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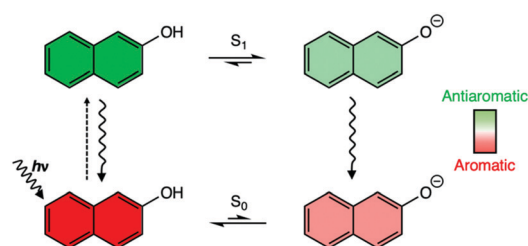
# How does excited-state antiaromaticity affect the acidity strengths of photoacids?<sup>†</sup>

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Photoacids like substituted naphthalenes ( $X = \text{OH}, \text{NH}_3^+, \text{COOH}$ ) are aromatic in the  $S_0$  state and antiaromatic in the  $S_1$  state. Nucleus independent chemical shifts analyses reveal that deprotonation relieves antiaromaticity in the excited conjugate base, and that the degree of “antiaromaticity relief” explains why some photoacids are stronger than others.

Baird first proposed a set of rules suggesting that the electron-counting rules of aromaticity and antiaromaticity reverse in the lowest triplet states of  $\pi$ -conjugated cycles.<sup>1</sup> Based on this set of rules,  $[4n + 2]$   $\pi$ -rings are antiaromatic and  $[4n]$   $\pi$ -rings are aromatic in the first  $\pi\pi^*$  state. These predictions were later extended to the first singlet  $\pi\pi^*$  states ( $S_1$ ), and explained the reactivities of many Baird-type antiaromatic  $[4n + 2]$   $\pi$ -systems.<sup>2–7</sup> Benzene, in the  $S_1$  state, is reactive and readily isomerizes to fulvene.<sup>7,8</sup> Among other  $[4n + 2]$   $\pi$ -ring systems, salicylic acid, in the  $S_1$  state, undergoes intramolecular proton transfer.<sup>9,10</sup> Here we show that, in the  $S_1$  state, differences in the acidity strengths of photoacids might be rationalized by the effects of “antiaromaticity relief” upon deprotonation, followed by a redistribution of electrons in the excited conjugate base.

Following the early independent works of Förster<sup>11,12</sup> and Weller,<sup>13–15</sup> it was recognized that some aromatic acids (e.g., with hydroxyl or ammonium groups) can turn into stronger Brønsted acids in their first excited ( $S_1$ ) states.<sup>11–16</sup> 2-Naphthol, a prototypical organic “photoacid,” is a weak acid in the ground state, but shows enhanced acidity in the  $S_1$  state ( $\text{p}K_a = 9.5$ ,  $\text{p}K_a^* = 2.8$ ,  $\Delta\text{p}K_a = -6.7$ ), and can deprotonate to the solvent producing an electronically excited conjugate base.<sup>15</sup> A Stokes’ shift in the fluorescence spectrum of 2-naphthol in water was interpreted by Förster as radiative decay emitting from the excited conjugate base



**Scheme 1** The Förster cycle of 2-naphthol. As indicated by the color scheme above, 2-naphthol is  $[4n + 2]$   $\pi$ -aromatic in the ground state (red) but becomes  $[4n + 2]$   $\pi$ -antiaromatic in the  $S_1$  state (green). Deprotonation relieves excited-state antiaromaticity, stabilizing the excited conjugate base (light green). In the  $S_0$  state, the conjugate base (light red) is only moderately less aromatic than the acid.

(see depiction of “Förster cycle” in Scheme 1).<sup>11</sup> Yet, despite a large body of theoretical and experimental efforts towards understanding excited-state proton transfer reactions in aromatic acids,<sup>17–27</sup> reasons underlying the occurrence of photoacidity remain unclear. The disparate effects of substituents on photoacidity are even more puzzling. 2-Naphthylammonium displays increased acidity in the  $S_1$  state ( $\text{p}K_a = 4.1$ ,  $\text{p}K_a^* = -0.8$ ,  $\Delta\text{p}K_a = -4.9$ ), deprotonating from an  $\text{NH}_3^+$  group, but the change in acidity,  $\Delta\text{p}K_a$ , is two-folds less than that of 2-naphthol.<sup>28</sup> Aromatic acids, like the 2-naphthoic acid, show the opposite effect and exhibit decreased acidity in the  $S_1$  state ( $\text{p}K_a = 4.2$ ,  $\text{p}K_a^* = 11.5$ ,  $\Delta\text{p}K_a = +7.3$ ) (Table 1).<sup>29</sup>

Why do substituents have such disparate effects on the photoacidities of aromatic acids? Here, we relate the effects of photoacidity to a switch in the ground and excited-state (anti)aromatic character of aromatic acids. According to the Hückel rule, cyclic  $\pi$ -conjugated rings with  $[4n + 2]$   $\pi$ -electrons are aromatic, and those with  $[4n]$   $\pi$ -electrons are antiaromatic.<sup>30</sup> But this electron-counting rule reverses in the first  $\pi\pi^*$  state following Baird’s rule.<sup>1–7</sup> 2-Naphthol is  $[4n + 2]$  Hückel aromatic (ten  $\pi$ -electrons in naphthalene) in the ground state, but becomes  $[4n + 2]$  antiaromatic in the  $S_1$  state. Upon deprotonation, an excited conjugate base forms, and negative charge on the  $\text{O}^-$  delocalizes into the ring, giving rise to a resonance structure with

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<sup>†</sup> Electronic supplementary information (ESI) available: Cartesian coordinates for all structures, computed ground state NICS data, HOMA analyses, and bond alternation analysis for the  $^1\text{L}_a$  and  $^1\text{L}_b$  states of naphthalene. See DOI: 10.1039/d0cc02952a

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**Table 1** Experimental  $pK_a$ ,  $pK_a^*$ , and  $\Delta pK_a$  values

| Compound               | $pK_a$ | $pK_a^*$ | $\Delta pK_a$ | Ref. |
|------------------------|--------|----------|---------------|------|
| 2-Naphthol             | 9.5    | 2.8      | −6.7          | 15   |
| 2-Naphthylammonium     | 4.1    | −0.8     | −4.9          | 28   |
| 2-Naphthoic acid       | 4.2    | 11.5     | +7.3          | 29   |
| 8-Cyano-2-naphthol     | 8.4    | −0.8     | −9.2          | 19   |
| 5,8-Dicyano-2-naphthol | 7.8    | −4.5     | −12.3         | 19   |
| Salicylamide           | −2.6   | −5.3     | −2.7          | 35   |

breached cyclic  $[4n + 2]$   $\pi$ -electron delocalization—this alleviates antiaromaticity in the  $S_1$  state of the acid (see Scheme 1 and resonance structure for the excited conjugate base in Fig. 1a, note delocalization of the negative charge into the ring). In this way, photoacidity might be considered as a consequence of antiaromaticity relief in the  $S_1$  states of aromatic acids.<sup>6</sup> Based on a more bond equalized  $S_1$  vs.  $S_0$  state of 2-naphtholate, Agmon *et al.* pointed out similarly that the excited conjugate base of 2-naphthol might be stabilized by increased aromatic character.<sup>19,20</sup> The effects of ground and excited-state (anti)aromaticity also have been recognized in other excited-state proton transfer processes.<sup>31,32</sup>

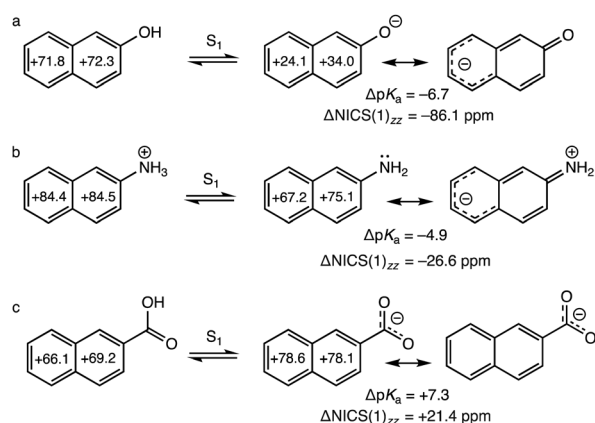
Even though not realized at the time, the possible effects of excited-state antiaromaticity relief were implied in Weller's original explanation (1950's) of photoacidity—argued based on a redistribution of ring  $\pi$ -electrons in the  $S_1$  states of the aromatic acids;<sup>13–15</sup> notably, these ideas were published roughly ten years prior to Dewar<sup>33</sup> and Zimmerman's<sup>34</sup> independent works and Baird's<sup>1</sup> proposal (60's–70's) of a reversed Hückel  $\pi$ -electron-counting rule for aromaticity and antiaromaticity in the first  $\pi\pi^*$  states of transition states<sup>33,34</sup> and of  $\pi$ -conjugated rings.<sup>1</sup> Weller reasoned that when 2-naphthol is electronically excited to the  $S_1$  state, intramolecular charge transfer from the hydroxyl oxygen to the aromatic ring increases acidity of the OH group. Later, it was suggested that even more pronounced charge redistribution happens upon deprotonation (as indicated by shortened C–O bond lengths and changes in dipole moments),<sup>21</sup>

stabilizing the excited conjugate base. We now relate the effects of “charge redistribution” in the excited conjugate base to “relief of excited-state antiaromaticity.” This rationale also may explain why photoacidity only is observed for aromatic acids (*i.e.*, with  $[4n + 2]$  ring  $\pi$ -electrons), but not for other hydroxyl, amine, or ammonium compounds.

Compared to 2-naphthol, the effect of charge redistribution for alleviating antiaromaticity in the excited conjugate base of 2-naphthylammonium is much weaker, since delocalization of a neutral nitrogen lone pair into the naphthalene ring is less effective (see Fig. 1b, note charge separated resonance form). In the  $S_1$  state of 2-naphthylammonium, deprotonation produces a neutral amine ( $NH_2$ ); proton transfer alleviates antiaromaticity of the excited naphthalene ring, but to a lesser degree compared to that of 2-naphthol. Notably, compounds with competing deprotonation sites like salicylamide<sup>35,36</sup> and 3-ammonium-2-naphthol<sup>37</sup> undergo proton transfer from  $NH_3^+$  in the ground state (*i.e.*, to retain aromaticity of the  $\pi$ -ring), but deprotonate from the OH group in the  $S_1$  state (*i.e.*, to alleviate excited-state antiaromaticity of the  $\pi$ -ring) when solvated in water. In the  $S_1$  state of 2-naphthoic acid, deprotonation of the carboxylic group gives a carboxylate ( $COO^-$ ). But negative charge is mostly delocalized between the two oxygen atoms, and does not help lessen excited-state antiaromaticity in the naphthalene ring (Fig. 1c).

We performed dissected nucleus independent chemical shifts,<sup>38,39</sup> NICS(1)<sub>zz</sub>, to quantify excited-state antiaromaticity in the  $S_1$  states<sup>40</sup> of the acids and conjugate bases of 2-naphthol, 2-naphthylammonium, and 2-naphthoic acid (Fig. 1). The computed ring NICS(1)<sub>zz</sub> values of excited 2-naphthol are large and positive (+71.8, +72.3 ppm, strongly antiaromatic) but become much less so in the excited conjugate base (+24.1, +34.0 ppm, weakly antiaromatic), suggesting decreased antiaromaticity upon deprotonation of the excited acid ( $\Delta$ NICS(1)<sub>zz</sub> = −86.1 ppm, Fig. 1a). 2-Naphthylammonium reveals a lesser degree of antiaromaticity relief upon deprotonation ( $\Delta$ NICS(1)<sub>zz</sub> = −26.6 ppm, Fig. 1b). Accordingly, the computed exocyclic CO and CN bond distances of 2-naphthol (1.350 Å) and 2-naphthylammonium (1.476 Å), are longer in the  $S_1$  state acid, and shorter in the excited conjugate base (1.247 Å and 1.373 Å, respectively), indicative of electron delocalization from the deprotonated site into the excited naphthalene ring (see optimized geometries in the ESI†).

In contrast, computed ring NICS(1)<sub>zz</sub> values for 2-naphthoic acid in the  $S_1$  state are positive for the acid (+66.1, +69.2 ppm, strongly antiaromatic) but become even more so in the excited conjugate base (+78.6, +78.1 ppm, strongly antiaromatic), suggesting increased antiaromaticity upon deprotonation of the excited acid ( $\Delta$ NICS(1)<sub>zz</sub> = +21.4 ppm, Fig. 1c). The exocyclic C–C bond of the excited acid is 1.471 Å (*cf.* 1.40 Å CC length of benzene), indicating modest  $\pi$ -conjugation between the carboxylic group and the naphthalene ring. In the excited conjugate base, the exocyclic C–C bond lengthens to 1.534 Å (*cf.* 1.53 Å CC length of ethane), suggesting little resonance between the exocyclic carboxylate group and the excited (antiaromatic) naphthalene (see geometries in the ESI†). In the ground state, deprotonation has less effect on the 10  $\pi$ -electron aromatic character of the naphthalene ring, in 2-naphthol ( $\Delta$ NICS(1)<sub>zz</sub> = +8.5 ppm), 2-naphthylammonium ( $\Delta$ NICS(1)<sub>zz</sub> = +2.2 ppm), and



**Fig. 1** Computed NICS(1)<sub>zz</sub> and  $\Delta$ NICS(1)<sub>zz</sub> (sum of NICS(1)<sub>zz</sub> values of the conjugate base minus that of the acid) (in ppm) values for the  $S_1$  states of the acid and conjugate base at CASSCF(12,12)/6-311+G(d,p), for (a) 2-naphthol, (b) 2-naphthylammonium, and (c) 2-naphthoic acid. Negative  $\Delta$ NICS(1)<sub>zz</sub> values indicate antiaromaticity relief, and positive values indicate antiaromaticity gain, upon formation of the excited conjugate base. Experimental  $\Delta pK_a$  values are included for reference.

2-naphthoic acid ( $\Delta\text{NICS}(1)_{zz} = -1.2$  ppm); positive/negative values indicate aromaticity loss/gain (see full data in the ESI†).

Naphthols with cyano (CN) substituents at the C5 and C8 positions are very strong photoacids: 8-cyano-2-naphthol ( $pK_a = 8.4$ ,  $pK_a^* = -0.8$ ,  $\Delta pK_a = -9.2$ ) and 5,8-dicyano-2-naphthol ( $pK_a = 7.8$ ,  $pK_a^* = -4.5$ ,  $\Delta pK_a = -12.3$ ) show increased acidities of up to 12 units in the  $S_1$  state.<sup>17,18,41,42</sup> These strong photoacids can undergo excited-state proton transfer reactions in methanol, methylsulfonyl, and other organic solvents in the absence of water, first expanding the possibility of studying proton transfer kinetics in non-aqueous solvents.<sup>41</sup> Computed ring NICS( $1)_{zz}$  values for the  $S_1$  state of 8-cyano-2-naphthol and its excited conjugate base ( $\Delta\text{NICS}(1)_{zz} = -137.2$  ppm, Fig. 2a) show significant excited-state antiaromaticity relief upon deprotonation, and the effect in 5,8-dicyano-2-naphthol ( $\Delta\text{NICS}(1)_{zz} = -167.3$  ppm, Fig. 2b) is even greater (*cf.*  $\Delta\text{NICS}(1)_{zz} = -86.1$  ppm, for 2-naphthol). As suggested by the resonance forms in Fig. 2, the electron-withdrawing CN groups help increase charge redistribution in the excited conjugate base, by inductive effects, but also by resonance stabilization (see resonance contributors with negative charges delocalized to the nitrogen atoms). We note that other known strong photoacids with electron-withdrawing groups, *e.g.*, sulfonyls, also have  $\pi$ -systems on the substituents capable of delocalizing negative charge of the excited conjugate base.

Computed deprotonation reaction energies ( $\Delta E$ ) based on the equation:  $\text{ArOH} + \text{H}_2\text{O} \rightarrow \text{ArO}^- + \text{H}_3\text{O}^+$ , document the energetic effects of (anti)aromaticity loss in the  $S_0$  and  $S_1$  states upon deprotonation, and agree with the conclusions based on NICS analyses. Compared to computed  $\Delta E$  values in the  $S_0$  state, deprotonation is less endothermic for the  $S_1$  states of 2-naphthol ( $\Delta\Delta E = -17.99$  kcal mol<sup>-1</sup>,  $\Delta\Delta E = \Delta E(S_1) - \Delta E(S_0)$ ), 2-naphthylammonium ( $-2.77$  kcal mol<sup>-1</sup>), 8-cyano-2-naphthol ( $-28.85$  kcal mol<sup>-1</sup>), and 5,8-dicyano-2-naphthol ( $-29.28$  kcal mol<sup>-1</sup>), but more endothermic for the  $S_1$  states of 2-naphthoic acid ( $+1.39$  kcal mol<sup>-1</sup>) (see full data in the ESI†).

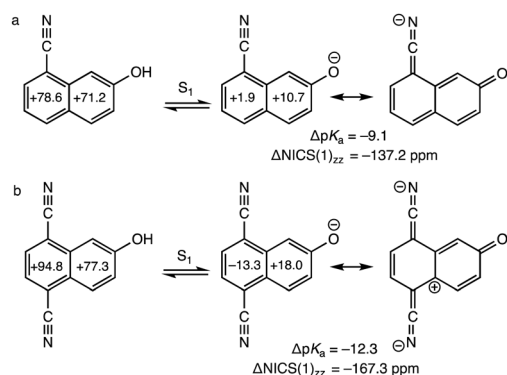


Fig. 2 Computed NICS( $1)_{zz}$  and  $\Delta\text{NICS}(1)_{zz}$  (sum of NICS( $1)_{zz}$  values of the conjugate base minus that of the acid) (in ppm) values for the  $S_1$  states of the acid and conjugate base at CASSCF(12,12)/6-311+G(d,p), for (a) 8-cyano-2-naphthol and (b) 5,8-dicyano-2-naphthol. Negative  $\Delta\text{NICS}(1)_{zz}$  values indicate antiaromaticity relief upon formation of the excited conjugate base. Experimental  $\Delta pK_a$  values are included for reference. Note resonance form showing delocalization of the negative charge into the CN groups.

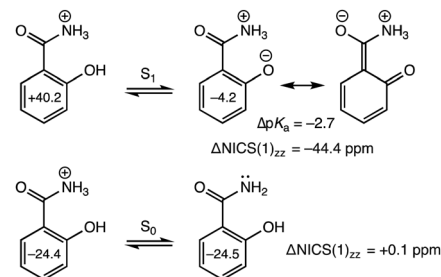
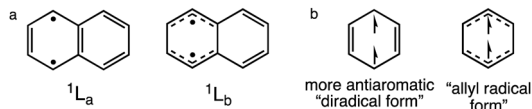


Fig. 3 Computed NICS( $1)_{zz}$  and  $\Delta\text{NICS}(1)_{zz}$  (sum of NICS( $1)_{zz}$  values of the conjugate base minus that of the acid) (in ppm) values for the  $S_0$  and  $S_1$  states of protonated salicylamide and its conjugate base at CASSCF(12,12)/6-311+G(d,p). Negative  $\Delta\text{NICS}(1)_{zz}$  values indicate antiaromaticity relief upon formation of the excited conjugate base. Positive  $\Delta\text{NICS}(1)_{zz}$  values in the ground state indicate aromaticity loss upon formation of the conjugate base. An experimental  $\Delta pK_a$  value is included for reference.

Aromatic acids with competing deprotonation sites like salicylamide undergo proton transfer from  $\text{NH}_3^+$  in the  $S_0$  state, but deprotonate from the OH group in the  $S_1$  state.<sup>35,36</sup> In the  $S_0$  state, computed NICS( $1)_{zz}$  for the acid ( $-24.4$  ppm) and conjugate base ( $-24.5$  ppm, deprotonated at  $\text{NH}_3^+$ ) give nearly the same values ( $\Delta\text{NICS}(1)_{zz} = +0.1$  ppm) (Fig. 3, bottom). But in the  $S_1$  state, the ring NICS( $1)_{zz}$  value for protonated salicylamide is large and positive ( $+40.2$  ppm) while that for the excited conjugate base is modestly negative ( $-4.2$  ppm, deprotonated at OH), documenting the effects of excited-state antiaromaticity relief ( $\Delta\text{NICS}(1)_{zz} = -44.4$  ppm) (Fig. 3, top). Notably, when a proton is removed from the  $\text{NH}_3^+$  site of the electronically excited acid, computed NICS( $1)_{zz}$  for the excited conjugate base ( $+65.8$  ppm) show increased excited-state antiaromaticity in the benzene ring.

A long-standing and much debated anomaly is the stronger photoacidity of 1-naphthol ( $pK_a = 9.2$ ,  $pK_a^* = -0.5$ ,  $\Delta pK_a = -9.7$ ) compared to its structurally similar 2-naphthol isomer ( $pK_a = 9.5$ ,  $pK_a^* = 2.8$ ,  $\Delta pK_a = -6.7$ ); note 3-fold  $\Delta pK_a$  difference!<sup>43–45</sup> Based on time-resolved emission spectroscopy and steady state spectrofluorometry experiments, 1-naphthol displayed a rate of deprotonation ( $k_1^*$ ) greater than 2-naphthol by 280 times.<sup>44</sup> It was proposed that the stronger photoacidity of 1-naphthol was a consequence of populating and emitting from the  $^1\text{L}_a$  state, while 2-naphthol shows near degenerate  $^1\text{L}_a$  and  $^1\text{L}_b$  states, emitting from the  $^1\text{L}_b$  state.<sup>17,46,47</sup> We speculate that naphthols promoted to the  $^1\text{L}_a$  state deprotonate more readily because of pronounced antiaromatic character in the  $^1\text{L}_a$  state of naphthalene. Computed geometric indices of aromaticity for excited naphthalene show a more bond alternated  $^1\text{L}_a$  state and a more bond equalized  $^1\text{L}_b$  state (see data in the ESI†). See also resonance structures of naphthalene in the  $^1\text{L}_a$  ( $B_{2u}$ ) state (“diradical form,” note “antiaromatic” Clar sextet structure in the  $^1\text{L}_a$ ) and  $^1\text{L}_b$  ( $B_{3u}$ ) state (“allyl radical form”) (Scheme 2). We note that Baird’s original paper on the effects of triplet (anti)aromaticity also recognized a more antiaromatic “diradical form” for triplet benzene.<sup>1</sup>

Although substituents are typically considered to have negligible effects on aromaticity (unless charged or in the presence



**Scheme 2** (a) Resonance structures for the  $1L_a$  and  $1L_b$  states of excited state naphthalene, and (b) the "diradical" ( $B_{1u}$ ) and "allyl radical" ( $B_{2u}$ ) forms of triplet benzene.

of other push/pull substituents),<sup>48,49</sup> they can easily perturb the excited-state antiaromaticity of organic compounds,<sup>50</sup> having tremendous effects on reactions such as the excited-state proton transfer of photoacids. These findings are another manifestation of the increasingly recognized effects of excited-state (anti)aromaticity on the photochemical reactivity of organic compounds.

**Computational methods:** Geometry optimizations for the ground ( $S_0$ ) and excited ( $S_1$ ,  $1\pi\pi^*$ ) states of all acid and conjugate base structures were performed at CASSCF(12,12)/6-311+G(d,p) with  $C_s$  symmetry, employing Molpro2012.1.<sup>51</sup> The  $S_0$  and  $S_1$  geometries of 5,8-dicyano-2-naphthol and its conjugate base were computed at CASSCF(10,10)/6-311+G(d,p) with  $C_s$  symmetry. Computed dissected nucleus-independent chemical shifts, NICS(1)<sub>zz</sub>,<sup>38,39</sup> were performed at CASSCF(12,12)/6-31G(d,p), employing the Dalton2016 program,<sup>52</sup> to quantify the magnetic effects of aromaticity and antiaromaticity in the  $S_0$  and  $S_1$  states<sup>40</sup> of the acids and conjugate bases. NICS(1)<sub>zz</sub> values were computed at 1 Å above each of the six membered ring centers and include only contributions from the "out-of-plane" (zz) tensor component perpendicular to the ring plane.  $\Delta$ NICS(1)<sub>zz</sub> values were calculated based on the sum of ring NICS(1)<sub>zz</sub> values of the conjugate base minus that of the acid.

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## Conflicts of interest

There are no conflicts to declare.

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