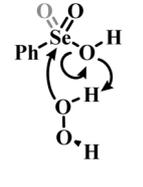
The strain energy was then decomposed into the contribution of each fragment *i.e.*, a  $\Delta E_{\text{str},\text{H}_2\text{O}_2}$  accounting for H<sub>2</sub>O<sub>2</sub> deformation only, and  $\Delta E_{\text{str},\text{Se(OS)}}$  associated with seleninic/selenonic acid deformation alone (Fig. 1b). What can be inferred is that the higher strain, and thus the higher activation energy predicted for H<sub>2</sub>O<sub>2</sub> addition to Se(VI), is due exclusively to the higher  $\Delta E_{\text{str},\text{Se(VI)}}$  when compared to  $\Delta E_{\text{str},\text{Se(IV)}}$ . Indeed, seleninic acid has a lower  $\Delta E_{\text{str},\text{Se(OS)}}$  along the whole *r.c.*, when compared to selenonic acid. We interpreted this result as an effect of the different environment around the Se nucleus in the two OSs. Indeed, selenonic acid has four groups in the surrounding of Se (a phenyl group, two oxygens and one hydroxyl group), while seleninic has only three groups (one Se=O less than selenonic acid); thus, the distortion required to collocate the new OOH group around Se is intuitively larger for the former than for the latter, as reflected by its higher  $\Delta E_{\text{str}}$ .

**Table 1** Activation energies (kcal mol<sup>-1</sup>) for three different H<sub>2</sub>O<sub>2</sub> activation mechanisms. Schematic transition states of the processes are shown below<sup>a</sup>

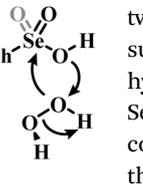
	$\Delta G^\ddagger$	
	Se(IV)	Se(VI)
<b>a</b>	22.1 <sup>b</sup>	31.7 <sup>b</sup>
<b>b</b>	36.9	48.2
<b>c</b>	61.9	74.0



**a**



**b**



**c**

<sup>a</sup> Grey portions of the Lewis structures represents the additional Se=O bond present in Se(VI) structures only. <sup>b</sup> The H<sub>2</sub>O<sub>2</sub> addition process sketched in **a** is represented for comparison with **b** and **c**, with its relative activation energy (TS1). The overall barrier towards peroxyseleonic/peroxyselenonic acid is slightly higher (TS2), corresponding to dehydration of the selenurane (I). However, it remains much lower than both processes **b** and **c**. Level of theory: COSMO-M06//OPBE.

### The oxidation of aniline to nitrobenzene

In the previous section, we showed that Se(VI) activates hydrogen peroxide in the form of a peroxyacid less efficiently than Se(IV). Nevertheless, in the context of a catalytic mechanism, to properly assess the performances of each oxidation state in aniline conversion, the full mechanism must be





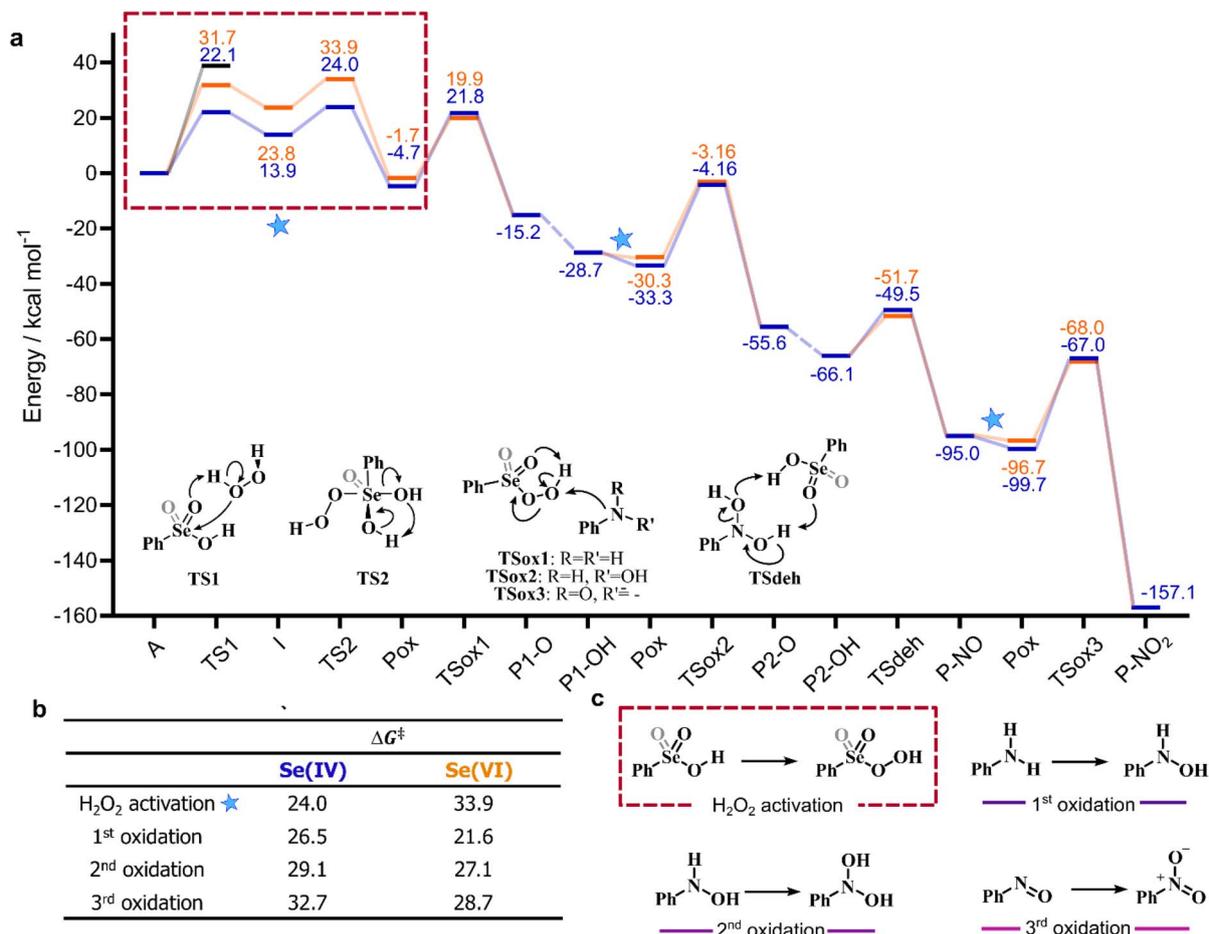


Fig. 2 (a) Gibbs free energy profile (kcal mol<sup>-1</sup>) for the oxidation of aniline to nitrobenzene: Se(IV) mediated process (blue); Se(VI) mediated process (orange). All energies are relative to the free seleninic/selenonic acid and H<sub>2</sub>O<sub>2</sub>. The black line represents the activation energy for direct aniline oxidation by H<sub>2</sub>O<sub>2</sub> (*i.e.*, the uncatalyzed process). Level of theory: COSMO-M06//OPBE. Lines are intended as a guide to the eye only. Only the first H<sub>2</sub>O<sub>2</sub> activation step is represented (within a dashed red box); the subsequent analogous steps are omitted, and their position is indicated by a star. Dashed lines are acid/base processes occurring without or with very low activation energies. (b) Activation energies (kcal mol<sup>-1</sup>) for the key steps of the mechanism. (c) Schematic representation of the H<sub>2</sub>O<sub>2</sub> activation process and of the three consecutive oxidative steps.

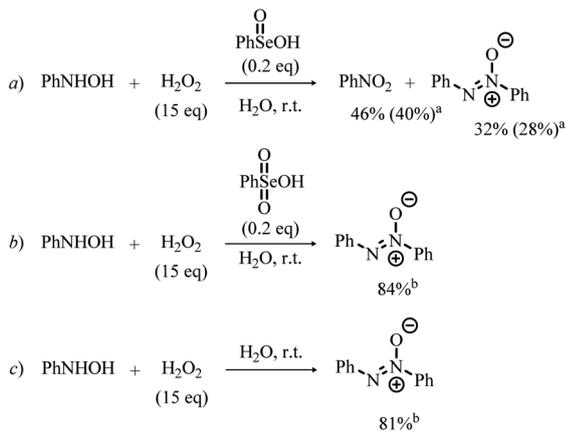
acid. In both cases, the acid (A) reacts first with H<sub>2</sub>O<sub>2</sub> to produce the corresponding peroxyacid (Pox). (*vide supra*). Three consecutive catalytic cycles progressively lead to the oxidation of aniline to the corresponding mono hydroxylamine (P1-OH), dihydroxylamine (P2-OH) and lastly to nitrobenzene (P-NO<sub>2</sub>). In each cycle, the peracid generated by H<sub>2</sub>O<sub>2</sub> activation (Fig. 2c) reacts with the substrate in a S<sub>N</sub>2 fashion, producing the *N*-oxide products (P1-O and P2-O) and the corresponding seleninic or selenonic acid. Under the acidic conditions in which the reaction takes place, the primary *N*-oxide is easily protonated by either **1** or **4** on the O position, and then easily deprotonated on the N position, leading to the mono hydroxyl amine and dihydroxylamine (P1-OH and P2-OH). For aniline *N*-oxide (P1-O), the process was found to proceed without any appreciable activation energy, while, for aniline hydroxylamine *N*-oxide (P2-O), the process occurs with a low activation energy of less than 2 kcal mol<sup>-1</sup> (Table S2†).

The involvement of *N*-phenylhydroxylamine in the mechanism of the Se-catalyzed oxidation of anilines was also

confirmed through the control experiments summarized in Scheme 7. In the presence of catalytic amounts of benzeneseleninic acid, *N*-phenylhydroxylamine reacted with H<sub>2</sub>O<sub>2</sub> to afford nitrobenzene, although significant amounts of the related azoxybenzene (diphenyldiazene oxide) was formed (Scheme 7, reaction a). Remarkably, the reaction of *N*-phenylhydroxylamine with H<sub>2</sub>O<sub>2</sub> in the presence of benzeneselenonic acid (Scheme 7, reaction b) as well as the control experiment conducted under Se-free conditions (Scheme 7, reaction c) did not afford nitrobenzene and only diphenyldiazene oxide was detected.

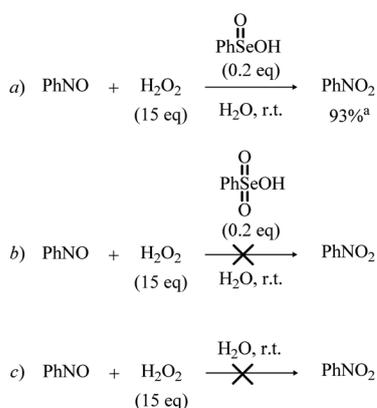
Before the 3<sup>rd</sup> catalytic cycle, P2-OH undergoes an acid catalyzed dehydration reaction to nitrosobenzene (P-NO), in which either **1** or **4** acts as amphoteric acid/base catalyst with an overall activation energy of 16.6 and 14.4 kcal mol<sup>-1</sup>, respectively, and a reaction energy of -28.9 kcal mol<sup>-1</sup>. Lastly, in the final oxidative cycle, P-NO is oxidized to the final species, nitrobenzene (P-NO<sub>2</sub>). The potentially competitive direct oxidation of P2-OH to the corresponding *N*-oxide followed by





**Scheme 7** Reaction of *N*-phenylhydroxylamine with  $\text{H}_2\text{O}_2$  in the presence of benzeneseleninic acid (reaction a), benzeneselenonic acid (reaction b) and Se-free control experiment (reaction c). <sup>a</sup>Conversion is reported (isolated yield is given in parentheses). <sup>b</sup>Isolated yield is given.

dehydration to P-NO<sub>2</sub> was found to be less favored kinetically than the dehydration to P-NO, even if it has a lower activation energy than the oxidation of P-NO to P-NO<sub>2</sub>. (Table S3†) Thus, Se-catalyzed dehydration to P-NO was deemed more likely. Additionally, the formation of P-NO was confirmed in previous Se-catalyzed oxidations of aniline, and it is likely the key intermediate in the formation of azoxy benzene, one of the side products of the reaction.<sup>11</sup> Direct evidence of the role of nitrosobenzene as a reaction intermediate in the Se-catalyzed process was provided by the control experiments reported in Scheme 8. Indeed, nitrobenzene was directly achieved in good yield upon treatment of nitrosobenzene with  $\text{H}_2\text{O}_2$  in the presence of a catalytic amount of benzeneseleninic acid (Scheme 8, reaction a). On the other hand, the reactions of Scheme 8 further highlighted the key role of Se(IV) species in the oxidation of aniline. The plausible nitroso intermediate was recovered unreacted after treatment with  $\text{H}_2\text{O}_2$  in the presence of Se(VI) catalysts such as benzeneselenonic acid (Scheme 8,



**Scheme 8** Reaction of nitrosobenzene with  $\text{H}_2\text{O}_2$  in the presence of benzeneseleninic acid (reaction a), benzeneselenonic acid (reaction b) and the Se-free control experiment (reaction c). <sup>a</sup>Isolated yield is given.

reaction b). As expected, no oxidation occurred in the absence of selenium-containing species (Scheme 8, reaction c).

The direct participation of this peroxy selenurane in the oxidation of the substrate was excluded on the basis of the activation energy of the process, *e.g.*, for Se(IV), aniline oxidation to P1-O occurs with an activation energy of *ca.* 45 kcal mol<sup>-1</sup> and of *ca.* 26 kcal mol<sup>-1</sup> when the peroxy selenurane (I), and the peroxyacid (Pox) acts as the oxidant, respectively (Table S4†).

Following  $\text{H}_2\text{O}_2$  activation, the three oxidations occur with increasing activation energies, regardless of the oxidation state of the catalyst. That is, oxidation of aniline has the lowest activation energy (Fig. 2b, 1st oxidation) and oxidation of P-NO the highest one (Fig. 2b, 3rd oxidation). Intuitively, since the amine acts as the nucleophile in the reaction, each increment of the N oxidation state reduces the nucleophilicity of the substrate due to the electron-withdrawing nature of each new binding oxygen group. Indeed, the Hirshfeld partial charge<sup>19,20</sup> on N increases from  $-0.212$  to  $-0.088$  and to  $-0.003$  going from aniline to P1-OH to P-NO, thus highlighting a reduced nucleophilicity of the oxidized intermediates. Similarly, the barrier for the three acid-catalyzed processes increases, *i.e.*, while the isomerization of P1-O proceeds without any appreciable activation energy, and the conversion of P2-O to P2-OH has a low activation energy of less than 2 kcal mol<sup>-1</sup> when seleninic acid **1** act as the acid catalyst, the dehydration of P2-OH to P-NO has an activation energy of 14–16 kcal mol<sup>-1</sup>. Thus, excluding the two previous almost barrierless acid catalyzed processes, this step has the lowest barrier of the overall mechanism by 10 kcal mol<sup>-1</sup> or more for both oxidation states.

Overall, the whole process from aniline to nitrobenzene is strongly exergonic by more than 150 kcal mol<sup>-1</sup>. All the reaction steps of the mechanism are exergonic or almost isergonic, apart from the  $\text{H}_2\text{O}_2$  addition to **1** or **4**, leading to the formation of the intermediate peroxy selenurane (I) which is destabilized with respect to the corresponding acids. In contrast, the reaction energy of each sequential oxidation becomes more and more negative along the pathway, starting from a  $\Delta G_r$  of  $-28.6$  kcal mol<sup>-1</sup> for the oxidation of aniline to P1-OH and reaching a  $\Delta G_r$  of  $-62.1$  for the final oxidation of P-NO to P-NO<sub>2</sub>.

### Comparison between Se(IV) and Se(VI) cycles

The overall representation of the energy profile leading to the conversion of aniline to P-NO<sub>2</sub>, in which a clear comparison between the performance of Se(IV) and Se(VI) species can be made, is shown in Fig. 2a. As initially highlighted, Se(IV) outperforms Se(VI) in the hydrogen peroxide activation step, leading to a more stable peroxyacid with a lower activation energy (Fig. 2b). Conversely, Se(VI) appears to perform consistently better in each of the following reaction steps, *i.e.*, the three substrate oxidations and the Se-catalyzed dehydration of P2-OH to P-NO. Focusing on the three former reactions, activation energies for Se(VI) mediated processes are 2–5 kcal mol<sup>-1</sup> lower than those for the corresponding Se(IV) mediated processes. Thus, peroxyseleonic acid **5** is a better oxidant from the kinetic point of view than peroxyseleonic acid **3**. Part of this activation energy lowering is clearly due to the lower stability of



peroxyselenonic than selenonic acid (as compared to the couple peroxyseleninic/seleninic acid) assessed on the basis of their  $\Delta G_{\ddagger}$ , as previously mentioned. Thus, while  $\text{Se}(\text{vi})$  appears to be the best oxidant for the reactions under investigation,  $\text{Se}(\text{iv})$  appears to be the best species in the  $\text{H}_2\text{O}_2$  activation process, with a much higher energy gap between the two PESs in this latter case.

### Theoretical comparison of the catalytic performance

While the analysis of the PESs already provides valuable insight into the catalytic potential of  $\text{Se}(\text{iv})$  and  $\text{Se}(\text{vi})$  species, a much more quantitative index of their catalytic potential is represented by the calculated turnover frequency (TOF) of each catalytic cycle, when the two different catalysts, *i.e.* seleninic and selenonic acid, are employed. This quantity is directly related to an experimentally accessible parameter, and accounts for all steps of each cycle at once in the evaluation of the performance of the catalyst. Within the energetic span model proposed by the Kozuch and Shaik, who elaborated the idea of Amatore and Jutand,<sup>21</sup> it is possible, starting from the energy landscape constructed with quantum mechanics calculations, to obtain a well-defined TOF value.<sup>22–24</sup> Additionally, this procedure allows to gain insight into the states which mostly (or totally) determine the value of the TOF, *i.e.* the TOF determining transition state (TDTS) and the TOF determining intermediate (TDI) (see additional Computational details in the ESI† for further information).<sup>25</sup>

In Fig. 3, a representative close-up of the first catalytic cycle is illustrated; the TDI and the TDTS are shown for both  $\text{Se}(\text{iv})$  and  $\text{Se}(\text{vi})$  catalyzed processes. The two downhill cycles bear close similarity until Pox, differing only in the substrate oxidations. Notably, the TDI and TDTS positions do not change in the other two cycles. For the  $\text{Se}(\text{vi})$  catalyzed processes, the TDI/TDTS are always identified within the  $\text{H}_2\text{O}_2$  activation step corresponding respectively to the selenonic acid and to the  $\text{Se}(\text{vi})$  peroxyselenurane dehydration (A and TS2 in Fig. 3). For all three  $\text{Se}(\text{vi})$

catalyzed oxidations, this step has the overall largest transition state – minimum energy difference, *i.e.* the energetic span of the cycle.

Conversely, for the  $\text{Se}(\text{iv})$  catalyzed processes, the TDI/TDTS are always identified within the substrate oxidation step, corresponding to the peroxyacid and to the  $\text{S}_{\text{N}}2$  oxygen-transfer transition state (Pox and TSox in Fig. 3), respectively. In the two downhill cycles, as previously outlined, this couple remains the TDI/TDTS, but since the three oxidations appear to have progressively increasing activation energies (Fig. 2b) the energetic span of  $\text{Se}(\text{iv})$  cycles increases along the overall mechanism, with the first cycle having the lowest and the last having the largest span, always corresponding to the substrate oxidation activation energy.

To verify how these mechanistic differences affect the overall process, the six TOFs for the three  $\text{Se}(\text{iv})/\text{Se}(\text{vi})$  catalyzed oxidations were computed, and their ratio was evaluated (Table 2) to quantify the different performance of  $\text{Se}(\text{vi})$  and  $\text{Se}(\text{iv})$  catalysis.

From these results, it emerges that the differences between  $\text{Se}(\text{vi})$  and  $\text{Se}(\text{iv})$  as catalysts go attenuating along the overall mechanism, with the greatest differences in the first cycle (highest TOF ratio) and the smallest differences in the last cycle (lowest TOF ratio). This result comes from the different nature of the TDI/TDTS couples when  $\text{Se}(\text{vi})$  selenonic and  $\text{Se}(\text{iv})$  seleninic acids are the catalysts, as above described. Indeed, since for  $\text{Se}(\text{vi})$  the TDI and the TDTS remain the same in the three cycles, and correspond to the same energetic span, all three  $\text{Se}(\text{vi})$  catalyzed processes have the same TOF. Conversely, since the activation energy of the three substrate oxidations increases along the mechanism, and this activation energy matches to the energetic span of  $\text{Se}(\text{iv})$  catalyzed processes, their TOF becomes lower along the overall conversion from aniline to nitrobenzene, thus narrowing down the TOF ratio from *ca.*  $10^{-6}$  to *ca.*  $10^{-1}$ . Most importantly, the data reported in Table 2 clearly reveal that  $\text{Se}(\text{iv})$  is always a better catalyst for aniline oxidation than the corresponding  $\text{Se}(\text{vi})$  species, since in all cases the TOF ratio is smaller than 1 by at least one order of magnitude. Similar conclusions hold true at different levels of theory (Fig. S2, Tables S5, S6, and relative discussion†). These results further corroborate the privileged role of seleninic acid 1 in the organoselenium catalyzed oxidation of aniline to nitrobenzene, that is, peroxyseleninic acid 3 appears to be a much better oxygen-transfer catalyst than peroxyselenonic acid 5, even if the latter is in principle a better oxidizing agent (Fig. 2b). The central role of  $\text{Se}(\text{iv})$  catalytic species in the oxidation of plausible reaction

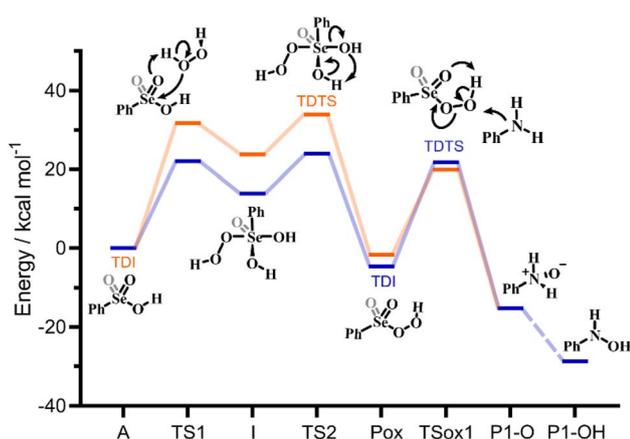


Fig. 3 Gibbs free energy profile, intermediates, transition states, and calculated TDI and TDTS for the 1st catalytic cycle:  $\text{Se}(\text{iv})$  mediated process (blue);  $\text{Se}(\text{vi})$  mediated process (orange). Grey portions on the Lewis structures represent the additional Se–O bond which is present only in  $\text{Se}(\text{vi})$  structures. Level of theory: COSMO-M06//OPBE.

Table 2 TOF ratio between  $\text{Se}(\text{vi})$  and  $\text{Se}(\text{iv})$  catalyzed oxidations for each catalytic cycle. Level of theory: COSMO-M06//OPBE

	$\text{TOF}^{\text{Se}(\text{VI})}$
	$\text{TOF}^{\text{Se}(\text{IV})}$
1st cycle	$3 \cdot 10^{-6}$
2nd cycle	$3 \cdot 10^{-4}$
3rd cycle	$1 \cdot 10^{-1}$



intermediates is further highlighted by the control experiments reported in Schemes 7 and 8, as previously highlighted.

### Insight into Se(IV) to Se(VI) interconversion

While the data above discussed highlight the privileged role of the Se(IV) oxidation state in the reaction under investigation, one could argue that Se(IV) seleninic acid cannot catalyze the three reactions simply because it is rapidly oxidized to Se(VI) selenonic acid under the conditions in which the reaction takes place.

According to Table 2, this conversion would result in a catalyst inactivation. However, we followed this hypothesis, since Back and coworkers reported the fast formation of Se(VI) species when diphenyl diselenide is treated with H<sub>2</sub>O<sub>2</sub>,<sup>6</sup> and Tanini and coworkers observed the formation of Se(VI) selenonic acid in the water recovered after the complete oxidation of aniline to nitrobenzene.<sup>11</sup>

Guided by the inspiring work of Back *et al.*, we first probed the mechanistic hypothesis that the peroxy selenurane (I) on the Se(IV) PES acts as the key intermediate in the interconversion. Indeed, it was previously postulated that while a non-redox dehydration would lead to peroxyacid, as previously described, an alternative redox dehydration should lead to selenonic acid (Scheme 9a). Unfortunately, while such interconversion is strongly favored thermodynamically, it proved to be quite troublesome kinetically (Fig. 4).

Particularly, with respect to peroxyseleonic acid (Pox in Fig. 4), the formation of selenonic acid (A in Fig. 4) is exergonic by *ca.* 30 kcal mol<sup>-1</sup>. As previously mentioned, the peroxy selenurane (I) is destabilized by *ca.* 13 kcal mol<sup>-1</sup> with respect to seleninic acid and its dehydration to peroxyseleonic acids occurs thorough a low-lying transition state (TS2). Conversely, its dehydration to selenonic acid goes through a transition state TS3 located way higher on the PES, such that the process is kinetically disfavored by *ca.* 24.40 kcal mol<sup>-1</sup>. Overall, with respect to the Se(IV) peroxyseleonic acid Pox, TS3 is located 53.1 kcal mol<sup>-1</sup> higher in energy. Thus, while the

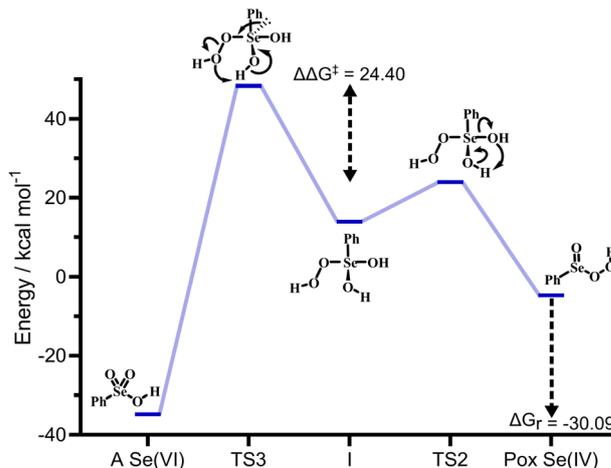
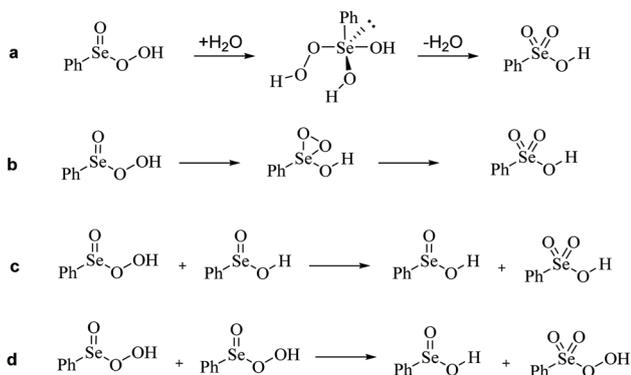


Fig. 4 Gibbs free energy profile (kcal mol<sup>-1</sup>) and relative structures for the interconversion of peroxyseleonic acid (Pox) to selenonic acid (A). All energies are relative to seleninic acid and H<sub>2</sub>O<sub>2</sub>. Level of theory: COSMO-M06//OPBE.

peroxyseleuranine is a key intermediate in H<sub>2</sub>O<sub>2</sub> activation, it does not seem to be involved in the interconversion between Se(IV) and Se(VI). Intrigued by this result, we explored a couple of alternative plausible mechanisms. To the best of our knowledge, the question of the formation of selenonic acid was first tentatively addressed by Syper and coworkers in 1987,<sup>26</sup> who invoked a peroxy transition state/intermediate already envisioned by Sharpless and Hori in 1978 (Scheme 9b).<sup>4</sup>

A transition state for the formation of such peroxy intermediate was found to have an activation energy of 52.5 kcal mol<sup>-1</sup> with respect to peroxyseleonic acid, not much lower than the redox-dehydration, which has an overall activation energy of 53.1 kcal mol<sup>-1</sup> with respect to peroxyseleonic acid. Thus, these two processes were deemed equally unlikely to occur. Further evolution of the peroxy intermediate to selenonic acid was not investigated, because the barrier for its formation was prohibitively high.

Since the active role of peroxyseleonic acid in the autocatalytic oxidation of organoselenides was observed in the past,<sup>15</sup> we checked whether a similar "autocatalytic" oxidation might operate also in this case (Scheme 9c). Such process, in which one equivalent of peroxyseleonic acid 3 oxidizes some residual seleninic acid 1 to selenonic acid 4, while being reduced back to seleninic acid, occurs with an activation energy of 37.5 kcal mol<sup>-1</sup> at our level of theory, way more favorable than the two previous proposed mechanisms, but still higher than the H<sub>2</sub>O<sub>2</sub> activation process of seleninic acid, which has an overall barrier of 23.98 kcal mol<sup>-1</sup> (Fig. 2b). Thus, such a process does not appear to be autocatalytic in nature. Most importantly, since all mechanistic proposals in Scheme 9a-c have an activation energy higher than each of the three substrate oxidations by peroxyseleonic acid (Fig. 2b), we conclude that no conversion to selenonic acid should occur, as long as there is some substrate left to undergo catalytic oxidation. This is consistent with the hypothesis by Tanini *et al.* that selenonic acid forms by overoxidation of the catalyst after all



Scheme 9 Plausible pathways for Se(IV) to Se(VI) interconversion. (a) Redox dehydration of peroxy selenurane. (b) Syper hypothesis, through a peroxy transition state/intermediate. (c) Direct oxidation of seleninic acid by peroxyseleonic acid. (d) Dismutation of peroxyseleonic acid to seleninic and peroxyseleonic acid.



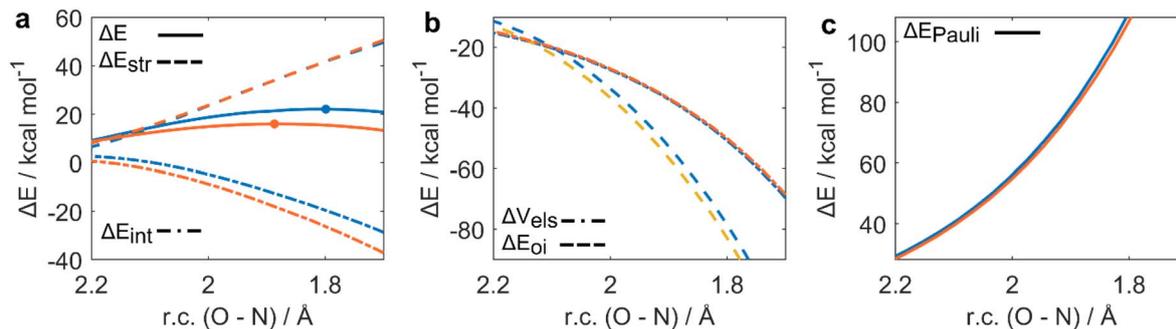


Fig. 5 Activation strain (a) and energy decomposition analysis (b and c) of aniline oxidation to aniline *N*-oxide (P1-O) by peroxyseleninic (blue) and peroxyselenonic (orange) acids. Level of theory: M06//OPBE.

aniline and derivatives have been fully oxidized to the final product.<sup>11</sup>

Lastly, we checked whether peroxyselenonic acid **3** might undergo a self-oxidation reaction directly to peroxyselenonic acid **5**, thus bypassing the high activation energy required by selenonic acid in H<sub>2</sub>O<sub>2</sub> activation (Scheme 9d). Selenonic acid would be finally produced from the oxidation of the organic substrate, which is a favored process (Fig. 2b). However, also this self-oxidation occurs with a rather high activation energy of 39.8 kcal mol<sup>-1</sup> at our level of theory, higher than that of the previously described oxidation to selenonic acid. Thus, also this process is not expected to take place in the presence of aniline and its intermediates towards nitrobenzene.

#### Activation strain and energy decomposition analysis of substrate oxidation

While we previously discussed the privileged nature of Se(IV) as both an hydrogen peroxide activator and an oxygen-transfer catalyst, our calculations also highlight its worst performance in the effective organic substrate oxidation. To fully untangle

the role of the oxidation state in organoselenium catalyzed oxygen-transfers, a clear molecular picture of the peroxyseleninic and peroxyselenonic acid performances in the actual oxygen-transfer step is desirable. To provide a quantitative discussion on the effect of the oxidation state on the S<sub>N</sub>2-like oxidation of aniline, ASA and EDA were performed (see Computational methods). Particularly, the oxidation of aniline to aniline-*N*-oxide by peroxyseleninic and peroxyselenonic acid was compared to provide a detailed insight into the different oxidation state effect in the 1st catalytic cycle. The oxidative potential of the two peroxyacids was analyzed (Fig. 5a-c).

In the actual oxygen-transfer step, peroxyselenonic acid acts as the best oxidant. To analyze this step, ASA was plotted along the O-N distance, which undergoes a well-defined change along the reaction. Analogous conclusions can be drawn by investigating the complementary reaction coordinate *i.e.*, O-O bond breaking (Fig. S1†). ASA (Fig. 5a) reveals how for the two reactions, similar values of  $\Delta E_{\text{str}}$  occur at consistent points of r.c. Conversely, the  $\Delta E_{\text{int}}$  value alone accounts for the trend in activation energy, being systematically more stabilizing for Se(VI) than for Se(IV), and thus correlating with the lower  $\Delta E$  of

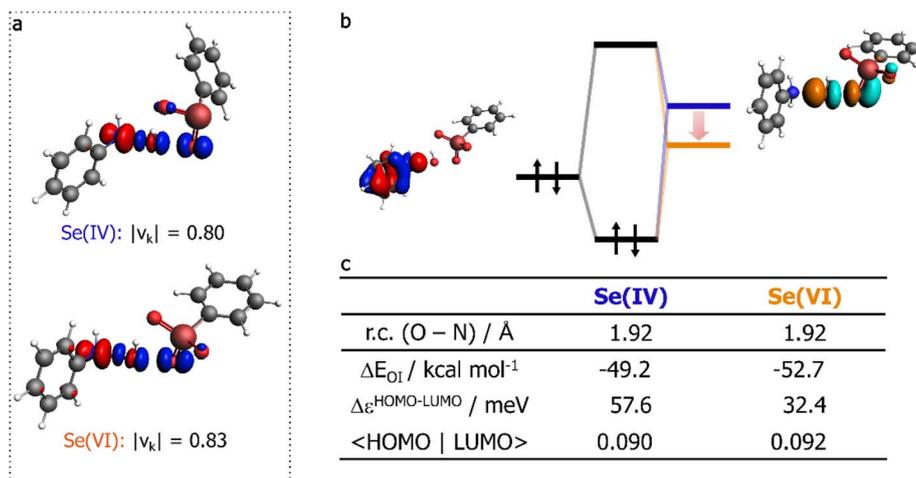


Fig. 6 Analysis at consistent geometries, r.c. (O-N) = 1.92 Å. (a) Deformation density associated with the NOCVs with the largest eigenvalue ( $\nu_k$ ). Red areas are associated with charge depletion, while blue lobes are associated with charge accumulation. (b) Schematic representation of the HOMO (aniline)-LUMO(peroxyacid) orbital interaction and KS-MO for Se(VI), isodensity value: (0.03). (c)  $\Delta E_{\text{OI}}$ , HOMO(aniline)-LUMO(peroxyacid) energy gap ( $\Delta \epsilon^{\text{HOMO-LUMO}}$ ) and HOMO(aniline)-LUMO(peroxyacid) orbital overlap ( $\langle \text{HOMO} | \text{LUMO} \rangle$ ). Level of theory: M06//OPBE.



the former. To understand what factors are responsible for this effect,  $\Delta E_{\text{int}}$  was partitioned according to EDA (Fig. 5b and c). The main factor responsible for the lower  $\Delta E_{\text{int}}$  of Se(vi) is  $\Delta E_{\text{OI}}$ , which is consistently more stabilizing for peroxyseleonic acid.

An EDA-NOCV calculation was performed to characterize the orbital interaction between the two reactants. Within the EDA-NOCV approach,  $\Delta E_{\text{OI}}$  can be further decomposed into contributions associated with NOCV. For each NOCV pair, a deformation density can be visualized, associated with a certain charge transfer degree from one fragment to the other (see Computational methods).

The deformation density associated with the NOCV with the greater eigenvalue (*i.e.*, to the greatest charge transfer between the two fragments), is represented in Fig. 6a. It appears that for both Se(IV) and Se(vi) the strongest orbital interaction leads to a charge depletion from the nitrogen lone pair, and to charge accumulation on the oxygen atom of the peroxy bond directly bonded to Se, and in the O–N internuclear region, due to the new bond formation. This representation is consistent with the  $S_{\text{N}}2$ -like nature of the reaction, in which aniline acts as the nucleophile and the peroxyacid as the electrophile. Additionally, the absolute value of the eigenvalue associated with the NOCV of the Se(vi) reaction is greater than that for Se(IV), being equal to 0.83 and 0.80, respectively, thus unveiling a higher charge-transfer character for the former. Since the nucleophile is the same, this can be associated with the better electron-accepting properties of the peroxyacid in the highest OS. In fact, the couple of canonical Kohn–Sham molecular orbitals (KS-MO) mainly associated with this charge-transfer interaction can be identified with the HOMO–LUMO couple, in which the HOMO of the nucleophile (mostly the nitrogen lone pair) interacts with the LUMO of the electrophile (mostly localized on the peroxide bond). The peroxyacid of Se(vi) has a lower LUMO that better matches to the energy of aniline's HOMO (Fig. 6b and c). Overall, this is the main factor responsible for the stronger Se(vi) orbital interaction, rationalizing the better performance of peroxyseleonic acid in the substrate oxidation step, if it is indeed produced in the reaction mixture.

## Conclusions

In this work, an extensive theoretical mechanistic investigation has been carried out on the on-water organoselenium catalyzed oxidation of aniline to nitrobenzene using  $\text{H}_2\text{O}_2$  as the final oxidant, with particular attention to the  $\text{H}_2\text{O}_2$  activation step. A careful analysis of the PES, carried out in the framework of the energetic span model, showed how the Se(vi) selenonic acid catalyst has a poorer catalytic performance (evaluated on the basis of calculated TOFs) in all three catalytic cycles progressively oxidizing aniline to nitrobenzene, highlighting the privileged role of Se(IV) seleninic acid species in the title catalytic oxygen-transfer reaction.

Additionally, our analysis pinpointed how peroxyseleonic acid 5 is indeed a stronger oxidant (at least from the kinetic point of view) than peroxyseleonic acid 3, since all three substrate oxidations occur with a lower activation energy. Thus, if peroxyseleonic acid 5 is indeed produced, a faster reactivity is

envisioned. Conversely, selenonic acid 4 performs worse than seleninic acid 1 in  $\text{H}_2\text{O}_2$  activation, thus rationalizing its poor catalytic performance. Through ASA, we interpreted the sluggish  $\text{H}_2\text{O}_2$  activation by selenonic acid as an effect of the higher distortion associated with this step when compared to the analogous one for seleninic acid. Conversely, the enhanced electrophilicity of peroxyseleonic acid was associated with its reinforced orbital interaction, mainly granted by its low-lying LUMO. Nevertheless, it is important to stress once again that these two opposite effects (*i.e.*, Se(IV) has a lower  $\Delta E_{\text{str}}$  in the  $\text{H}_2\text{O}_2$  activation step, and Se(vi) has a stronger  $\Delta E_{\text{OI}}$  in the substrate oxidation step) lead to a TOF which remains consistently in favor of the Se(IV) oxidation state. Thus, in the context of a catalytic mechanism, the less favorable orbital matching of peroxyseleonic acid with the organic substrate is favorably counterbalanced by the lower distortion associated with its  $\text{H}_2\text{O}_2$  activation step, which makes the Se(IV) reaction pathway overall more productive.

Lastly, the conversion of seleninic to selenonic acid under oxidizing conditions was found to proceed with a higher activation energy than the three substrate oxidations by peroxyseleonic acid. Thus, the formation of the Se(vi) species is expected only when the substrate is fully converted to nitrobenzene.

This work provides a detailed mechanistic picture on the catalytic oxygen-transfer behavior of the popular organoselenium catalysts derived from the commercial diphenyl diselenide, shedding light on the physico-chemical role of the oxidation state in their catalytic performance, spanning from the  $\text{H}_2\text{O}_2$  activation to their oxidative potential toward the organic substrate. Additionally, to the best of our knowledge, this is the first comprehensive theoretical attempt to rationalize the oxygen-transfer catalytic cycles of seleninic acid and derivatives, providing a theoretical complementary basis to the existing experimental knowledge and tackling the general problem of identifying the active and inactive intermediates in organoselenium catalyzed reactions. Based on these results, we can hypothesize that the way in which the specific organic substrate and the reaction conditions (*e.g.*, choice of the solvent, homogeneous *vs.* heterogeneous conditions) affects the balance between the hydrogen peroxide activation step and the actual substrate oxidation step might shift the catalysis either toward Se(IV) or Se(vi) based catalytic cycles. Indeed, if Se(vi) can be formed competitively with the substrate oxidation, and the environment somewhat facilitates Se(vi) hydrogen peroxide activation, both oxidation states might contribute to the catalysis. How this balance is modified in other organoselenium catalyzed oxygen transfer is currently under investigation.

## Experimental section

### Synthetic procedures

Aniline, nitrosobenzene, and benzeneseleninic acid were purchased and used as received, without further purification. Benzeneselenonic acid was prepared according to the literature.<sup>6,11,26</sup> *N*-Phenylhydroxylamine was synthesized upon reduction of nitrobenzene.<sup>27</sup> Experimental details are reported in the ESI.†



## Computational details

DFT calculations were carried out with the ADF2019.307 software.<sup>28,29</sup> All geometries were optimized employing OPBE density functional<sup>30</sup> combined with a small-core TZ2P basis set. Scalar relativistic corrections have been included in all calculations at the zeroth-order regular-approximation (ZORA) as implemented in ADF.<sup>31</sup> Thus, all optimizations are at the ZORA-OPBE/TZ2P level of theory. This level of theory was found to properly reproduce geometrical features of organochalcogenides<sup>32</sup> and is recognized as one of the best GGAs to tackle S<sub>N</sub>2 reactions,<sup>30,33</sup> such as the oxidations studied in this work. Vibrational analysis was performed to confirm the nature of each optimized geometry: minima displayed all positive frequencies, while transition states displayed one imaginary frequency associated with the motion of the atoms along the reaction. Accurate single points energies were then computed with two different functionals, the hybrid OPBE0 and the meta-hybrid M06,<sup>34,35</sup> combined with an all-electron TZ2P basis set. Both functionals were previously benchmarked on organochalcogen reactions in different oxidation states and provided results within a couple of kcal mol<sup>-1</sup> when compared against highly correlated reference calculations.<sup>36</sup> OPBE, OPBE0 and M06 provide a consistent, qualitatively analogous, description of the PES (Tables S5, S6 and Fig. S2†). Along the manuscript, only M06 energies are discussed. Solvation effects were included employing the conductor-like screening model (COSMO) as implemented in ADF,<sup>37,38</sup> using the standard parameters (dielectric constant and effective radius) for water. The radii of the atoms were taken to be the MM3 van der Waals radii divided by 1.2, as per default in ADF. Thermodynamics corrections were evaluated at the ZORA-OPBE/TZ2P level of theory under perfect gas approximation using standard statistical thermodynamic relationships and added to the electronic energy calculated at the COSMO-ZORA-M06/TZ2P-ae level of theory. This level of theory is labelled M06//OPBE.

DLPNO-CCSD(T) calculations with the cc-pVTZ-DK basis set and Douglas-Kroll-Hess (DKH) scalar relativistic correction<sup>39,40</sup> have been performed on critical steps of the reaction mechanism to corroborate the DFT based discussion.<sup>41</sup> The electronic energy was obtained by DLPNO-CCSD(T) single point calculations with TightPNO criteria, using ZORA-OPBE/TZ2P optimized geometries. Orca 4.2 was used for all DLPNO-CCSD(T) calculations.<sup>42,43</sup> The thermodynamic correction and solvation energy were estimated at the ZORA-OPBE/TZ2P level of theory, and COSMO-ZORA-OPBE/TZ2P level of theory respectively. We label this level of theory DLPNO-CCSD(T)//OPBE. The results are comparable to those obtained at the DFT level of theory employed in the manuscript and in the ESI.† (Table S7 and S8†). DLPNO-CCSD(T)//OPBE calculations were also used to probe the effect of one water molecule in the reaction mechanism, which was found to be modest (Table S9†).

Activation strain analysis (ASA, eqn (1)) and energy decomposition analysis (EDA, eqn (2)) were applied along a suitable reaction coordinate (r.c.)  $\zeta$  to gain further insight into the factors at the origin of the investigated reactivity.<sup>44–48</sup>

$$\Delta E(\zeta) = \Delta E_{\text{strain}}(\zeta) + \Delta E_{\text{int}}(\zeta) \quad (1)$$

Within ASA, any energy along a reaction profile can be decomposed in a fragment-based approach in a distortion contribution ( $\Delta E_{\text{strain}}(\zeta)$ ) and in an interaction contribution  $\Delta E_{\text{int}}(\zeta)$ . This term can be further partitioned into chemically meaningful contribution according to the EDA scheme:

$$\Delta E_{\text{int}}(\zeta) = \Delta V_{\text{elstat}}(\zeta) + \Delta E_{\text{Pauli}}(\zeta) + \Delta E_{\text{OI}}(\zeta) \quad (2)$$

in which  $\Delta V_{\text{elstat}}(\zeta)$  is the electrostatic interaction between the unperturbed charged densities of the two fragments,  $\Delta E_{\text{Pauli}}(\zeta)$  is the Pauli repulsion which accounts for two center-four electron destabilizing interactions, and  $\Delta E_{\text{OI}}(\zeta)$  is the orbital interaction which accounts for two center-two electron interactions. The most stabilizing orbital interaction was identified and characterized *via* the Natural Orbital for Chemical Valence (NOCV) method.<sup>49–51</sup> Additional computational details can be found in the ESI.†

Along the manuscript, only M06//OPBE results are discussed, since the EDA partitioning scheme here employed is not compatible with DLPNO-CCSD(T) calculations. The absolute value of the TOF at M06//OPBE is reported in the ESI (Table S10†), while in the main text only TOF ratios are discussed.

## Data availability

The datasets supporting this article have been uploaded as part of the ESI.†

## Author contributions

Conceptualization: all the authors; investigation: A. M. (theory and computations) and D. T. (experiments); data curation and formal analysis: A. M., methodology: A. M. and D. T.; writing – original draft: A. M. and D. T.; writing – review & editing: all the authors; fundings and supervision: L. O.

## Conflicts of interest

The authors declare no competing financial interest.

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