Chem Soc Rev

REVIEW ARTICLE

View Article Online
View Journal | View Issue

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

Check for updates

Cite this: *Chem. Soc. Rev.*, 2018, **47**, 5786

Electrochemical strategies for C–H functionalization and C–N bond formation

Markus D. Kärkäs

Received 17th March 2018

DOI: 10.1039/c7cs00619e

rsc.li/chem-soc-rev

Conventional methods for carrying out carbon–hydrogen functionalization and carbon–nitrogen bond formation are typically conducted at elevated temperatures, and rely on expensive catalysts as well as the use of stoichiometric, and perhaps toxic, oxidants. In this regard, electrochemical synthesis has recently been recognized as a sustainable and scalable strategy for the construction of challenging carbon–carbon and carbon–heteroatom bonds. Here, electrosynthesis has proven to be an environmentally benign, highly effective and versatile platform for achieving a wide range of nonclassical bond disconnections *via* generation of radical intermediates under mild reaction conditions. This review provides an overview on the use of anodic electrochemical methods for expediting the development of carbon–hydrogen functionalization and carbon–nitrogen bond formation strategies. Emphasis is placed on methodology development and mechanistic insight and aims to provide inspiration for future synthetic applications in the field of electrosynthesis.

1. Introduction

Synthetic chemists are constantly faced with the challenge of producing molecules, whether it be small-molecule synthesis or accessing increasingly complex target structures, in a selective and efficient manner. The methods and strategies of organic synthesis are continuously being improved and augmented in order to expand the chemical toolbox, thereby enabling new routes to solve ever-increasingly complex problems in chemical research.[1–6] Over the years, the use of renewable chemical feedstocks,[7] biocatalysis[8] and base-metal catalysis,[9] in combination with the development of cascade/tandem processes[10] has become paramount for realizing atom-economic and more sustainable reaction platforms.[11] Recently, research in the chemical sciences has experienced renewed interest in visible light photocatalysis[12,13] and electrochemistry.[14–17] These versatile synthetic platforms have the ability to promote challenging bond constructions under mild reaction conditions.

Department of Chemistry, Organic Chemistry, KTH Royal Institute of Technology, SE-100 44 Stockholm, Sweden. E-mail: karkas@kth.se

Markus D. Kärkäs

Markus D. Kärkäs earned his PhD degree in 2013 from Stockholm University under the direction of Prof. Björn Åkermark. His research focused on the development and mechanistic insight of artificial water oxidation catalysts. He carried out postdoctoral research in the group of Prof. Corey Stephenson at the University of Michigan working on photochemical methods for valorization of lignin. His research interests include photoredox catalysis, organic electrochemistry and artificial photosynthesis.

1.1. Fundamental principles and general considerations of preparative electrochemistry

Electrolysis is carried out in an electrochemical cell, a reactor that is comprised of an electroactive species/substrate, electrolyte, solvent and (at least) two electrodes (an anode and a cathode). The anode is connected to the positive pole of a power source (galvanostat or potentiostat) and is oxidative while the cathode is connected to the negative pole and is reductive. For construction of the cell, two different options exist: undivided cells and divided cells (Fig. 1). In undivided cells the anode and cathode are not separated and are thus placed in the same compartment. This setup is preferred due to its ease of construction and allows for both reduction and oxidation to occur within the same compartment, thereby enabling the substrate to be exposed to all species present in the reaction. When employing this arrangement, it is crucial to consider the possible reactions that may occur at the auxiliary electrode.[18] However, examples exist where a species produced at the auxiliary electrode interferes with the electrochemical process at the working electrode. This issue can be ameliorated using a divided cell. In this arrangement, the two (anodic and cathodic) compartments are physically separated through the use of a small, porous frit. This allows for the transfer

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

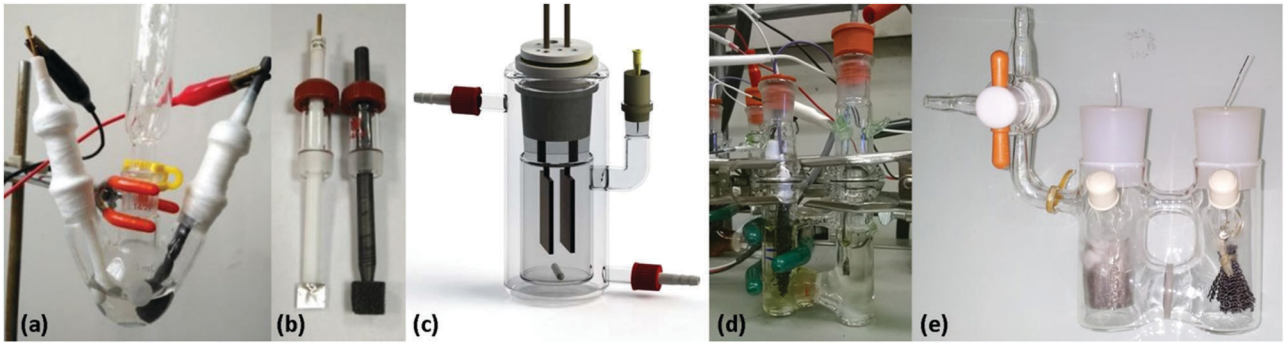


Fig. 1 Selected electrolysis setups. (a) Three-necked round-bottom flask equipped with (b) a reticulated vitreous carbon anode and a platinum plate cathode (Reprinted with permission from ref. 926. Copyright 2017 Wiley-VCH Verlag GmbH & Co. KGaA, Weinheim). (c) Undivided beaker-type glass cell equipped with an isostatic graphite anode and a platinum cathode (Reproduced from ref. 918 with permission from The Royal Society of Chemistry). (d) H-Type divided cell equipped with a reticulated vitreous carbon anode and a platinum wire cathode (Reprinted with permission from ref. 398. Copyright 2018 American Chemical Society). (e) H-Type divided cell equipped with a 4G glass filter, an anode made of carbon strings and a platinum plate cathode (Reprinted with permission from ref. 958. Copyright 2017 Wiley-VCH Verlag GmbH & Co. KGaA, Weinheim).

of charge and enables the two half-reactions of the electrolysis to occur separately.[19,20]

In principle, electrodes can be constructed from any inert material that enables electron transfer in solution. Examples of electrodes include carbon-based materials (graphite and glassy carbon), magnesium, platinum and stainless steel. Another appealing electrode material is reticulated vitreous carbon (RVC), a material that combines the properties of carbon with glass, which has a high surface area, high void volume, and chemical resistance. At the end of the reaction, the electrodes can simply be physically removed as long as they are stable to the applied reaction conditions.[21,22] Acceptable solvents that can be employed in electrochemical processes include, but are not limited to acetonitrile, dichloromethane, methanol, or dimethylformamide. Here, the use of nonconventional organic solvents, such as trifluoroethanol,[23] trifluoroacetic acid,[24] ionic liquids[25] and supercritical fluids,[26] has also received attention. In order for the reaction solution to become conductive, a supporting electrolyte has to be added. These are typically simple salts, such as alkali metal perchlorates and tetraalkyl ammonium salts, and are mainly used for increasing the conductivity of organic systems.

The electron transfer events that occur at the anode or cathode enable one to classify the electrochemical reactions as either oxidations or reductions, respectively. They occur on the surface of the electrodes and are therefore heterogeneous processes. The resulting species that is generated after the initial electron transfer between an electrode and an organic substrate is subsequently diffused into solution, which is followed by a secondary reaction of the initially produced radical species.[27–31] Here, the potential of the electrode (E)—that is the difference between the potential at the electrode of interest and the selected reference electrode, such as the saturated calomel electrode (SCE)—will determine whether a specific electron transfer process is thermodynamically feasible, and is given by eqn (1):

$$\Delta G = -nFE \tag{1}$$

where ΔG is the free energy change, F is Faraday's constant (96 485 Coulombs mol^{-1}), and n is the number of electrons involved in the overall reaction. An advantage of preparative electrolysis is that an initial electroanalytical evaluation using, for example, cyclic voltammetry (CV) enables convenient assessment of the reaction system. Such simple analyses provide a great deal of vital information with relatively limited experimental effort and insight into which functional groups that may be oxidized or reduced, as well as the magnitude of the free energy that is needed to promote electron transfer in the studied electrochemical process.[32]

It is important to note that the electrolysis experiments can be conducted using either a controlled-current or controlled-potential approach. In a controlled-current experiment, the current is held constant while the voltage gradually increases until the potential is reached for the electroactive species. Once depleted, the potential continues to rise until a second electroactive species or solvent molecule is oxidized or reduced. It is thus critical to consider the redox potentials of the substrate as well as all other species in solution in order to assure that the initial redox reaction does not involve participation of the solvent or other additives. However, if the potential needs to be tightly controlled, a controlled-potential experiment where the potential is set to a certain value will allow for a selective redox reaction to occur.[33] With this setup, it is important to have in mind that the decrease in substrate concentration is accompanied by a decrease in the current, and thus the rate of the reaction.

1.2. Historical advances in synthetic electrochemistry

Applying electrical current to affect a chemical reaction can be traced back to the 1830s when Faraday electrolyzed an acetate solution to produce hydrocarbons.[34] However, the Kolbe electrolysis constitutes one of the earliest carbon–carbon (C–C) bond forming reactions that was studied in detail.[35,36] Here, carboxylate anions are oxidized at an anode. The initial electron transfer triggers extrusion of CO_2 to generate carbon-centered free radicals, which subsequently undergo homocoupling to produce the new C–C bond. The reaction

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

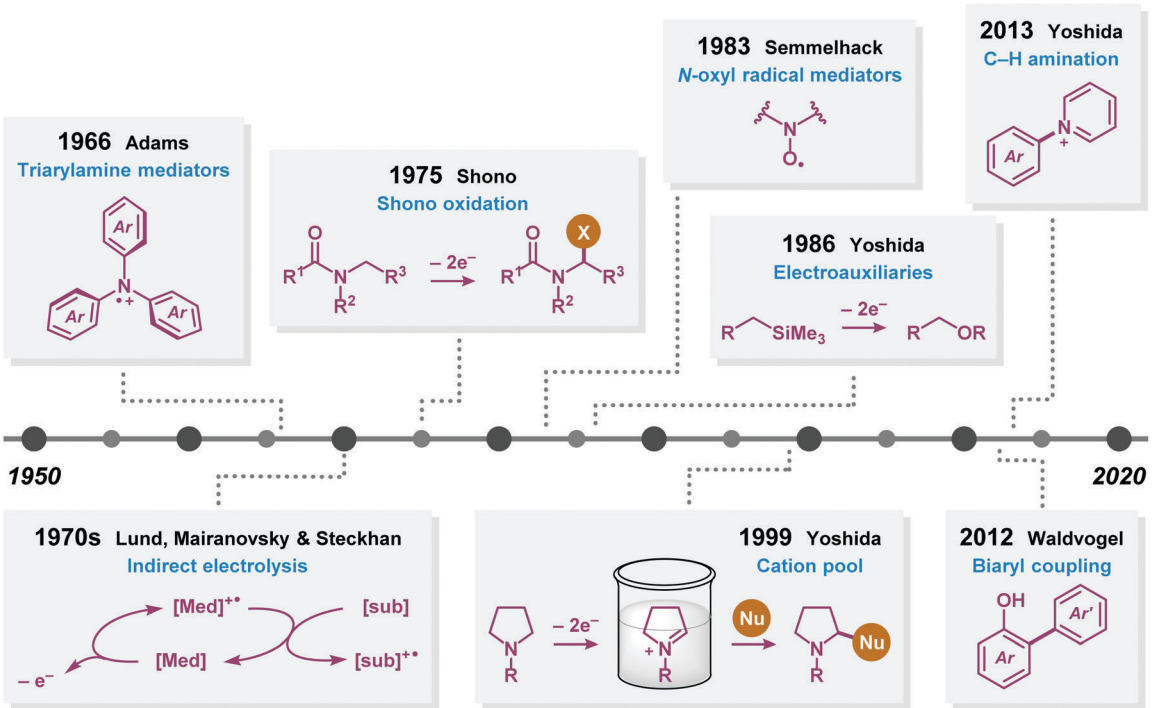


Fig. 2 Advances in organic electrochemistry.

still serves as a versatile method for C–C bond formation in contemporary organic synthesis.[37] Preparative electrolysis has also received significant interest from the chemical industry.[38–40] Classical examples include the chloralkali and the Hall–Héroult processes, as well as production of organic compounds, such as adiponitrile and ethylene glycol.[41]

Additional advances in preparative electrolysis include the use of indirect electrolysis in which a redox mediator undergoes electron transfer at an electrode to afford an electrochemically generated reagent that triggers the reaction of interest (Fig. 2).[42] This approach has a long history[43] and one of the earliest industrial applications concerned the oxidation of glucose to calcium gluconate using bromide as mediator.[44] During the last four decades, commonly employed organic redox mediators for anodic oxidations are triarylamines[45] and nitroxyl radicals.[46,47] Since the active species of the mediator is continuously regenerated at the electrode it enables the use of substoichiometric quantities. Advantages of redox-mediated electrolysis (indirect electrolysis) are that problems such as are that problems associated with heterogeneous electron transfer, such as overpotentials,[48] can be avoided and that electrolysis can be conducted at lower potentials than the redox potential of the substrate, accelerating the reaction rate while affording higher selectivities by circumventing potential side reactions. It should be noted that there exist several similarities between electrochemically-mediated reactions and visible light photoredox catalysis (Fig. 3). In this visible light-mediated platform a photocatalyst absorbs photonic energy. Upon absorption of visible light, a long-lived redox-active excited state is produced, which can engage in single-electron transfer (SET) events with, for example, organic substrates.

In the late 1980s, Yoshida introduced the concept of using electroauxiliaries.[49,50] Typical electroauxiliaries include enol ethers,[51] organostannanes,[52] silyl groups,[53] and thioacetals.[54] The electroauxiliary is functionality appended to a compound in order to aid the oxidation or reduction of the substrate by reducing its oxidation or reduction potential. This facilitates either oxidation or reduction of the starting material in the presence of the desired nucleophile and reduces the likelyhood of overoxidation or overreduction of the resultant product by promoting electron transfer in a more selective and predictable fashion. Moreover, noticeable contributions during the 21st century include Waldvogel's work on selective biaryl cross-coupling

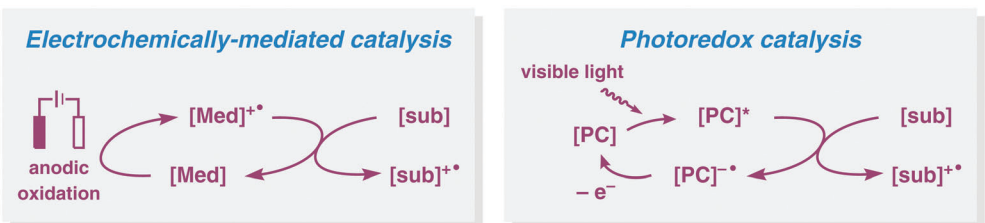


Fig. 3 Comparison of electrochemically-mediated catalysis and visible light photoredox catalysis. PC = photocatalyst.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

platforms[55] and Yoshida's strategy for C–H amination[56] of aromatic compounds using the "cation pool" method.[57,58]

1.3. Advantages of applying electrochemical methods in organic synthesis

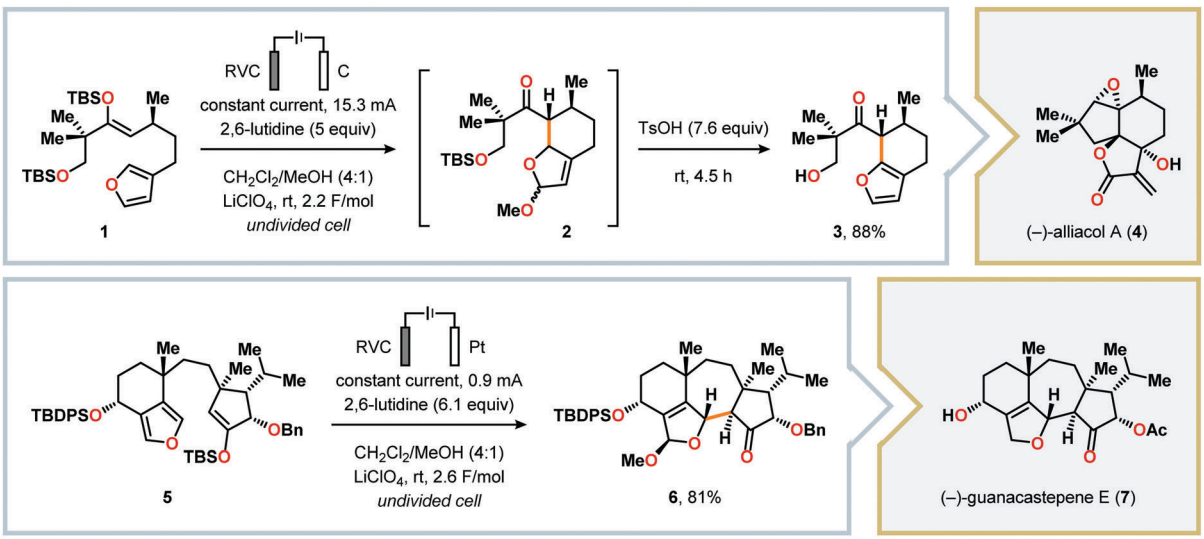
The use of electrochemistry is considered a "green" and sustainable alternative to traditional, toxic redox-based methods. It has the ability to maximize atom efficiency while replacing stoichiometric redox reagents, such as NaH, OsO_4 and $Pb(OAc)_4$, with an electrical current, thereby minimizing production of reagent waste. Moreover, since the energy of the electrochemical system can be governed by the applied electrode potential, a majority of the preparative electrochemical reactions, even the ones with high activation energies, can be carried out at ambient temperature. Therefore, the application of electrochemistry in organic synthesis constitutes an appealing strategy for generating reactive radical intermediates under mild conditions.[59,60]

Radical cyclization reactions, for example, have typically been conducted in the presence of stoichiometric quantities of hazardous and toxic reagents, such as tin hydrides and azobis(isobutyronitrile) (AIBN). However, employing electrochemistry allows the development of reagent-free protocols that operate under mild reaction conditions, providing an environmentally friendly alternative.[32,61] In addition to the aforementioned advantages of applying electrochemistry in organic synthesis, another feature of electrochemistry is that it can also be used to trigger umpolung reactions.[62–64] Two elegant examples of using such a strategy can be found in Moeller's synthesis of alliacol A (**4**)[65,66] and Trauner's synthesis of guanacastepene E (**7**)[67,68] in which silyl enol ethers are coupled with furans *via* intramolecular electrooxidative coupling (Scheme 1).

It is evident that organic electrochemistry represents an enabling technology for the organic chemistry research community for generating reactive but controllable radical intermediates under neutral reaction conditions. However, for the technique to become part of the standard chemical toolbox, the barrier to adoption needs to be eliminated. The necessity to construct your own home-made electrochemical apparatus has certainly contributed to the general lack of interest in pursuing electrochemistry, for example, as part of a total synthesis. The barrier to adoption becomes even higher if there exist alternative methods for accomplishing the desired transformation. Furthermore, the numerous reaction variables, such as cell type, experiment type, electrode material and electrolyte, might be overwhelming, especially for practicing organic chemists without any prior experience in electrochemistry. Organic electrochemistry would benefit from being presented in a more user-friendly manner. Access to a commercially available standardized device for organic electrochemistry would certainly mitigate this barrier to adoption and allow the field to develop more rapidly.[69]

1.4. Scope of this review

Over the years, a variety of review articles have been published that summarize the impressive advances made in the field of organic electrochemistry.[70] Pioneering reviews include those by Wawzonek[71] and Weinberg.[72] Anodic oxidation processes have been examined by Adams,[73] Eberson[27,74] and Shono,[75] and more recently by Boydston,[76] Chiba,[77] Moeller,[78–81] and Schäfer.[82,83] Reviews detailing both anodic oxidation and cathodic reduction processes have been summarized by Schäfer,[84,85] Wright[86] and Yoshida.[87] Progress in indirect electrolysis has been reviewed by Nikishin,[42c] Steckhan,[42a,b] as well as Francke and Little[42d] while the synthetic endeavors of using electrochemistry in complex settings have been described by Baran.[88] In this review an overview of the electrochemical strategies for carrying out carbon–hydrogen (C–H) functionalization as well as carbon–nitrogen (C–N) bond formation is presented with an emphasis on methods development and mechanistic insight.[89] The review is organized based on the aforementioned reaction types and are grouped according to the electrochemical methods employed. While focus is on representative examples from the past two decades, early pioneering work is also highlighted. The aim of this review is to encourage



Scheme 1 Synthesis of alliacol A and guanacastepene E *via* anodic coupling reactions.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

researchers to explore and to adopt organic electrochemistry, a technique with considerable potential, to the general synthetic toolbox.

2. Carbon–hydrogen (C–H) functionalization under oxidative electrochemical control

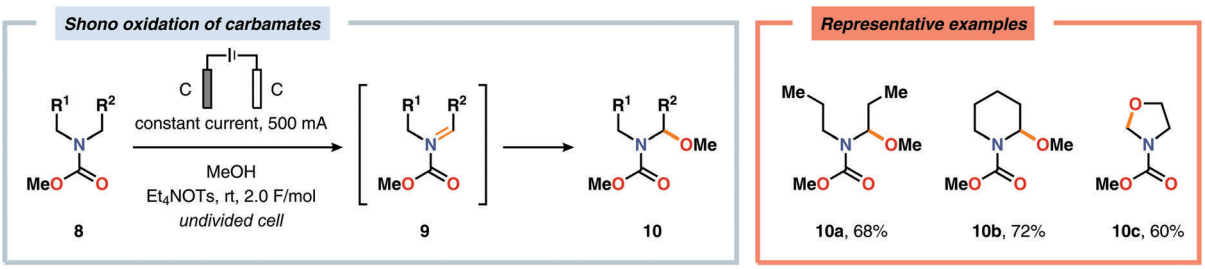
Over the past decade, transition metal-catalyzed C–H functionalization has enabled the development of a variety of nontraditional and innovative bond constructions in contemporary synthetic organic chemistry. The direct functionalization of C–H bonds represents a powerful strategy for selective C–C and carbon–heteroatom (C–X) bond formation, thereby improving atom- and step economy as well as streamlining chemical synthesis. However, despite intensive efforts and increased understanding of the mechanistic aspects of the C–H functionalization reaction manifolds, the inherent difficulty of activating kinetically inert C–H bonds has limited the development of catalytic platforms that operate efficiently under mild reaction conditions.[90,91] In this chapter, various electrochemical concepts to expedite the development of C–H functionalization platforms are therefore surveyed.

2.1. Shono-type anodic oxidations

Nitrogen-based heterocycles constitute an essential structural motif and are commonly found in natural products. In 1975, Shono and coworkers disclosed an electrochemical method for oxidation of carbamates to *N*-carbamoyl iminium ions.[92–95] The reaction proceeds through the initial formation of a nitrogen-centered radical (*vide infra*), which is subsequently oxidized to an iminium ion intermediate that can be trapped with an alcoholic solvent molecule.[96] This allows for functionalization of the α-position adjacent to the nitrogen atom in heterocycles.[97–110] For example, carrying out the anodic oxidation in methanol gives access to α-methoxylated products **10** (Scheme 2), which are established precursors to *N*-acyl iminium ions. If the substrate contains an alcohol moiety, intramolecular trapping produces oxazoline derivatives, such as **10c**.[92] Upon treatment with Brønsted or Lewis acids, the generated *N*,*O*-aminals can be converted back to the iminium ion species, which allows for trapping with allyl silanes,[111–113] cyanide,[114–116] fluoride,[117] furans,[118] isocyanides,[119] silyl enol ethers,[120] and trialkyl phosphites.[121] Alternatively, functionalization can be carried out to yield a diversity of scaffolds as shown in Scheme 3.[122] Attempts have also been focused on developing asymmetric versions of the Shono oxidation using chiral cyclic dipeptides,[123–126] oxazolines,[127] oxazolidinones[128] and pyrrolidones[129–131] as well as ketopinic acid[132] and carbamate-,[133–136] phosphorus-[137,138] and sulfur-based[139] chiral auxiliaries to give poor to high stereoselectivities in the alkylation step (Scheme 4).

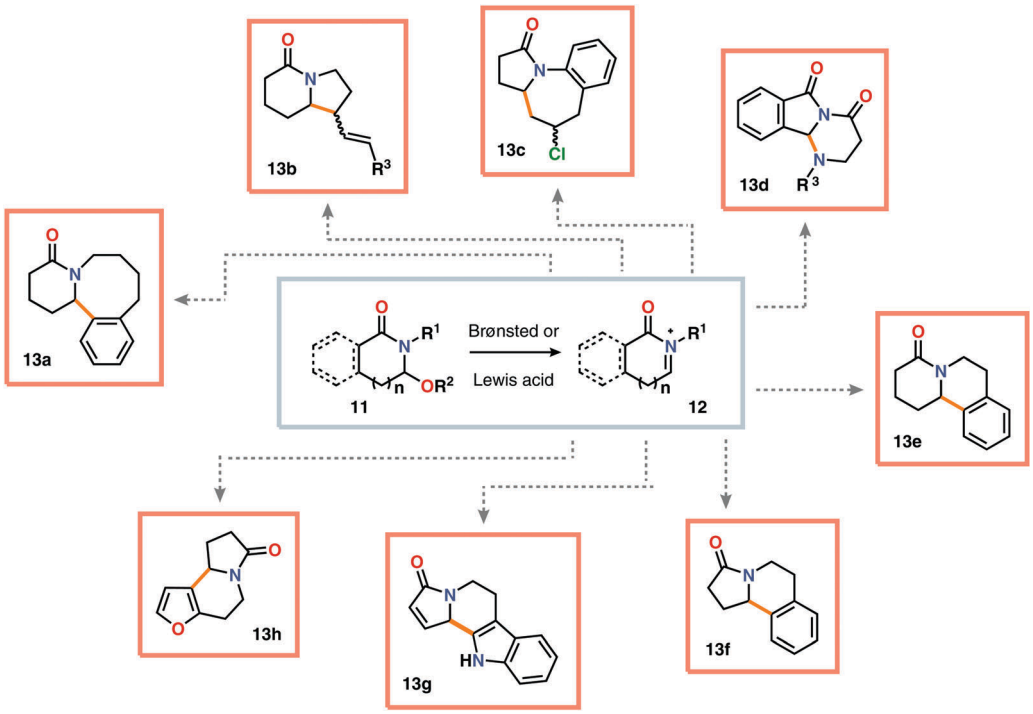
Shono-type anodic oxidations have in general been carried out with substrates and nucleophiles, such as methanol and cyanide, which possess high oxidation potentials. Although amines have lower oxidation potentials, which should facilitate the electrochemical oxidation process, protocols utilizing amines have been employed less frequently compared to their amide and carbamate counterparts. Huang and Gong recently disclosed a method for the electrochemical oxidation of γ-lactams. Direct trapping of the generated electrophilic acyliminium ions with anilines could be accomplished despite their low oxidation potential (Scheme 5).[140] Key to achieving high yields of the substituted γ-lactam products was to carry out the oxidation in a "quasi-divided cell"[141,142] with a Pt wire anode and a Pt foil cathode in which the large difference in surface areas of the working and counter electrodes provided high current density at the anode, thereby favoring oxidation of the lactam. The protocol was applicable to a wide range of anilines containing electron-withdrawing groups, such as CF_3, CO_2Et, and OCF_3, as well as electron-donating groups, such as Me and OMe. Halogen-containing anilines were also effective substrates for the developed reaction, which provides an opportunity for further synthetic elaborations.[140] Fuchigami and Baba have also demonstrated anodic oxidation of imidates and imines to yield α-methoxylated or α-acetoxylated products (Scheme 6).[143]

Of the limited protocols that exist that rely on using amines as substrates,[144,145] Gallardo and Vilà reported that hindered secondary alkyl diamines **28** could be produced from anodic oxidation of hindered secondary alkyl amines **27** (Scheme 7).[146–148] They subsequently demonstrated an electrosynthetic procedure to afford imidazolium and tetrahydropyrimidinium derivatives **30** from secondary alkyl diamines **29** (Scheme 8).[149] The Matsumura group has also shown that the introduction of a nucleophile to the α-position of an L-prolinol derivative (**31**) upon Shono-type oxidation is viable (Scheme 9).[150] Furthermore, Hurvois and coworkers prepared an α-amino nitrile through anodic oxidation in a synthesis of (+)-myrtine.[151] Shono-type oxidation of aliphatic amines can also be extended to the synthesis of sulfonyl amidines.[152] Different tertiary amines (**36**) were initially evaluated and were

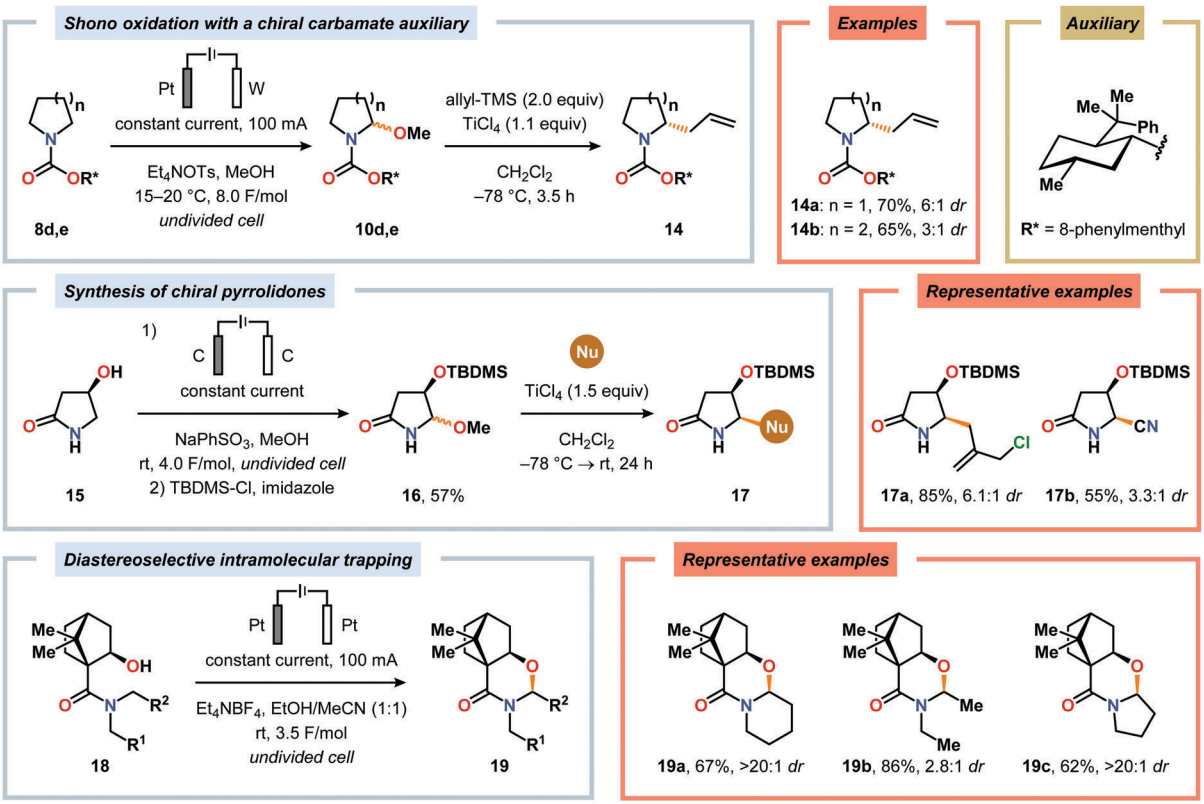


Scheme 2 Shono oxidation of carbamates.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



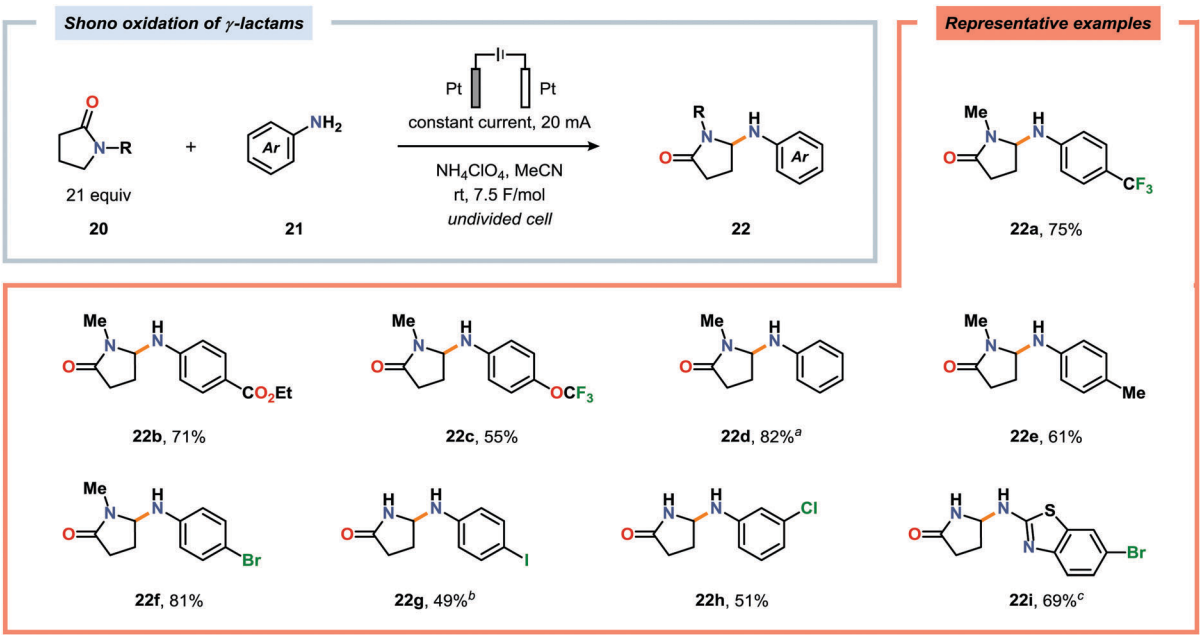
Scheme 3 Scaffold diversity from *N*-acyl iminium ions.



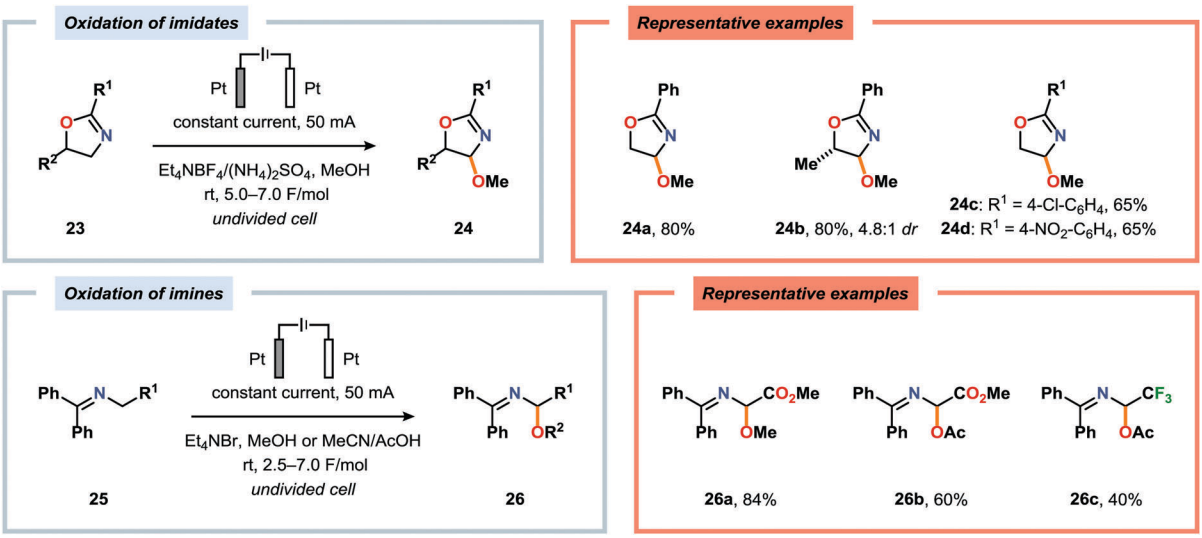
Scheme 4 Selected strategies for stereoselective functionalization of Shono oxidation products.

shown to efficiently afford the sulfonyl amidine derivatives **38** upon reaction with sulfonyl azides **37** (Scheme 10). Employing Et_3N as the amine these transformations presumably include anodic oxidation of the aliphatic amine to produce an iminium intermediate (**39**), which tautomerizes into the corresponding enamine (**40**). Subsequent 1,3-dipolar cycloaddition between the

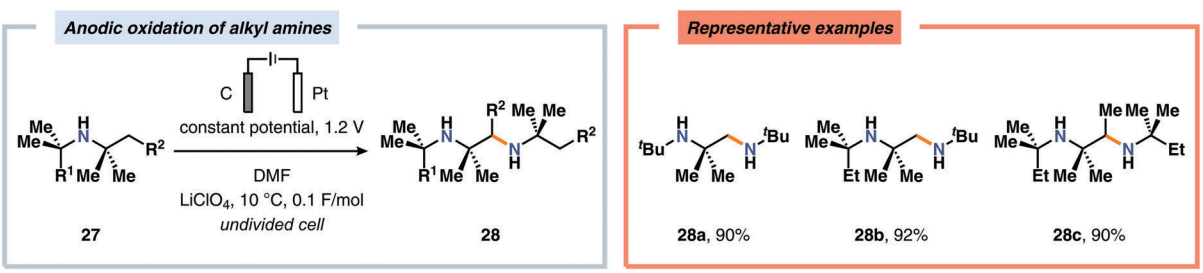
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 5 Shono oxidation of γ-lactams using anilines as the trapping nucleophiles. [a] Carried out in MeCN/EtOAc (3 : 2), Et_3N (1.7 equiv.), constant current (10 mA) and 10.5 h. [b] After 1.5 h. [c] After 5 h.

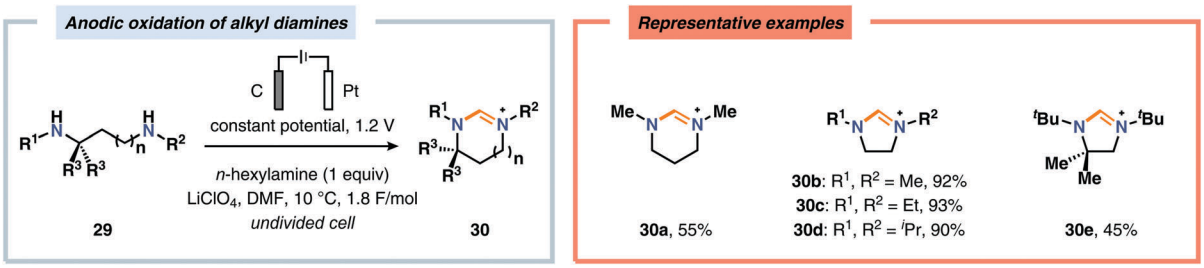


Scheme 6 Anodic oxidation of imidates and imines.

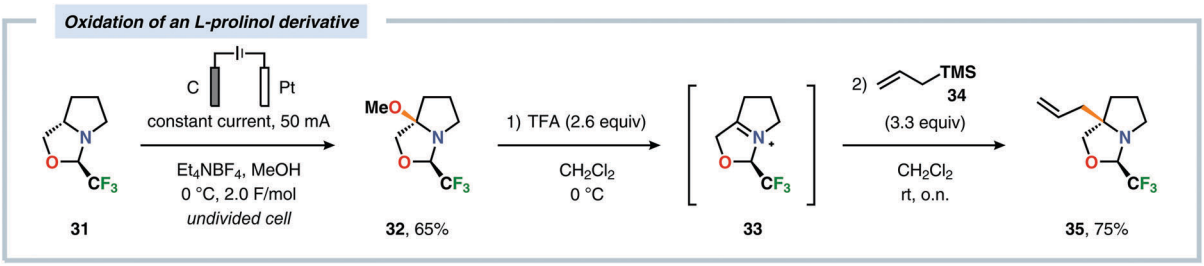


Scheme 7 Shono-type anodic oxidation of hindered secondary alkyl amines.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 8 Shono-type oxidation of secondary alkyl diamines.

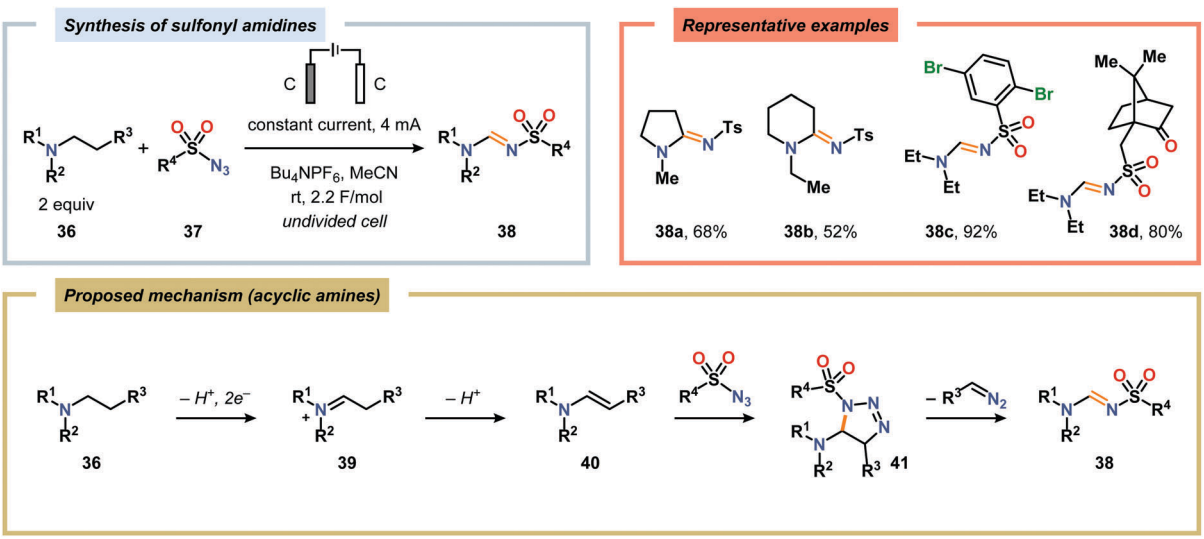


Scheme 9 Electrochemical oxidation of an L-prolinol derivative.

generated enamine and tosyl azide furnishes cycloadduct **41**. Finally, extrusion of CH_2N_2 delivers the desired amidine product **38**. The synthesis of amidine derivatives was also accomplished using secondary and primary aliphatic amines.

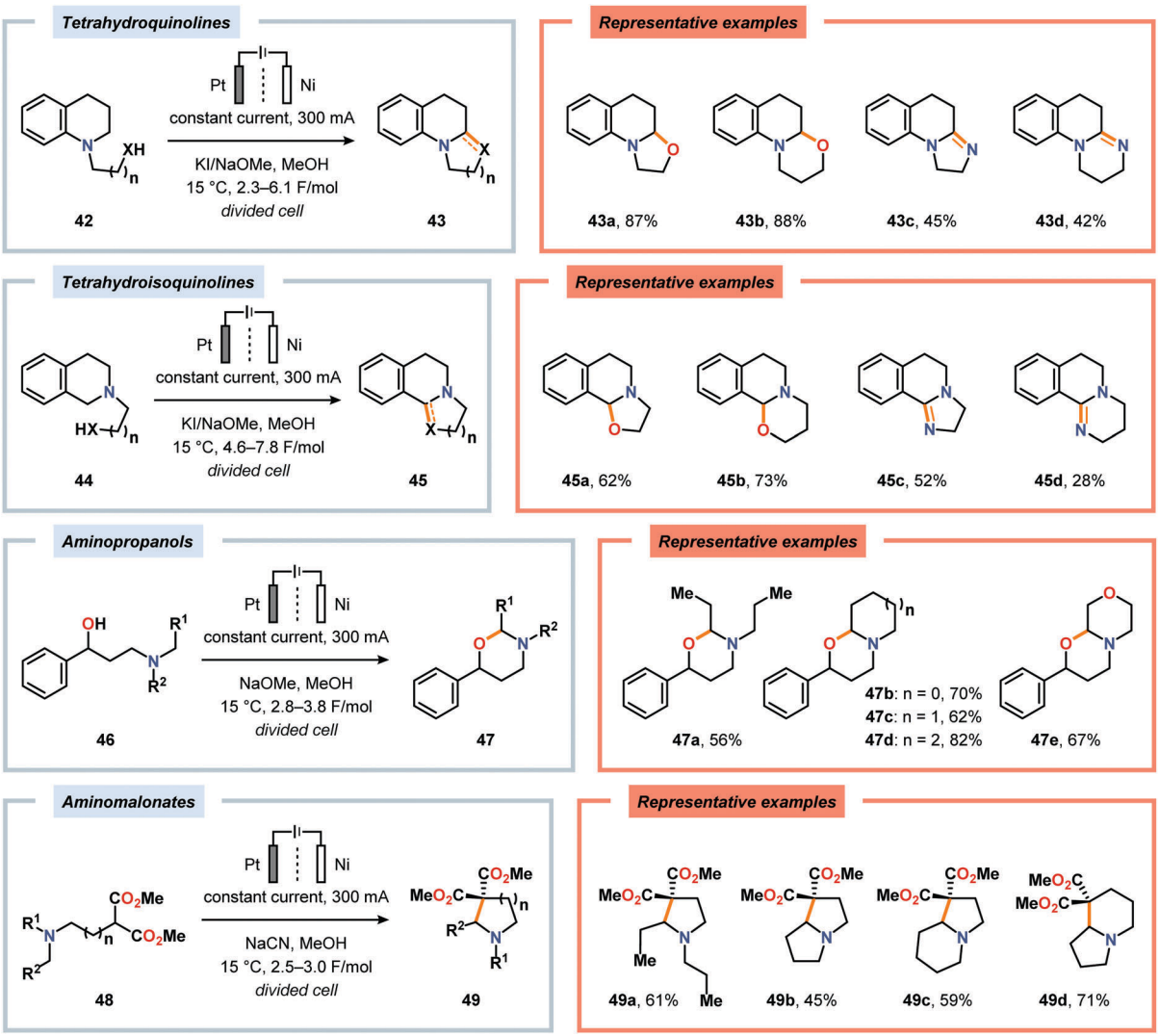
Okimoto and coworkers have also demonstrated that Shono-type oxidation of various amine-containing motifs carrying pendant nucleophiles can be employed to access a diverse set of heterocyclic compounds through intramolecular C–C or C–X bond formation processes (Scheme 11).[153–159] Using a combination of NaOMe and KI as supporting electrolyte in MeOH proved to be the optimal for affecting electrochemical oxidation and subsequent intramolecular cyclization of tetrahydro(iso)-quinolines **42** and **44** housing pendant alcohol or amine groups. For the tetrahydroisoquinoline-derived substrates **44a** and **44b**, selective oxidation at the benzylic position afforded **45a** and **45b**.[157] The electrochemical oxidation/cyclization strategy could also be applied on 3-dialkylamino-1-phenylpropanols **46** to give the 1,3-oxazinane derivatives **47** in moderate to high yields.[158] Furthermore, the authors also investigated the electrooxidation of dimethyl aminomalonates **48**. Here, the use of NaCN as the supporting electrolyte resulted in the highest yields for the cyclized products **49**.[157,160]

The Luo laboratory recently reported a visible light-mediated protocol for the enantioselective coupling of tetrahydroisoquinolines and simple alkyl ketones to furnish *syn*-Mannich-type adducts with good to excellent diastereo- and enantioselectivities.



Scheme 10 Electrochemical synthesis of sulfonyl amidines from aliphatic amines and sulfonyl azides.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



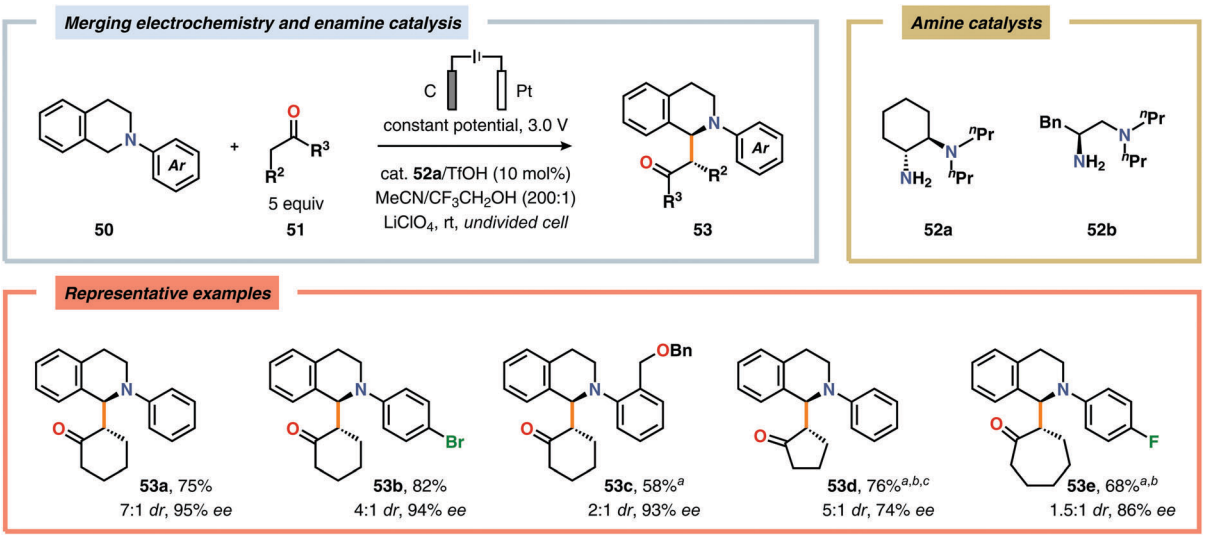
Scheme 11 Electrooxidative cyclizations of amine-containing motifs.

However, the process required the use of substoichiometric amounts of 3-nitrobenzoic acid as sacrificial oxidant.[161] In order to improve the utility and eliminate the use of a chemical oxidant, Luo and coworkers developed a protocol based on enamine catalysis and electrochemical C–H oxidation. The authors identified the chiral primary amine **52a** as the optimal catalyst and trifluoroethanol (CF_3CH_2OH) as the ideal additive. The established oxidant-free coupling reaction tolerated an array of *N*-aryl substituted tetrahydroisoquinolines and various simple ketones to provide the asymmetric oxidative coupling products **53** in good yields with good to excellent diastereo- and enantioselectivities (Scheme 12).[162] *N*-Aryl substituted tetrahydroisoquinolines are also susceptible to anodic nitro-Mannich C–C bond formation using nitromethane as the nucleophile.[163] Furthermore, Shono-type oxidation of tetrahydroisoquinolines has also been used for the oxidative functionalization of benzylic C–H bonds, mediated by a dual bromide ion/2,2,6,6-tetramethylpiperidinyl-*N*-oxyl (TEMPO) redox catalyst system, to produce dihydro-isoquinolinones.[164,165]

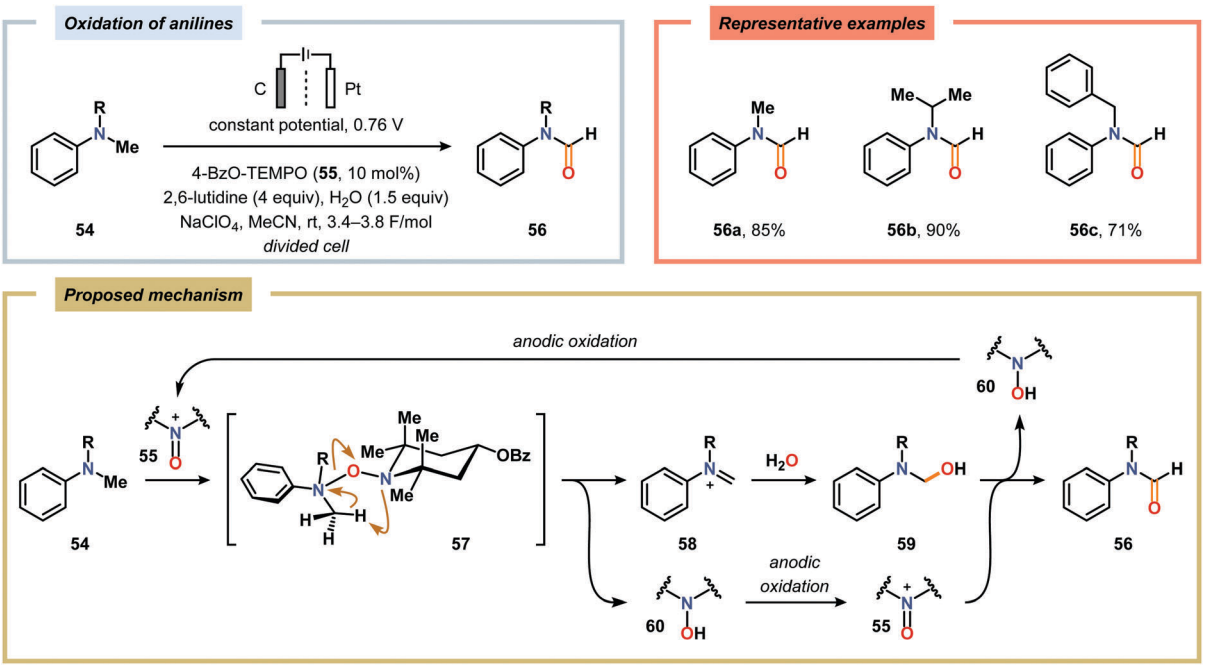
Kashiwagi and Anzai explored the electrochemical oxidation of *N*-alkyl-*N*-methylanilines (**54**) in aqueous media using a nitroxyl-based mediator. *N*-Alkylformanilides (**56**) were the major products and were obtained in 71–90% yield. The mechanism presumably proceeds through an iminium ion intermediate (**58**) which is hydrolyzed to produce hemiaminal **59**. This species undergoes a second oxidation event, mediated by the electrochemically generated oxoammonium species **55**, to furnish the corresponding formanilide **56** (Scheme 13).[166]

A potential strategy to expand the scope of the nucleophiles is to employ the "cation pool" method.[57] The "cation pool" method involves generation and accumulation of cations, such as *N*-acyliminium ions, through electrolysis at low temperatures. The nucleophile is subsequently introduced to the reaction mixture under non-oxidative conditions, which allows for easily oxidized nucleophiles to be used. Early work from Yoshida and coworkers involving the generation of "cation pools" from carbamates by low-temperature electrolysis enabled the use of a collection of carbon nucleophiles, including allyl silanes, enol

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 12 Oxidative coupling of tertiary amines with simple ketones through the merger of electrochemical C–H oxidation and enamine catalysis. [a] Hexafluoroisopropanol (HFIP) as additive. [b] **52b** as amine catalyst. [c] Cell potential of 2.0 V.

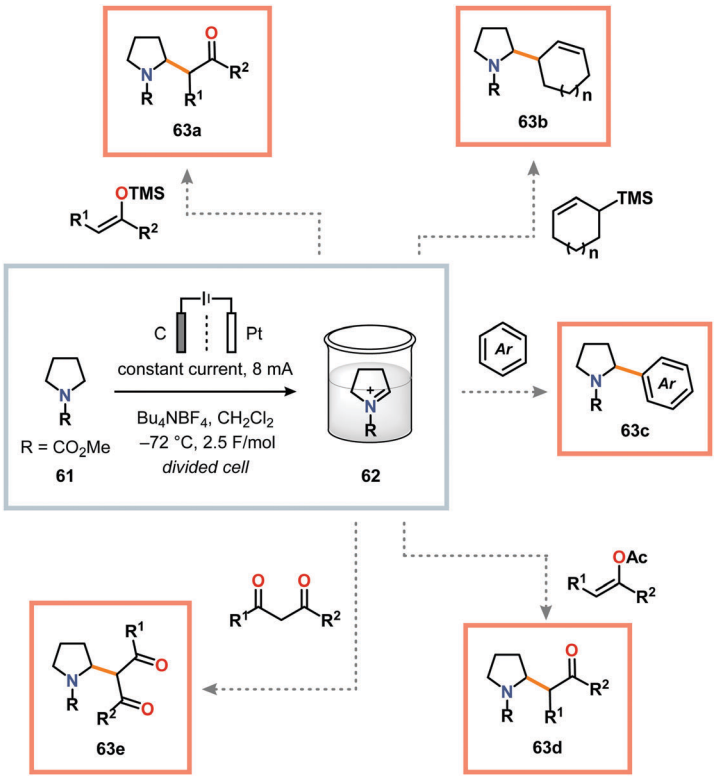


Scheme 13 Electrochemical oxidation of *N*-alkyl-*N*-methylanilines using a nitroxyl mediator.

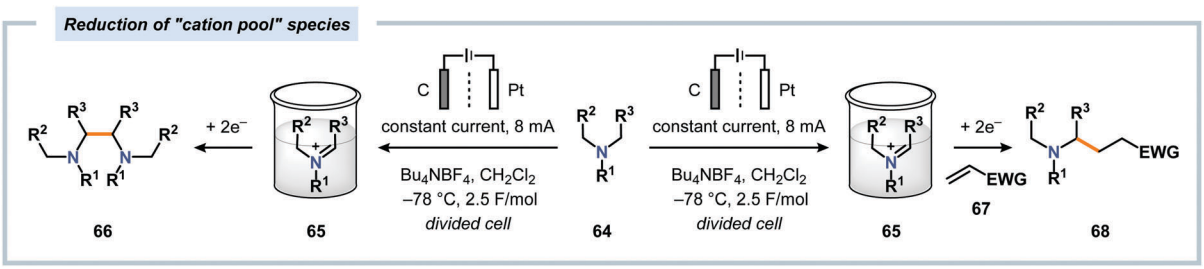
silyl ethers, enol acetates as well as aromatic and 1,3-dicarbonyl compounds (Scheme 14).[167] Other compatible nucleophiles consist of allyl stannanes, benzyl silanes, Grignard reagents and organoaluminum compounds.[168–170] These oxidative C–C bond forming reactions can also be carried out in a flow system, thus enabling continuous sequential combinatorial synthesis.[171–175] The Yoshida group has also highlighted that electrochemical reduction of *N*-acyliminium ions, generated by the "cation pool" method, affords α-amidyl radicals. These radicals can be harnessed to produce homocoupled products or be intercepted with electron-deficient olefins in Giese-type[176,177] reactions (Scheme 15).[178,179] The latter transformation serves as a formal addition of C–H bonds to alkenes and offers new redox-based opportunities to manipulate free radical species.

Additional strategies that allow for direct trapping of electrochemically generated *N*-acyliminium ions include the use of parallel laminar flow in micro-flow reactors,[180,181] "site isolation"[182,183] and acoustic emulsification.[184–188] The Atobe group has illustrated that the direct allylation of carbamates can be accomplished through the use of micro-flow reactors in which the flow is stable and laminar, thereby enabling a stable liquid–liquid interface and mass transfer between the two streams occurs only through diffusion. This concept affords selective anodic oxidation of substrates without affecting the oxidation of nucleophile and provides effective trapping of

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



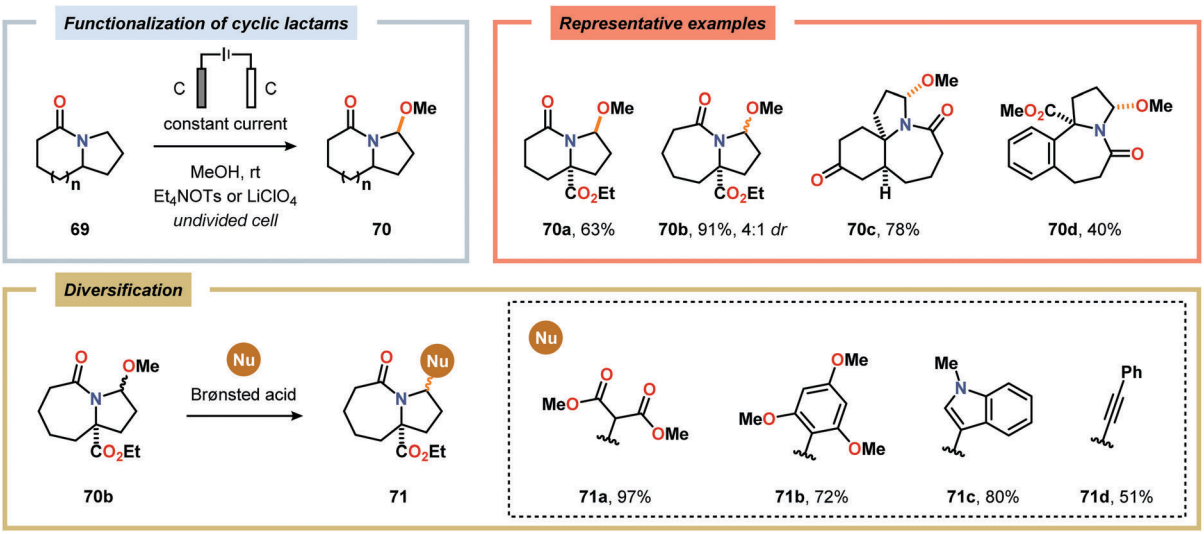
Scheme 14 Carbon–carbon bond formation of carbamates using the "cation pool" method.



Scheme 15 Electrochemical reduction of "cation pool" generated acyliminium ions.

potentially unstable cationic intermediates.[180,181] Atobe and co-workers have also established the feasibility of using acoustic emulsification in Shono oxidations. Here, chemoselective anodic oxidation of the substrate is realized by selecting an electrolytic medium in which the nucleophile is insoluble in. Upon ultrasonication, the nucleophile is dispersed as submicrometer range droplets, which react with the electrogenerated carbocations when the two species are in close proximity.[186,187] The Tajima laboratory has demonstrated the direct oxidative cyanation based on the concept of "site isolation". Using a polystyrene-supported quaternary ammonium cyanide (PS-$Me_3N^+CN^-$) as the heterogeneous cyanating reagent enabled the direct cyanation of anodically produced *N*-acyliminium ions, generated *via* Shono oxidation of carbamates. This highlights that the site isolation between an anode and, for example, the cyanating reagent (PS-$Me_3N^+CN^-$) dramatically suppresses the undesired oxidation of CN^- and allows for anodic cyanation of organic compounds, even ones that have higher oxidation potentials compared to CN^-.[182] Furthermore, utilizing recyclable solid-supported bases for *in situ* generation of electrolytes enables the development of systems where protic organic solvents serve as both the solvent and the supporting electrolyte.[189–196] This allows for easy separation of the solid-supported base through filtration and provides, for example, the methoxylated Shono-type product, which can be subjected to the subsequent C–C bond forming reaction without having to replace the solvent.[197,198] A complementary strategy for expanding the scope of Shono-type oxidations consists of using electroauxiliaries.[50] This concept relies on the inherent properties of the electroauxiliaries to lower the oxidation potential of the substrate, thereby facilitating and providing enhanced control of the anodic oxidation of the substrate. The electroauxiliaries employed in Shono-type oxidations have typically been silyl-based,[199–213] although 2,4,6-trimethoxyphenyl[214] and sulfur-based[215–217] electroauxiliaries have also been employed.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 16 Electrochemical modification and diversification of cyclic lactams.

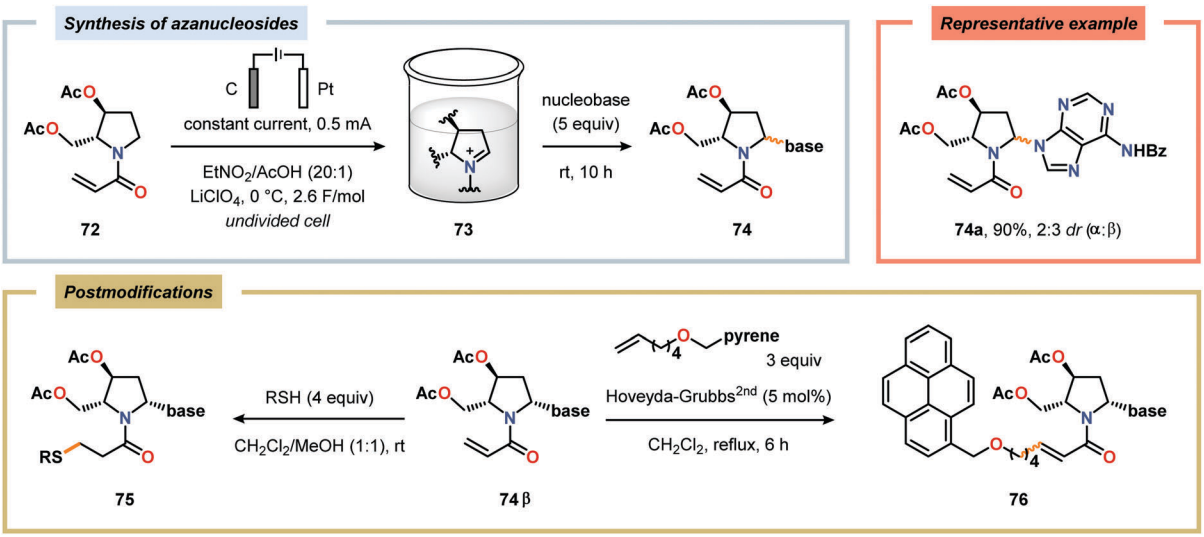
Considering the versatility of Shono-type oxidation reactions, it is perhaps unsurprising that these reactions have found numerous synthetic applications. The groups of Moeller and Aubé developed a practical two-step protocol for the late-stage functionalization of lactams by combining an intramolecular Schmidt reaction of keto azides followed by electrochemical anodic oxidation. The applicability of the Shono oxidation was exemplified across a range of ring systems. Finally, the versatility of the produced methoxyhemiaminals (**70**) was illustrated by synthesizing an array of functionalized lactam products (**71**) through the addition of various nucleophilic partners to the subsequent *in situ* generated *N*-acyliminium ions (Scheme 16).[218] Azanucleosides and related analogues have been found to exhibit anticancer and antiviral activity. Furthermore, incorporation of azanucleoside motifs into oligonucleotide sequences results in increased tolerance towards degradation by nucleases.[219] In this regard, the Matsumura group has exploited the Shono oxidation for preparation of azasugars,[220] and Chiba and coworkers recently applied the Shono oxidation in the development of a synthetic method for accessing azanucleosides. Initial studies revealed that various nucleophiles and nucleobases could be introduced at the α-position upon the anodic modification of *N*-carbamate protected prolinol derivatives.[221,222] Subsequent work showcased that *N*-acryloyl proline derivatives **72** could also be converted into the corresponding azanucleosides **74** by use of the Shono reaction. Moreover, the presence of the *N*-acryloyl group allowed for substrate postsynthetic modification through thiol conjugate addition or olefin cross-metathesis with a fluorescent dye (Scheme 17).[223] Electrochemical oxidation has also proven to be a viable strategy towards the synthesis of γ-amino acids[224] and preparing peptidomimetics.[225–235] Schmalz and coworkers utilized the Shono oxidation in the context of accessing a tricyclic dipeptide mimetic (**80**) starting from L-proline (**77**). The anodic oxidation of L-proline *tert*-butyl ester **78** afforded gram scale quantities of the desired methoxylated product **79** in virtually quantitative yield (Scheme 18). Building block **79** was subsequently converted into diproline mimetic **80** through peptide coupling, ring-closing metathesis and protecting group adjustment.[236]

The Shono oxidation has been exploited as a general synthetic method for accessing β-substituted indoles (Scheme 19). The products derived from the Shono oxidation served as masked aminoaldehydes, which upon treatment with arylhydrazines and $ZnCl_2$ afforded the corresponding β-substituted indoles through a Fischer indole synthesis pathway.[237] Ley and coworkers identified flow electrochemistry as an enabling technology for the Shono oxidation of various *N*-protected cyclic amines. Here, the Shono oxidation furnished the corresponding cyclic α-methoxyamines in excellent yields. Applying a Pictet–Spengler reaction between the electrosynthesized α-methoxyamines and substituted tryptamines afforded the biologically active indole alkaloid nazlinine (**88c**) and numerous related unnatural congeners (Scheme 20).[238] Other alkaloids that have been prepared include (−)-crispine A[239] and (±)-pumiliotoxin C.[240] Here, anodic cyanation furnished the desired α-amino nitriles, which upon further manipulations furnished the target alkaloids. The Santos laboratory has also demonstrated the utility of the Shono oxidation in total syntheses of ropivacaine[241] (**93**, Scheme 21), (−)-quinolactacin B[242] (**96**, Scheme 22) as well as (+)-lennoxamine[243] (**99**, Scheme 23). Finally, the Kam group reported that electrochemical oxidation could be applied to aspidofractinine-type alkaloids for construction of kopsidines A and B (Scheme 24), thus highlighting the potential of carrying out late-stage functionalization of complex molecules.[244] Additional synthetic applications of Shono-type oxidations include the synthesis of an angiotensin-converting enzyme inhibitor,[245] preparation of methoxylated analogues of the anticancer drugs ifosfamide and cyclophosphamide,[246] as well as dealkylation,[247–250] deallylation and debenzylation[251] of amines and amides.

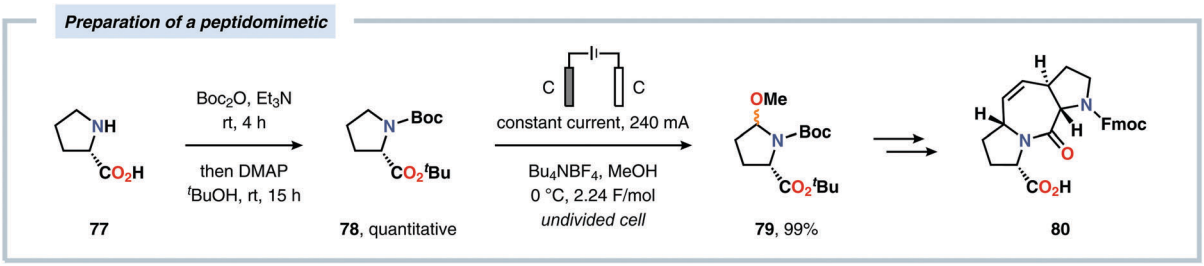
2.2. Anodic halogenation

Incorporation of fluorine into organic compounds has found broad applications in medicinal,[252–254] agrochemical[255,256] and material sciences.[257] Over the past decade, significant advances

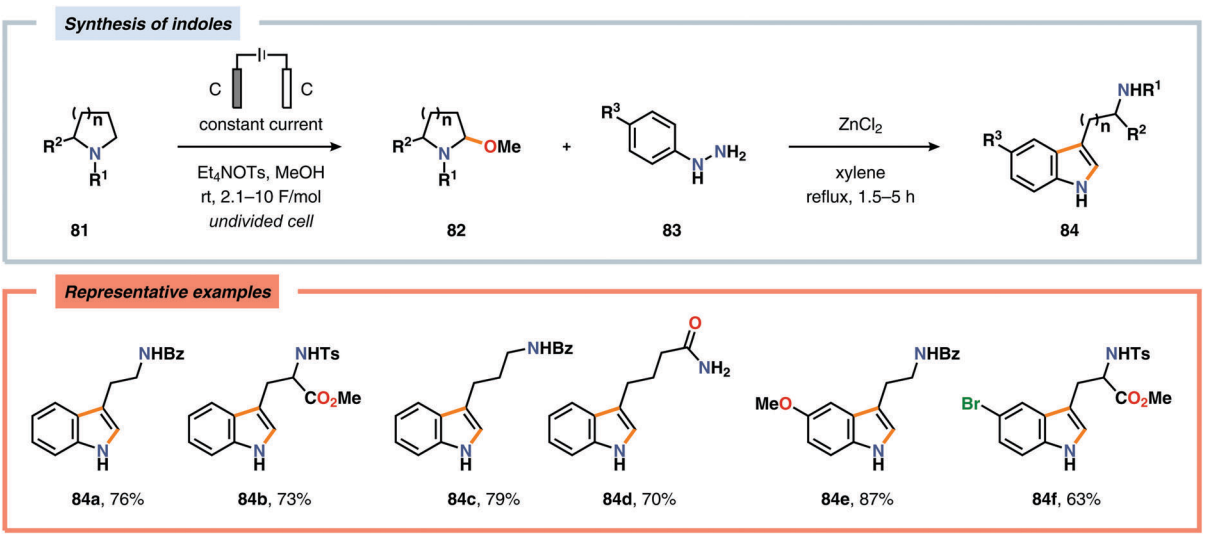
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 17 Electrochemical synthesis and postmodification of *N*-acryloyl azanucleosides.



Scheme 18 Application of the Shono oxidation in the preparation of a peptidomimetic.

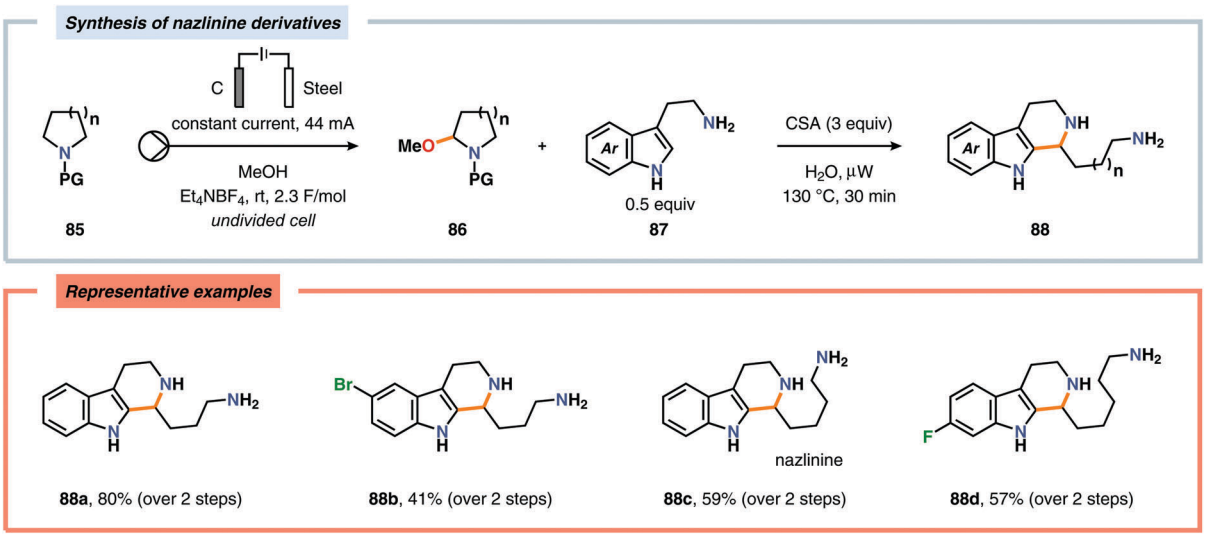


Scheme 19 Synthesis of β-substituted indoles *via* the Shono oxidation.

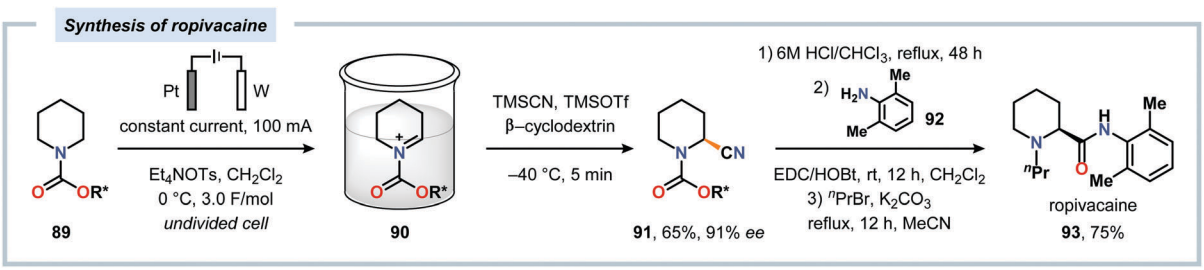
in chemo- and stereoselective fluorination of molecules with high functional and structural complexity have been made. In general, three different strategies exist for synthesis of organofluorine compounds: electrophilic,[258] nucleophilic[259] and radical[260] fluorination.[261,262]

Seminal studies by Rozhkov and coworkers in 1970 highlighted that selective electrochemical fluorination of arenes could be achieved in MeCN using $Et_4NF{\cdot}3HF$ as the ionic liquid, albeit in low yields. These reactions were proposed to proceed *via* arene radical cations, generated at the anode, that

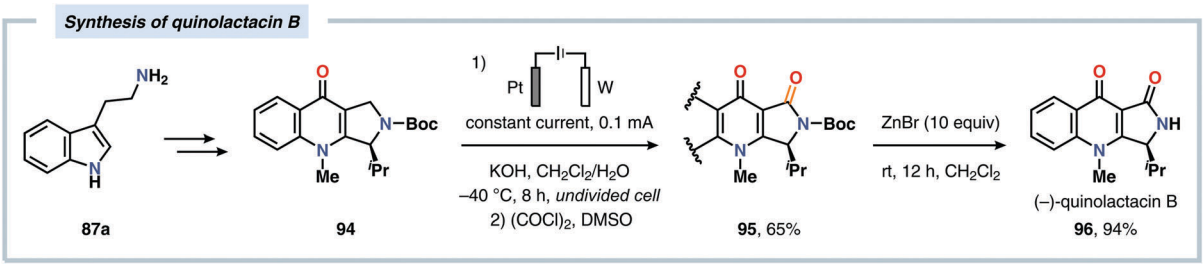
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



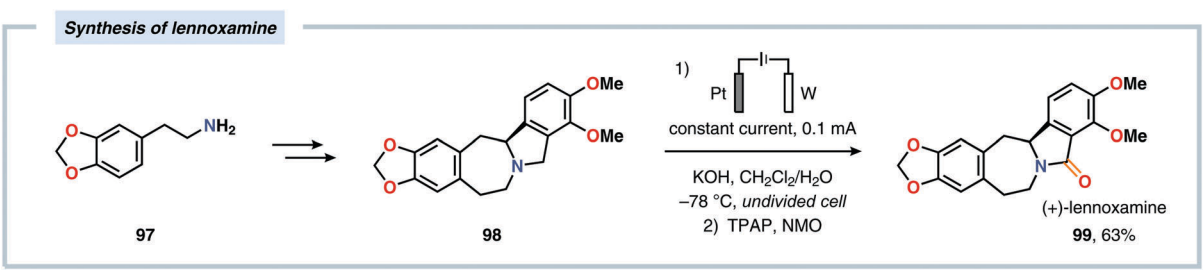
Scheme 20 Synthesis of nazlinine and unnatural congeners *via* a two-step method.



Scheme 21 Synthesis of ropivacaine employing the "cation pool" strategy. R* = 8-phenylmenthyl.

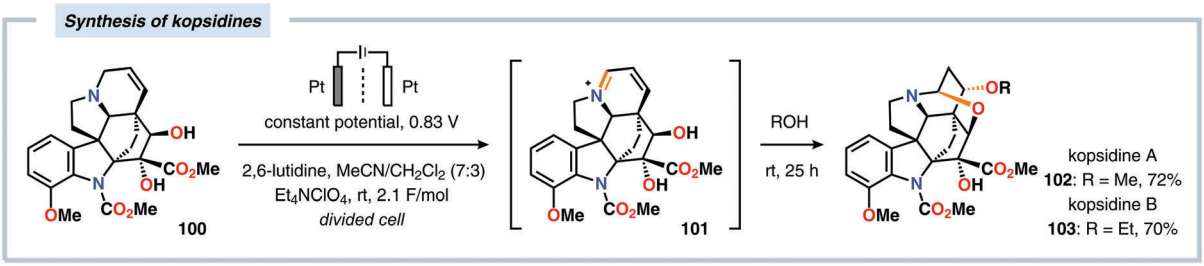


Scheme 22 Application of a late-stage Shono oxidation in the synthesis of (–)-quinolactacin B.



Scheme 23 Application of a late-stage Shono oxidation in the synthesis of (+)-lennoxamine.

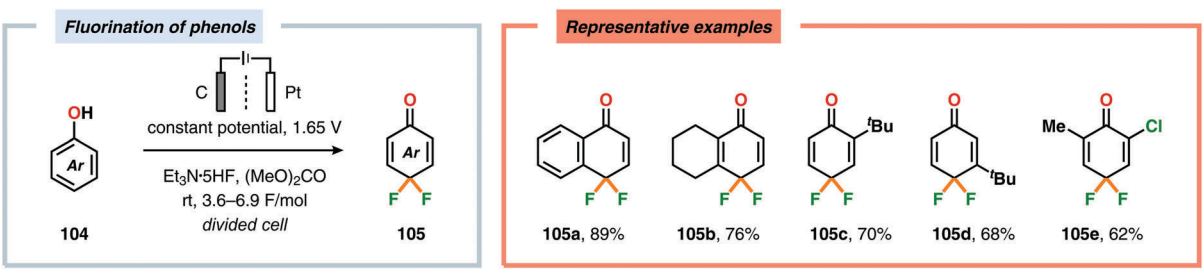
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 24 Shono-type oxidation of aspidofractinine-type alkaloids.

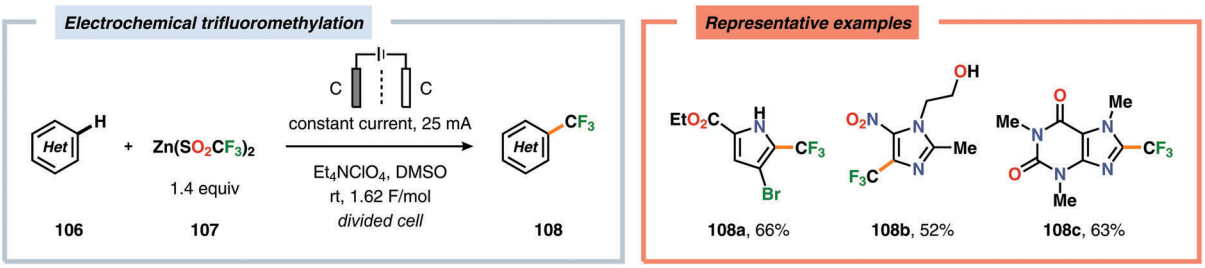
subsequently were trapped by fluoride to produce the carbon–fluorine (C–F) bond.[263–265] More recently, attempts to achieve selective fluorination of simple aromatic compounds include the work by Matsuda and coworkers. However, the yields of the fluorobenzenes were typically low.[266–269] Unlike unsubstituted arenes, such as benzene and naphthalene, electrochemical fluorination of toluene and related derivatives remains a challenge, as mixtures of products are formed originating from aromatic substitution, side chain substitution and addition reactions.[270–275] Heteroaromatics[276–283] and nitrogen-containing heterocycles[284–286] have also been reported to react in a similar manner. Anodic oxidation of phenols using $Et_3N{\cdot}5HF$ as the electrolyte was examined by the groups of Hara and Yoneda, and selectively afforded the corresponding 4,4-difluorocyclohexadienone derivatives in good yields (Scheme 25).[287–291] Furthermore, trifluoromethanesulfinate ($CF_3SO_2^-$) has recently been exploited as a source for electrochemical generation of trifluoromethyl radicals ($CF_3^{\bullet}$) in the presence of electron-rich aromatics[292] and heterocycles.[293] In this regard, the Baran and Blackmond groups demonstrated that electrochemical oxidation of zinc trifluoromethanesulfinate $\{Zn(SO_2CF_3)_2\}$ in DMSO in the presence of heterocycles, such as pyrazoles, pyrroles and benzothiazoles, afforded the corresponding trifluoromethylated products (Scheme 26). Moreover, the authors successfully highlighted that the developed electrochemical protocol was able to provide the functionalized heterocyclic products more effectively than conventional peroxide radical-initiated protocols for a majority of the screened substrates.[293]

Electrochemical fluorination of arylsulfides constitutes another transformation that has been extensively studied by several research groups. Early work during the 1990s from the Fuchigami[294–302] (Scheme 27) and Laurent[303,304] (Scheme 28) laboratories demonstrated that the introduction of an electron-withdrawing group at the α-position to sulfur had a dramatic influence on the reactions. These α-fluorination transformations were believed to proceed through a Pummerer-type mechanism where the electron-withdrawing group facilitates deprotonation of the α-carbon to produce intermediate **110**, thereby promoting fluorination.[305] The same effect could also be achieved using aromatics containing strongly electron-withdrawing groups, such as *p*-CN, *p*-NO_2 and *p*-SO_2Ph, as demonstrated by Simonet and coworkers.[306,307] These seminal studies constitute the first examples of selective electrochemical fluorination of chalcogen compounds. Since then, a variety of distinct sulfide-based motifs, such as 2*H*-benzo[*b*][1,4]-thiazin-3(4*H*)-one (**114j**),[308] benzothiazole (**114o**),[309–311] 2,3-dihydrothiochroman-4-one (**114k**),[312] 1,3-dithiolan-4-one (**114c**),[313] lactam (**114a**),[308] 1,3,4-oxadiazole (**114l**),[311,314] 1,3-oxathiolan-5-one (**114b**),[315] pyrimidin-4(3*H*)-one (**114h**),[316] phosphonate (**114d**),[317,318] pyridine (**114e**),[310,319] pyrimidine (**114g**),[310,311,319,320] quinazolin-4(3*H*)-one (**114i**),[319] quinoline (**114f**),[310] 1,3,4-thiadiazole (**114m**),[311,314] and 1,2,4-triazole (**114n**),[314] have been shown to be amenable to regioselective monofluorination (Scheme 29). Because of their good conductivity, non-flammability, non-volatility and thermal stability, ionic liquids, such as $Et_3N{\cdot}nHF$, have found widespread use as solvents in electrochemical fluorination reactions.[321–326] Recently, Fuchigami and coworkers demonstrated miscellaneous approaches to improve the electrochemical fluorination. One approach makes use of ultrasonication to promote the mass transport of the substrate due to the high viscosity associated with the ionic liquids. This was shown to increase the current efficiencies, selectivities and yields in the anodic fluorination of a variety of organosulfur compounds.[327] The authors have also developed an electrochemical system based on the cation-exchange reaction between alkali-metal fluorides and solid-supported acids. In the presence of 2,6-lutidine, the exchange reaction results in the *in situ* generation of 2,6-lutidine·HF, which functions as the fluorinating agent and supporting electrolyte, thus

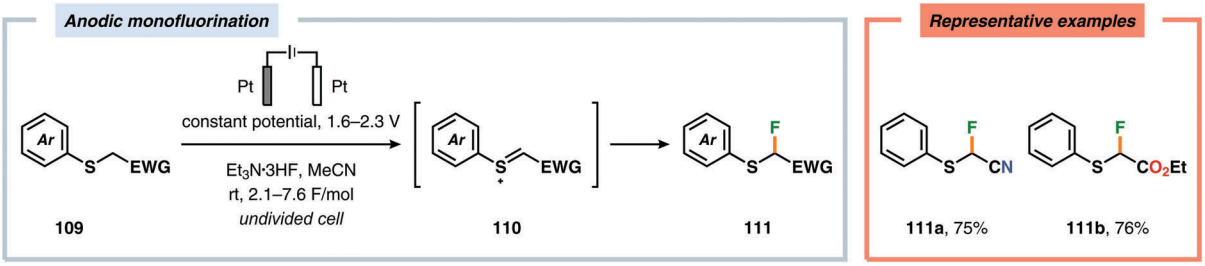


Scheme 25 Synthesis of difluorocyclohexadienones through electrochemical oxidation of phenols.

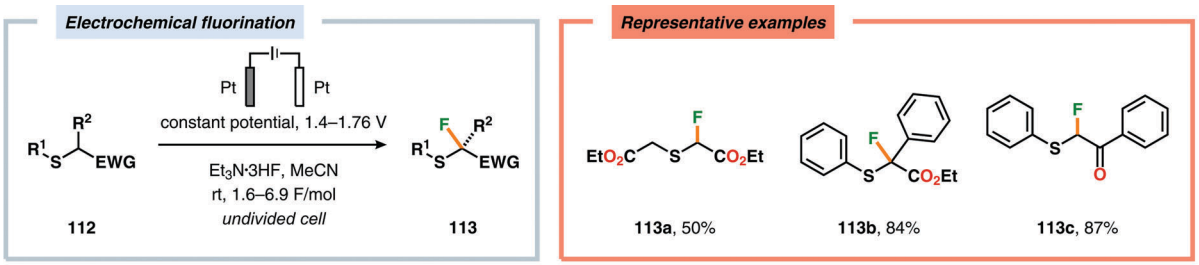
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 26 Electrochemical trifluoromethylation of heterocycles.



Scheme 27 Anodic monofluorination of organosulfur compounds.



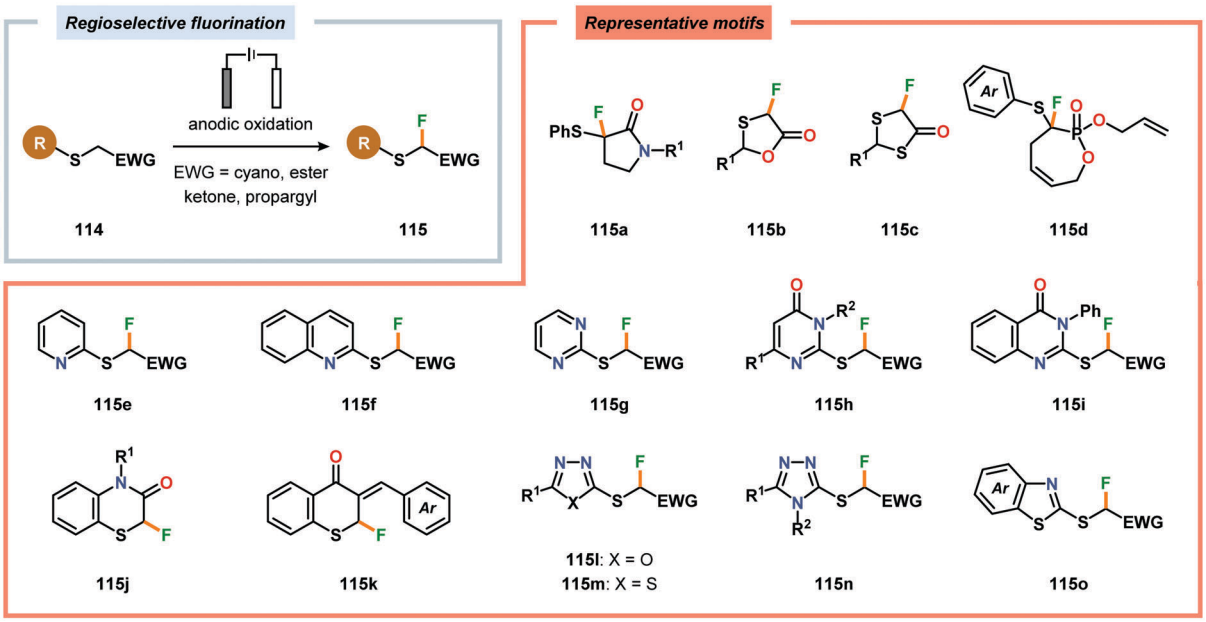
Scheme 28 Electrochemical fluorination of organosulfur compounds.

obviating the need to handle hazardous HF.[328] The Fuchigami group has also shown that indirect anodic fluorination can be realized using a hypervalent iodoarene difluoride ($ArIF_2$) mediator, which can be anodically produced from the corresponding iodoarene species and a HF-based ionic liquid ($Et_3N{\cdot}3HF$). Here, indirect reaction with the hypervalent iodoarene mediator proceeded efficiently to furnish the desired monofluorinated product, thus limiting anode passivation (the formation of a nonconducting polymer film on the electrode surface that suppresses faradaic current), a drawback that is often encountered when carrying out direct electrochemical fluorination.[329] Attempts to develop diastereoselective fluorination protocols have also been reported using chiral auxiliaries derived from camphorsulfonic acid,[330] (*R*)-(−)-3-chloro-1,2-propanediol,[331] L-cysteine[332] and menthol,[333] providing the monofluorinated products in modest to good diastereoselectivities.

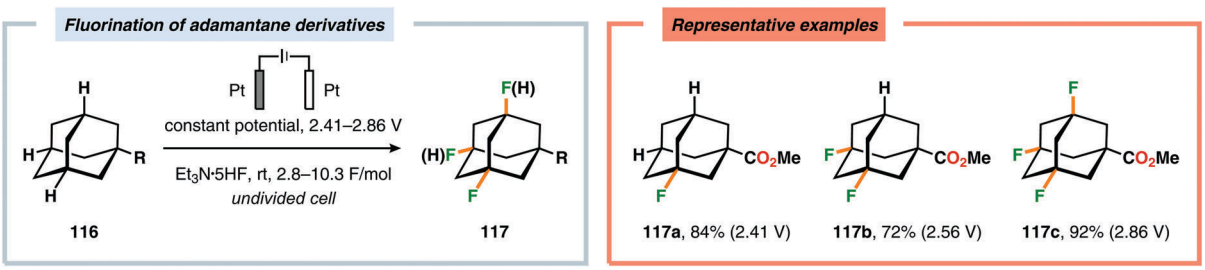
Recently, electrochemical fluorination of adamantane derivatives has been studied by Hara and coworkers. By controlling the oxidation potential, the authors were able to guide the fluorination process to selectively yield the mono-, di-, tri- or tetrafluorinated adamantane derivatives (Scheme 30).[334,335] Moreover, regioselective electrochemical monofluorination has also been reported for a collection of oxygen-containing heterocycles, including carbonates,[336,337] chroman-4-ones,[338–340] esters,[337] ethers[336,337] and lactones.[336,337] Additionally, electrochemical protocols for accomplishing fluorodesulfurization,[341–349] difluorination,[350,351] fluoro-selenylation,[352] iodofluorination[353] and thiofluorination[354] of electron-deficient olefins as well as partial fluorination of conjugated dienes[355] have been devised.[356]

In contrast to the electrochemical fluorination strategies, the anodic oxidation protocols involving chlorides, bromides and iodides typically proceed *via* generation of the corresponding electrophilic molecular counterpart (X_2) due to the relatively low oxidation potentials of these halogen ions.[357–367] Due to the limited solubility of alkali metal halide salts in organic solvents, a biphasic system consisting of an aqueous phase and a halogenated organic solvent is commonly employed to affect chlorination of aromatics[368,369] and heterocycles[370] as well as bromination of toluene derivatives.[371] The electrodes are placed in the upper layer of the aqueous phase, thus enabling selective oxidation of the halide. The electrochemically produced X_2 or hypohalite subsequently reacts with the organic substrate at the aqueous/organic

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



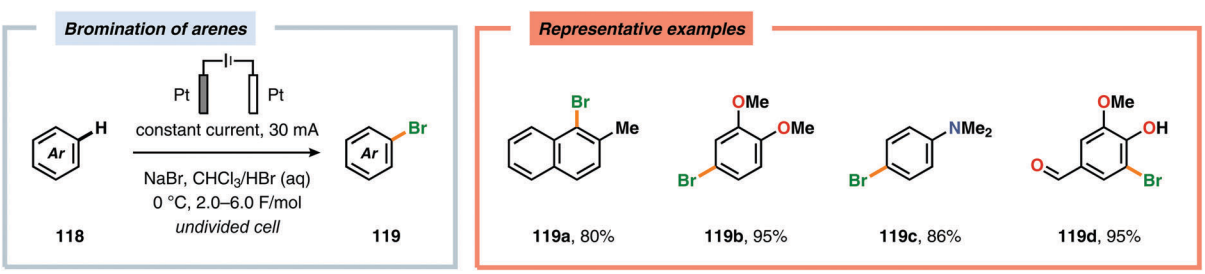
Scheme 29 Electrochemical regioselective fluorination of organosulfur compounds.



Scheme 30 Selective electrochemical fluorination of adamantane derivatives.

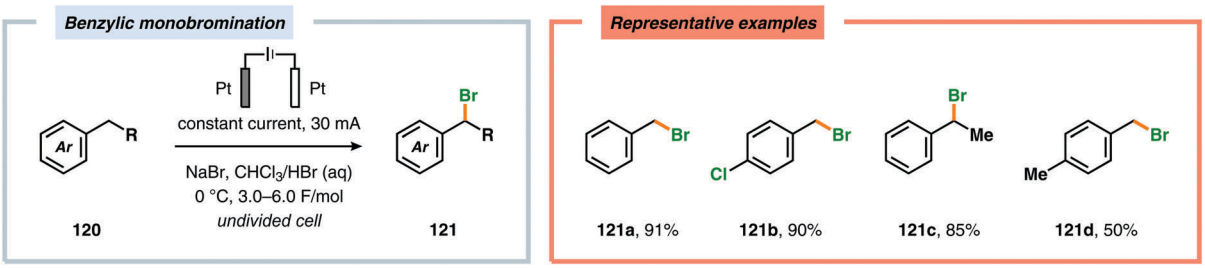
interface to ultimately produce the desired halogenated product. However, this can to some extent be circumvented using poly-(ethylene glycol) as an additive, which enhances the solubility of the inorganic salts in aprotic organic solvents.[372] By employing a biphasic system, bromination of various arenes[373] (Scheme 31) as well as benzylic mono-[374] (Scheme 32) and dibromination[375] (Scheme 33) of alkyl aromatics has been achieved. In addition to these examples, electrochemical chlorination of 1,3-dicarbonyl compounds using Cu catalysis,[376] azulene derivatives,[377,378] olefins,[379–384] phenols[385] and pyrazolecarboxylic acids,[386] as well as bromination of heterocycles[387] and olefins[388–391] have been reported.

By employing a similar electrochemical strategy as used for the chlorination of pyrazolecarboxylic acids,[386] Petrosyan and coworkers developed an electrochemical protocol for iodination of pyrazole and derivatives thereof. The iodination was carried out in a biphasic system consisting of water and chloroform using KI as the iodine source. As expected, the presence of electron-donating groups was found to facilitate the electrochemical iodination.[392,393] The Yoshida group has elegantly

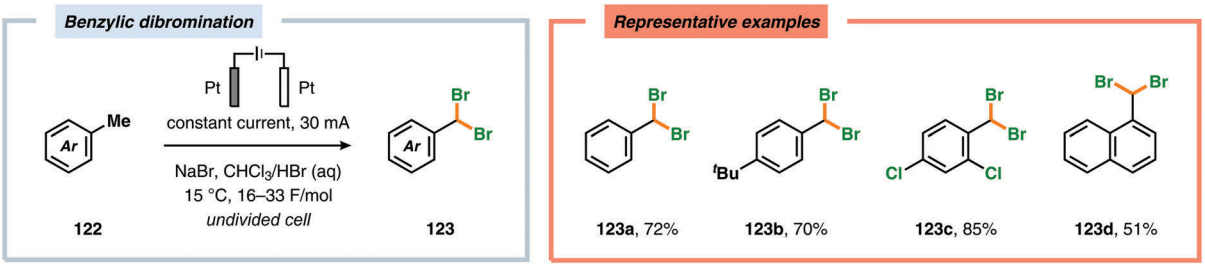


Scheme 31 Electrochemical bromination of arenes.

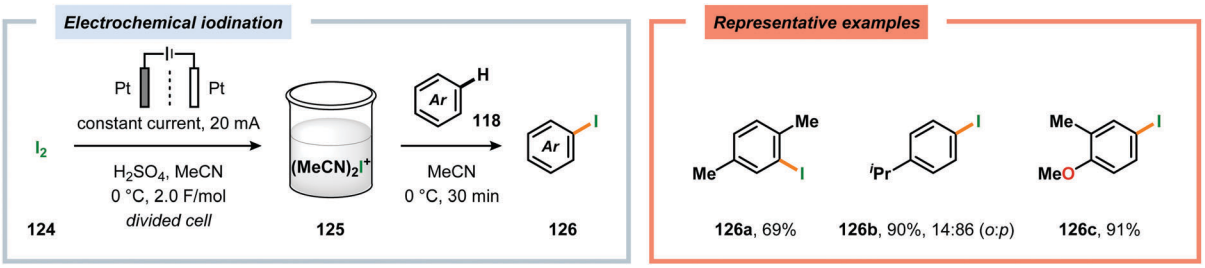
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 32 Benzylic monobromination of alkyl aromatics.



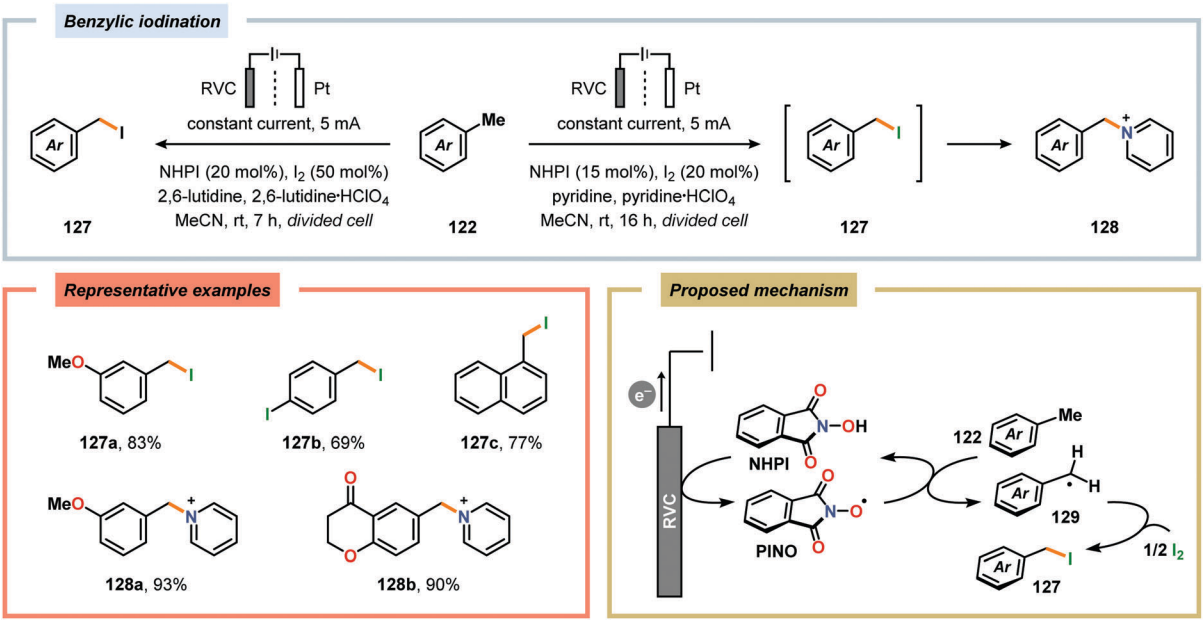
Scheme 33 Benzylic dibromination of alkyl aromatics.



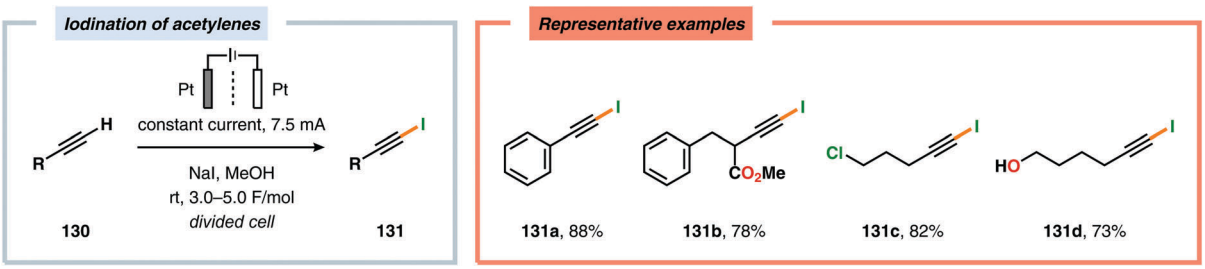
Scheme 34 Electrochemical iodination of aromatic compounds.

demonstrated iodination of aromatic compounds using electroaccumulated I^+ (Scheme 34). Initially, a "cation pool" approach was employed in which an MeCN solution containing iodine (I_2) was electrolyzed for generation of the electrophilic iodination agent, which was proposed to involve $(MeCN)_2I^+$. Subsequently, the anodic solution was added to a solution containing the arene. Alternatively, running the reactions in flow was shown to both increase the selectivity for the mono- *versus* the diiodinated compound and the yields compared to a batch reactor. Furthermore, the flow rate had a dramatic influence on the product selectivity where lower flow rates dramatically decreased the selectivity of the monoiodo compound due to less efficient mixing.[394,395] In addition, Hilt and coworkers recently revealed that electrochemical synthesis of aryl iodides can be achieved through anodic iododesilylation.[396] Anodic oxidation of arenes containing alkyl side chains provides aryl radical cations, which can subsequently be further oxidized to afford benzylic cations upon release of a proton, thus presenting a strategy for carrying out benzylic functionalization.[397] Recently, the Stahl group disclosed an alternative approach to benzylic C–H functionalization (Scheme 35).[398,399] The developed concept consisted of employing *N*-hydroxyphthalimide (NHPI) as a mediator.[400–402] Here, NHPI undergoes proton-coupled electron transfer at the anode to give phthalimido-*N*-oxyl (PINO), which subsequently mediates hydrogen atom transfer (HAT) from weak C–H bonds.[403] Using toluene derivatives and in the presence of a base, such as 2,6-lutidine, Stahl and coworkers were able to generate benzylic radicals at relatively modest oxidation potentials. I_2 was shown to be an effective radical trapping agent, thereby providing the corresponding benzyl iodide. The authors had to use a divided cell configuration in order to avoid reduction of the trap at the cathode. The developed benzylic C–H iodination method tolerated several functional groups, including esters, ethers, halides and ketones. *In situ* nucleophilic substitution using pyridine/pyridinium as the electrolyte proved to lead to efficient displacement of iodide by pyridine, leading to the corresponding benzylpyridinium product under electrochemical control. Iodide is produced in the nucleophilic displacement step, which can be reoxidized to iodine at the anode, allowing these reactions to be performed with substoichiometric amounts of iodine (20 mol%). Finally, using the *in situ* functionalization protocol enabled the authors to prepare three vital pharmaceutical building blocks,

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 35 Electrochemical iodination of toluene derivatives using NHPI as mediator.



Scheme 36 Electrochemical iodination of terminal acetylenes.

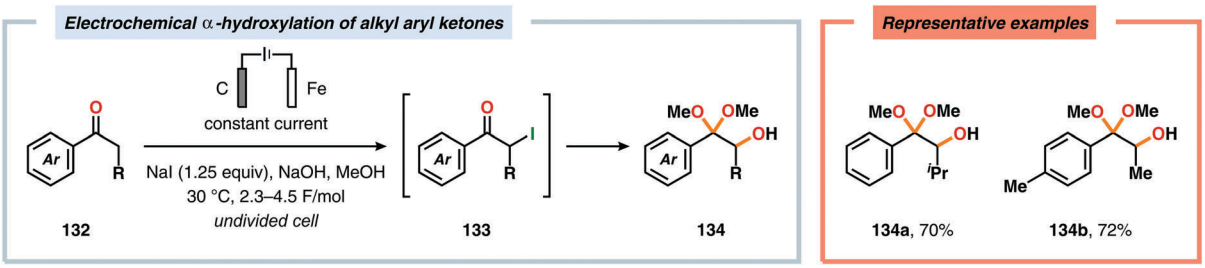
highlighting the concept of using toluene derivatives as alkylating agents.[398] Moreover, Nishiguchi and coworkers exploited the electrochemical oxidation of terminal acetylenes using a divided cell in the presence of NaI as the supporting electrolyte. The electrochemical method efficiently provided the desired 1-iodoacetylenes in good to excellent yields (Scheme 36).[404]

2.3. Electrochemical C–H bond oxygenation

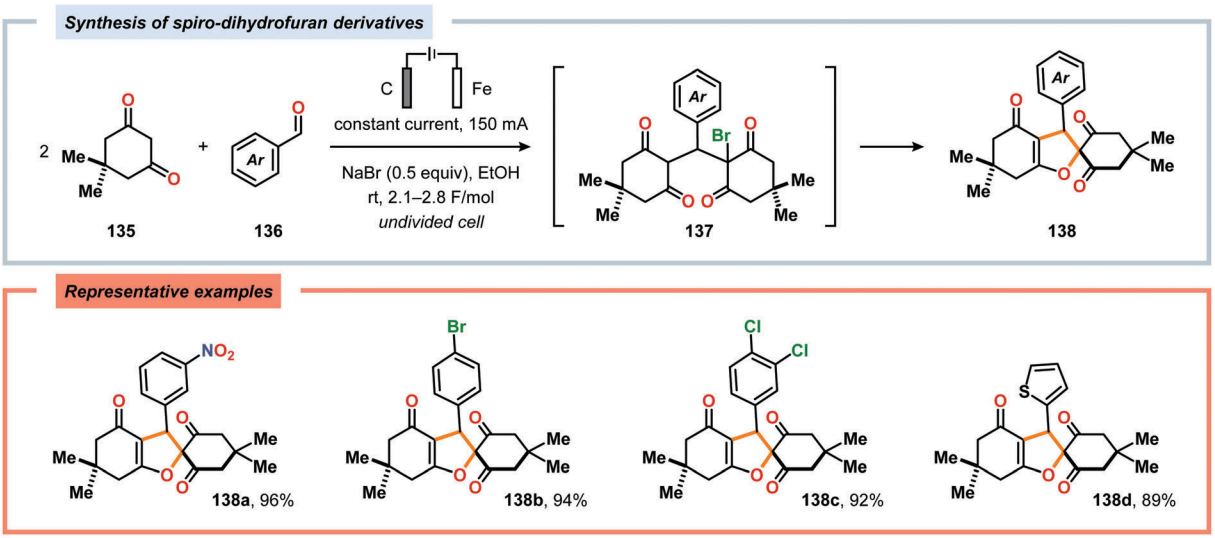
α-Oxygenation of carbonyl compounds can be accomplished using alkali metal halide salts. These halide salts can undergo anodic oxidation, generating α-halo compounds or carbon-centered α-radicals. Seminal work by Shono and coworkers demonstrated that iodide-mediated α-oxidation of aldehydes and ketones delivered the corresponding α-hydroxylated acetals or ketals.[405] More recently, Elinson and coworkers have also studied the sodium halide/sodium hydroxide system for indirect electrochemical α-oxidation of carbonyl compounds.[406,407] Based on these results, the authors employed the mediatory system NaI/NaOH for converting alkyl aryl ketones into the corresponding α-hydroxy ketals (Scheme 37). These reactions were proposed to proceed *via* α-iodoketone **133**.[408] The iodide-mediated oxidation system was subsequently employed for α-hydroxylation of cyclohexanones[409] and piperidin-4-ones.[410] Bromide-mediated α-oxidation has also been used for accessing spiro-dihydrofuran derivatives from dimedone and aldehydes (Scheme 38).[411] Furthermore, both intermolecular[412] and intramolecular[413–428] protocols for C–C bond formation based on halide-mediated anodic oxidation of carbonyl compounds and related derivatives have also been disclosed.

The Wang group has shown that iodide-mediated anodic oxidation of aryl methyl ketones can be exploited for synthesis of α-ketoesters (Scheme 39),[429–431] α-ketoamides (Scheme 40)[432] and isatins (Scheme 41).[433] Here, a radical-based pathway in which an iodine radical ($I^{\bullet}$) is proposed to undergo a HAT event with the aryl methyl ketone to deliver hydrogen iodide and an α-carbon-centered radical (**140**). The intermediate radical can subsequently be trapped with dioxygen to afford peroxyl radical **141**, which is readily converted to the corresponding ketoaldehyde compound. Finally, nucleophilic addition to this intermediate and a further oxidation event yields the desired product. α-Oxygenation of aliphatic aldehydes has been accomplished through a dual electrochemical/organocatalytic strategy (Scheme 42).[434] In this transformation, the generated α-iminyl radical cation is intercepted with TEMPO to afford the α-oxygenated product **148** upon

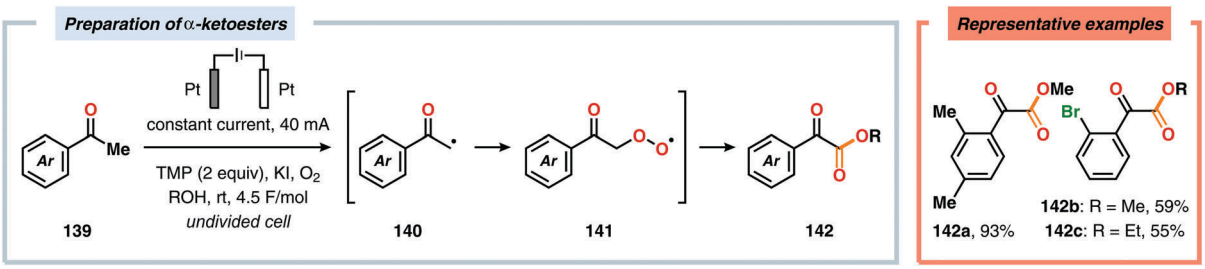
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



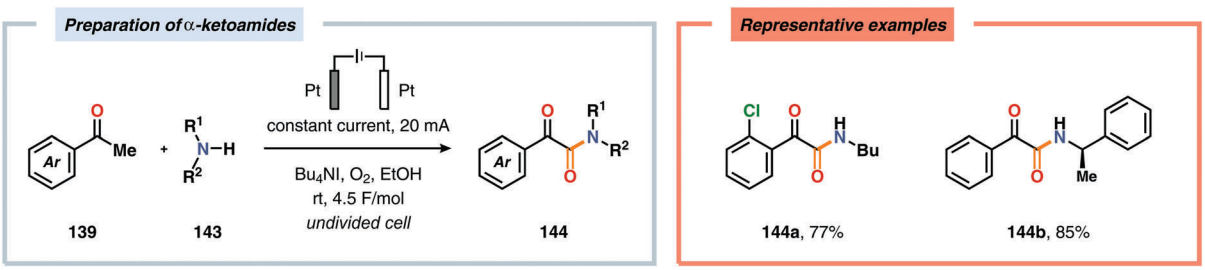
Scheme 37 Indirect electrochemical α-hydroxylation of alkyl aryl ketones.



Scheme 38 Electrochemical synthesis of spiro-dihydrofuran derivatives.

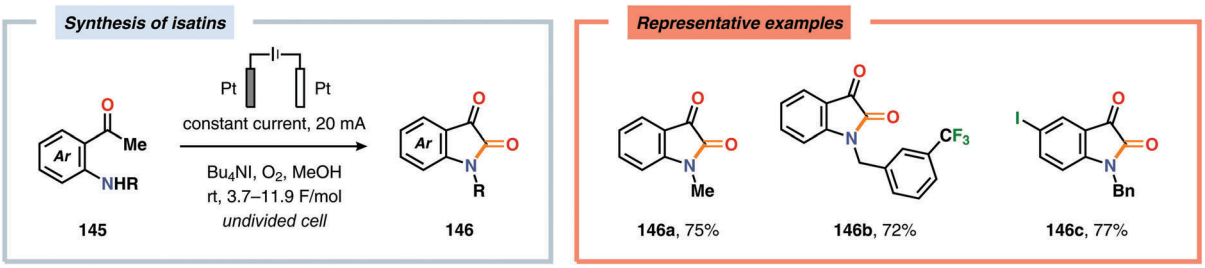


Scheme 39 Electrochemical preparation of α-ketoesters. TMP = 2,2,6,6-tetramethylpiperidine.

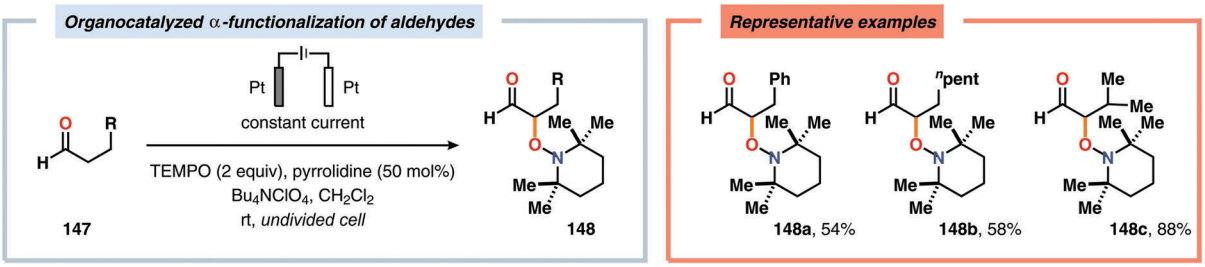


Scheme 40 Preparation of α-ketoamides through anodic oxidation.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

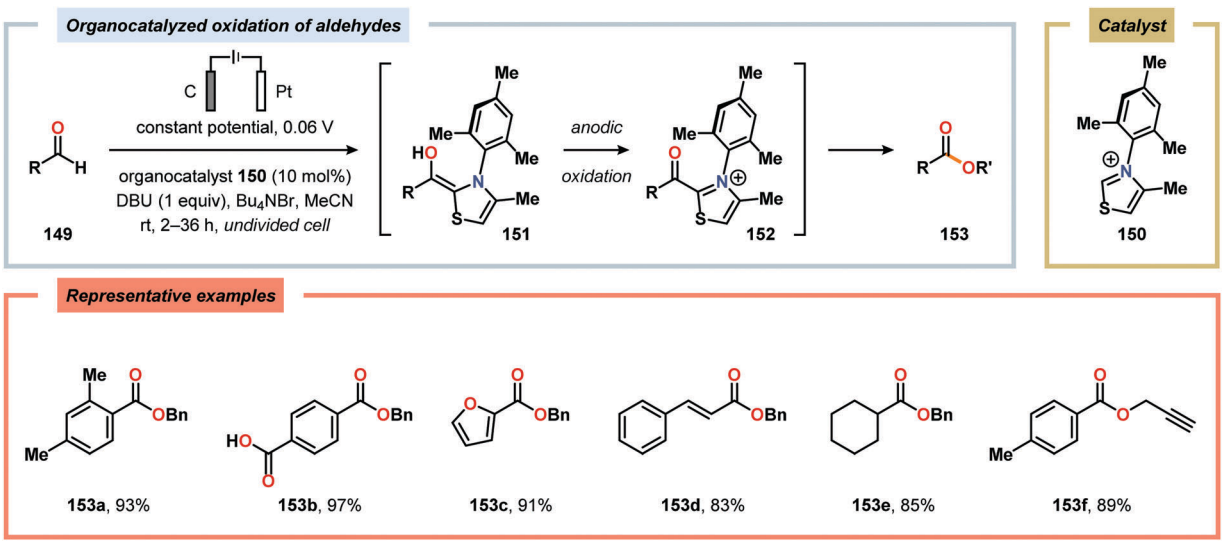


Scheme 41 Iodide-mediated synthesis of isatins.



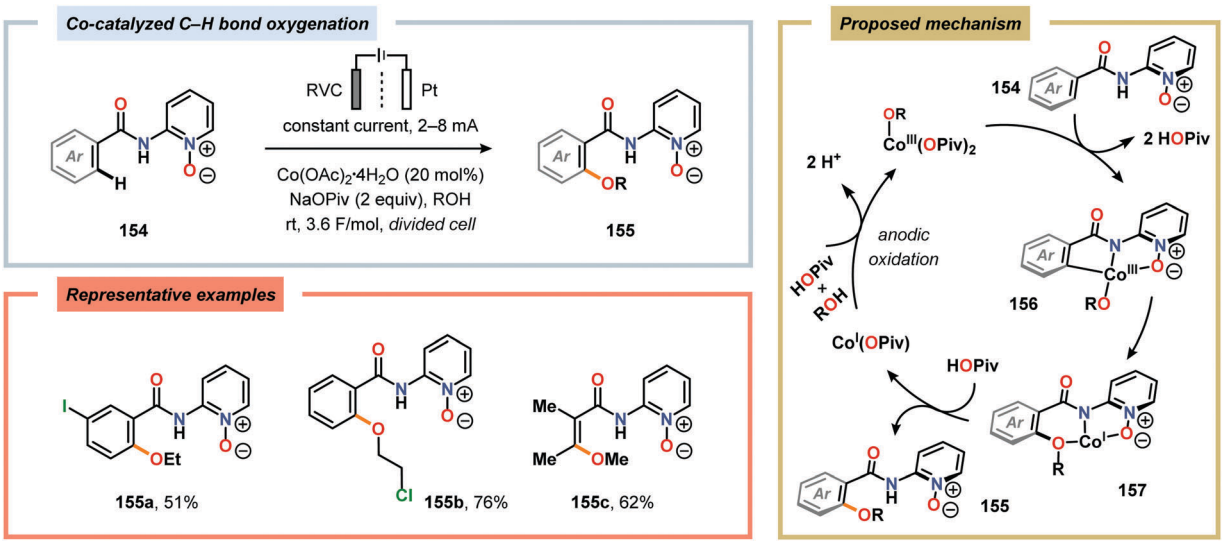
Scheme 42 Dual electrochemical/organocatalyzed α-oxygenation of aldehydes.

hydrolysis of the resultant iminium ion. In addition, various chiral *sec*-amines were evaluated in order to develop an enantioselective protocol. However, only moderate enantioselectivities were obtained using the Hayashi–Jørgensen catalyst.[435,436] The merger of electrochemistry with other organocatalytic activation modes was utilized by Boydston and coworkers for conversion of aldehydes to esters (Scheme 43). The method interfaced N-heterocyclic carbene-based organocatalysis with anodic oxidation for achieving efficient two-electron oxidation of the *in situ* generated Breslow intermediate (**151**) to produce an electrophilic 2-acylazolium species (**152**).[437–439] Subsequently, the authors disclosed a method for the direct conversion of aldehydes to thioesters *via* integration of organocatalysis and electrosynthesis.[440] Based on this dual catalytic activation mode, Brown and coworkers successfully demonstrated that the oxidative esterification of aldehydes[441] as well as amidation of aldehydes[442] could be conducted in an undivided electrochemical flow cell. Recently, the Ackermann laboratory exploited the powerful nature of dual catalytic methods for achieving C(sp^2)–H oxygenation of aromatic and alkene moieties.[443] Here, the combination of cobalt catalysis and electrochemical oxidation enabled C–H alkoxylation under mild reaction conditions at ambient temperature (Scheme 44). Employing electrical current as an oxidant obviated the need of stoichiometric amounts of Ag(I) salts through the use of an earth-abundant cobalt redox manifold. Furthermore, the developed protocol exhibited high levels of chemo-, regio- and diastereoselectivity. Electrooxidative C–H

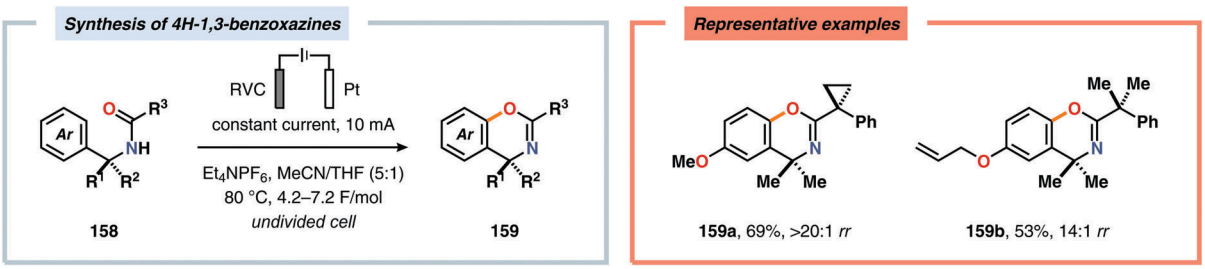


Scheme 43 Organocatalyzed anodic oxidation of aldehydes.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 44 Electrochemical cobalt-catalyzed C(sp^2)–H alkoxylation.

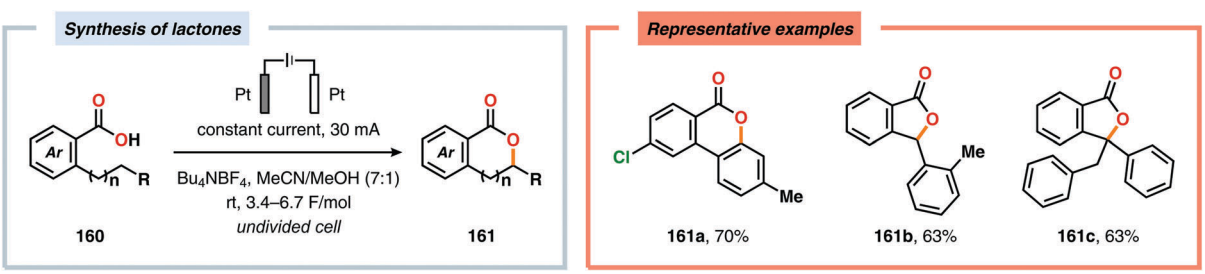


Scheme 45 Synthesis of 4*H*-1,3-benzoxazines through electrochemical C–H oxygenation.

activation with weakly coordinating benzoic acids has also been accomplished using rhodium-[444] or ruthenium catalysis.[445]

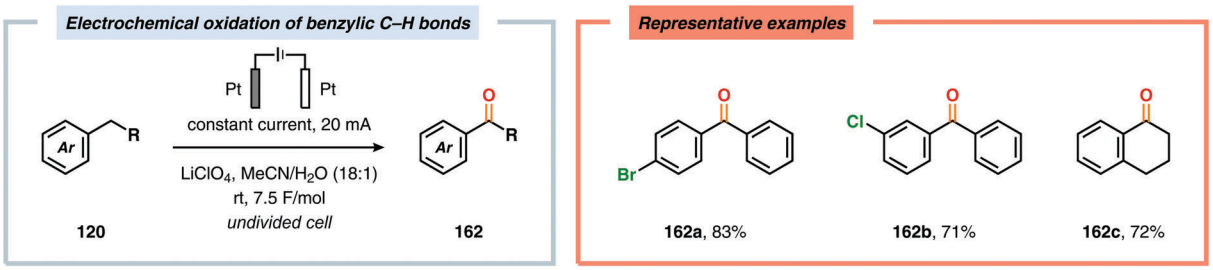
Electrochemical aromatic C(sp^2)–H oxygenation has also been utilized for carrying out oxidative annulations. Xu and coworkers disclosed an electrochemical method featuring the direct electrolysis of *N*-benzylamides for preparing a range of substituted 4*H*-1,3-benzoxazines (Scheme 45). The electrolysis was proposed to facilitate anodic oxidation of the electron-rich aromatic nucleus, triggering regioselective cyclization to afford a cyclohexadienyl radical. Finally, rearomatization is realized upon abstraction of an electron and a proton, furnishing the desired benzoxazine **159**.[446] The Zeng group demonstrated that carboxylic acids can be anodically oxidized to generate the corresponding carboxylate radical. The generated radical could subsequently undergo intramolecular addition to electron-rich aromatic rings to produce the C–O coupled lactone product (Scheme 46).[447–450] In addition to these examples, electrochemical oxygenation of furan derivatives[451–453] has also been reported and has, for example, been utilized for oxygenation of hispanolone[454] as well as preparation of dideoxynucleoside analogues.[455]

Development of strategies for oxygenation of C(sp^3)–H bonds constitutes a compelling approach for delivering alcohol and carbonyl derivatives. Historically, chemists have confided to using stoichiometric, and often toxic, oxidants for carrying out these transformations, resulting in the formation of considerable



Scheme 46 Electrochemical lactonization of C–H bonds.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



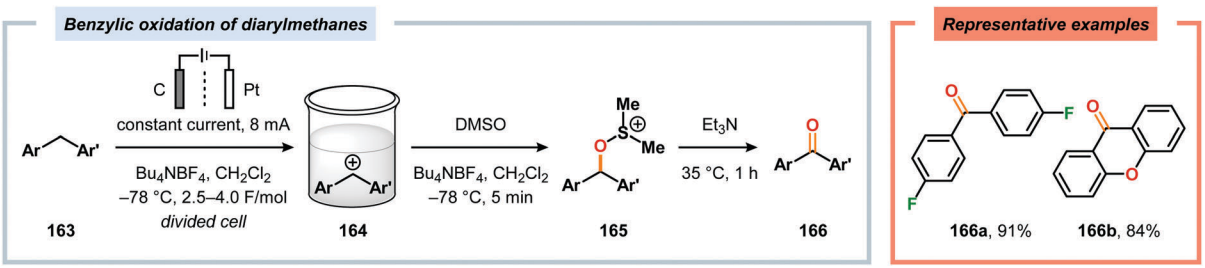
Scheme 47 Electrosynthesis of ketones through anodic oxidation of benzylic methylenes.

amounts of waste. However, the demand for more sustainable and atom economical methodologies has gained significant attention among the scientific community.[456] Electrochemical activation of toluene derivatives through initial oxidation of the aromatic core is an appealing platform for realizing C(sp^3)–H oxygenation of benzylic carbons. The aryl radical cation that is generated under these reaction manifolds can subsequently undergo deprotonation and a further oxidation to furnish a benzylic cation which can be trapped with various oxygen-based nucleophiles. Such an approach was implemented by the Wang group in which the benzylic cation was intercepted with water to furnish the benzylic alcohol, which underwent additional oxidation to give the corresponding ketone (Scheme 47).[457] Diarylcarbenium ions can successfully be generated and accumulated by low-temperature electrochemical oxidation using the "cation pool" method.[458–466] In the presence of DMSO, alkoxysulfonium ions serve as the key intermediates. Treatment with a base, such as Et_3N, or hydrolysis selectively affords the corresponding ketone[467] (Scheme 48) or alcohol[468] (Scheme 49). The fact that the initial electrochemical oxidation and the product-forming steps are separated implies that the products are not exposed to the electrolysis conditions, thereby limiting overoxidation. An electrochemical protocol for anodic acyloxylation based on the acid–base reactions between acetic acid or trifluoroacetic acid and a solid-supported base has also been reported (Scheme 50).[469,470] It was found that the silica gel supported base was stable under the electrochemical conditions, permitting easy recovery and reuse of the solid-supported base. Indirect electrolysis using various mediators, such as TEMPO/Br^-,[164] 2,3-dichloro-5,6-dicyano-1,4-benzoquinone (DDQ)[471,472] and NHPI (Scheme 51),[473] can also be harnessed for facilitating oxygenation of benzylic methylenes. In addition, Little and coworkers demonstrated that the use of triarylimidazoles[474–477] as redox mediators allowed for selective benzylic oxygenation of unsymmetrical benzyl ethers (Scheme 52).[478–480]

The oxidation of unactivated C–H bonds typically requires high redox potentials, thus limiting the functional group compatibility and necessitates the use of inert solvents. Allylic oxidations are generally based on the use of toxic reagents, such as chromium and selenium, or scarce transition metal catalysts, such as Pd and Rh, which causes complications in industrial settings. The Baran group has addressed the issue of oxidizing unactivated C–H bonds through indirect electrolysis. After careful evaluation of potential mediators, a NHPI derivative (Cl_4NHPI) was demonstrated to be the optimal mediator for the allylic oxidation of a range of natural product inspired compounds, including sesquiterpene- and steroid-based substrates (Scheme 53).[481–483] The described protocol exhibited broad substrate scope, operational simplicity and high chemoselectivity, and could even be conducted on a 100 g scale. The reaction is believed to involve hydrogen atom abstraction from the olefinic substrate to the electrochemically generated *N*-oxyl radical species, resulting in regeneration of Cl_4NHPI (**178**) and an allylic radical (**179**). Interception of the allylic radical with electrochemically produced ^{t}BuOO$^\bullet$ furnishes allylic peroxide **180**, providing enone **181** upon elimination of ^{t}BuOH. Furthermore, direct electrochemical oxidation of cholesterol at the allylic position has been demonstrated by Sobkowiak and coworkers.[484–489] Baran and coworkers subsequently reported a method for selective functionalization of unactivated methylene and methine moieties utilizing quinuclidine (**183**) as redox mediator (Scheme 54). Here, an assortment of functional groups were demonstrated to be compatible with the developed chemoselective process.[490,491]

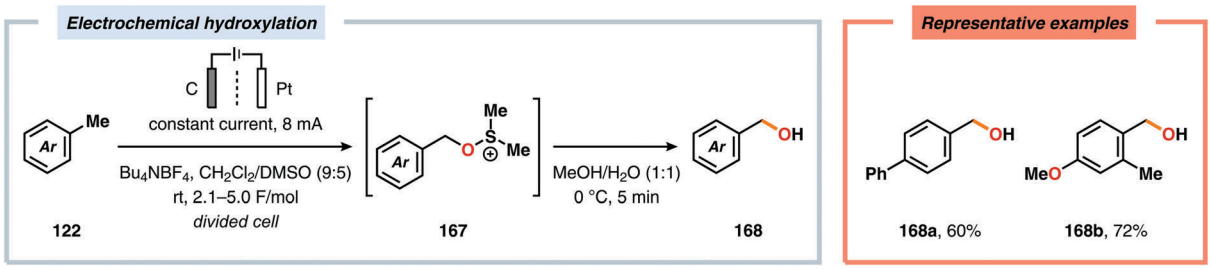
2.4. Palladium-catalyzed anodic C–H functionalization strategies

Palladium-catalyzed cross-coupling reactions have revolutionized C–C bond forming processes. Over the years, a variety

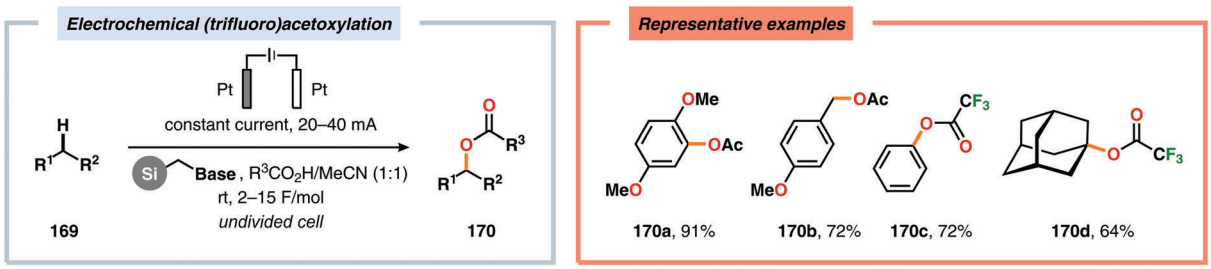


Scheme 48 Electrochemical oxygenation using alkoxysulfonium ions.

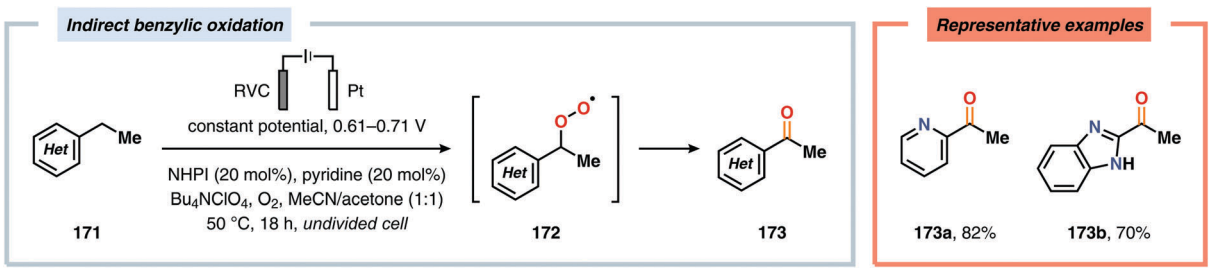
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



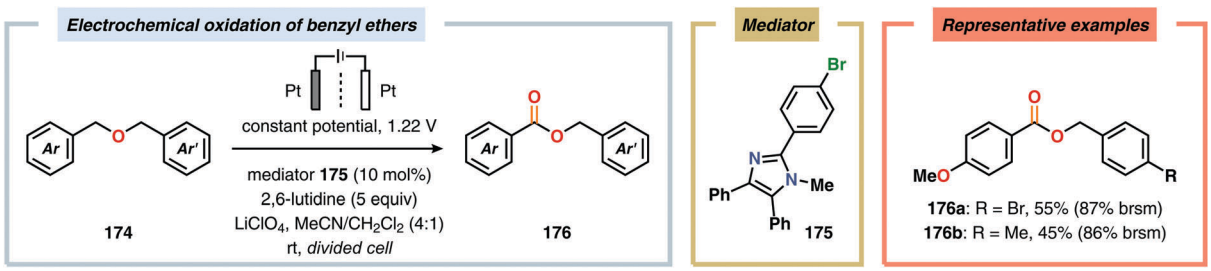
Scheme 49 Electrochemical hydroxylation using alkoxysulfonium ions.



Scheme 50 Electrochemical (trifluoro)acetoxylation of C(sp^2)–H and C(sp^3)–H bonds. Base = morpholine.



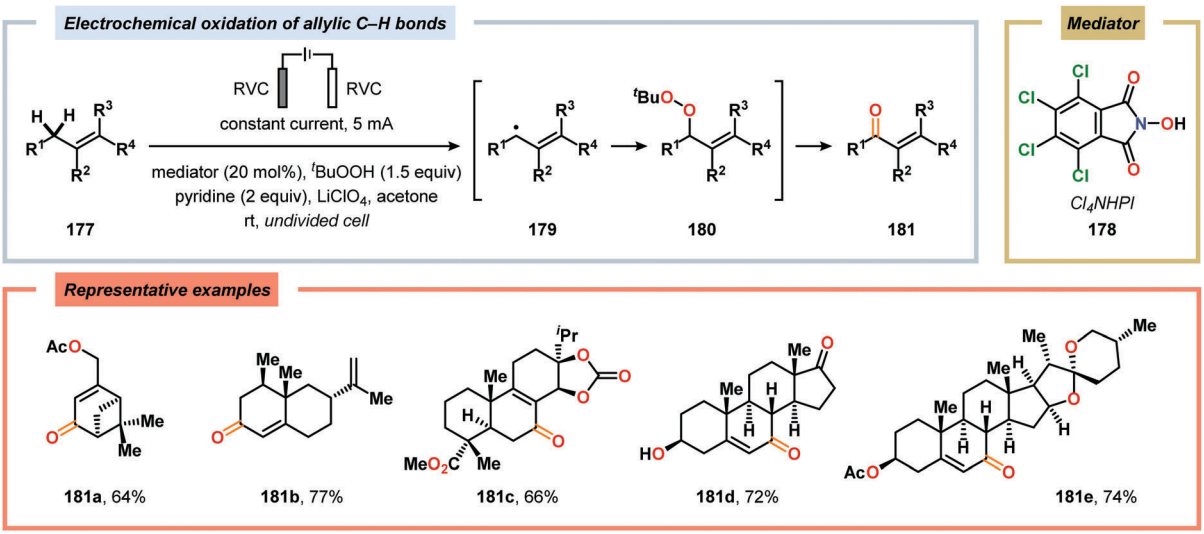
Scheme 51 Indirect electrochemical oxygenation using NHPI as mediator.



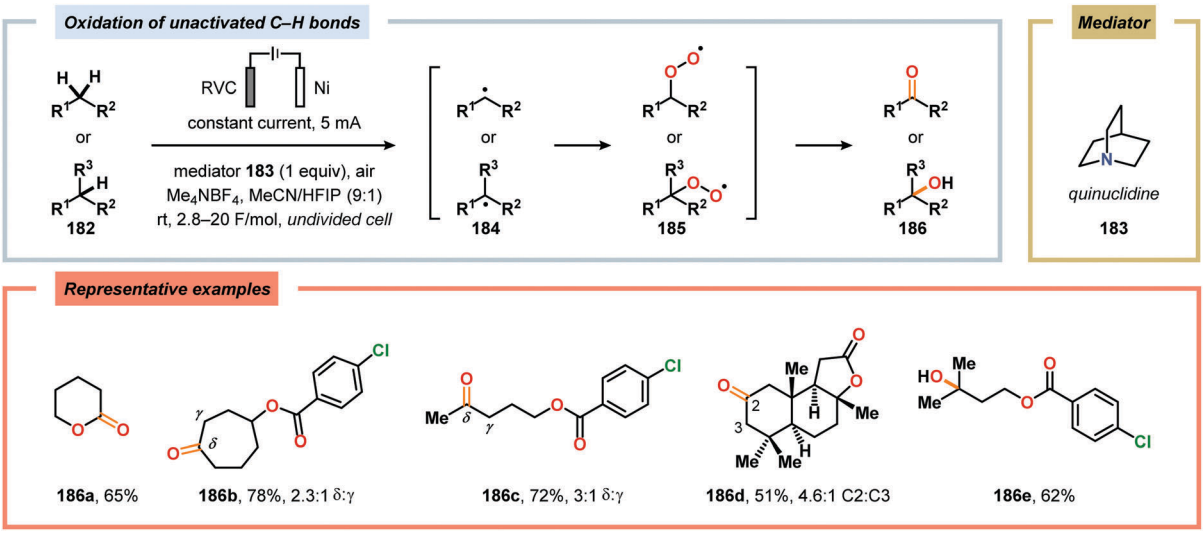
Scheme 52 Benzylic oxygenation using a triarylimidazole mediator.

of catalytic platforms, such as carbon–heteroatom coupling, α-arylation and decarboxylative coupling, based on Pd have been designed.[492] It is therefore not surprising that the synergistic merger of electrochemistry and Pd catalysis has been the subject of keen interest by the synthetic community.[70,493–497] Pioneering studies by Tsuji and Minato demonstrated a variant of the Wacker oxidation of terminal olefins by combining Pd-catalysis and electrochemical oxidation. In this system, benzoquinone is regenerated at the anode, which subsequently triggers the turnover-limiting reoxidation of Pd^0 to Pd^{II}.[498,499] More recently, related approaches have also been employed for carrying out Heck-type reactions (Scheme 55),[500,501] Glaser–Hay coupling[502] as well as homocoupling of arylboronic acids and derivatives thereof,[503] and oxidation of alcohols under anaerobic conditions.[504] Dual catalytic platforms based on Pd/TEMPO have been reported for electrooxidative synthesis of biaryls from arylboronic acids,[505–507] cross-coupling between terminal alkynes and arylboronic acids[508] as well as Wacker-type reactions[509] and cyclizations.[510] Furthermore, Moeller and coworkers have demonstrated that triarylamines are also effective mediators and have

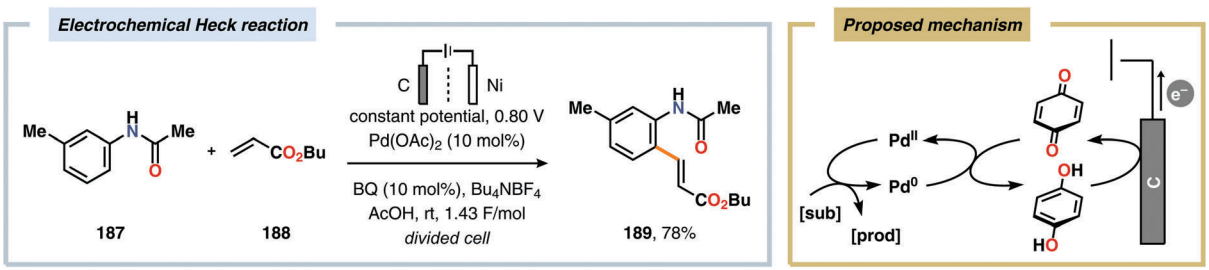
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 53 Indirect electrochemical allylic C–H oxidation.



Scheme 54 Indirect electrochemical oxidation of unactivated C–H bonds.

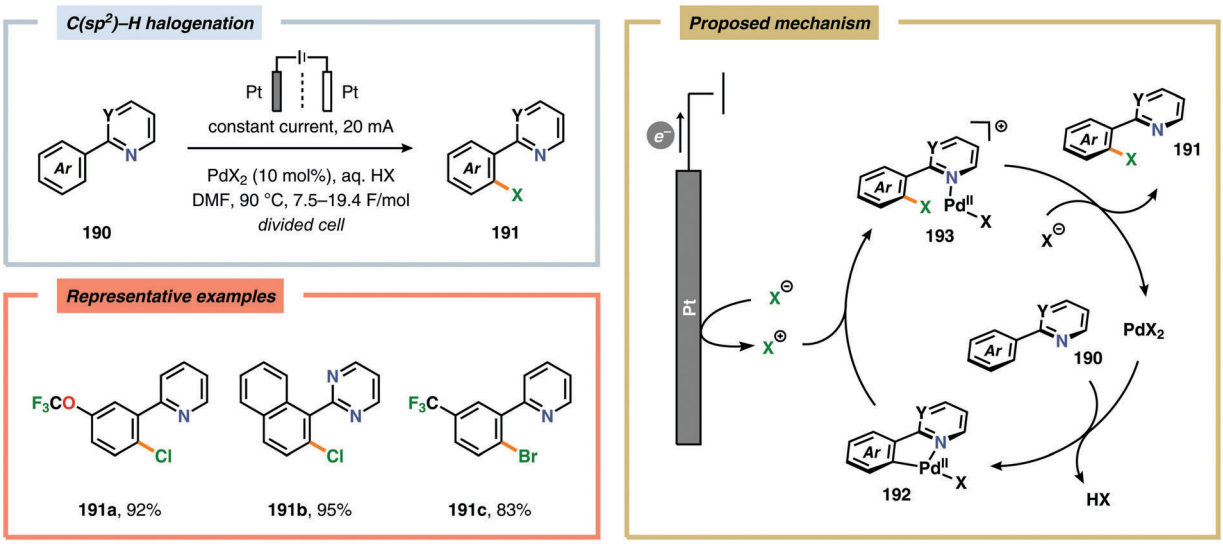


Scheme 55 Electrochemical Heck reaction using benzoquinone as mediator.

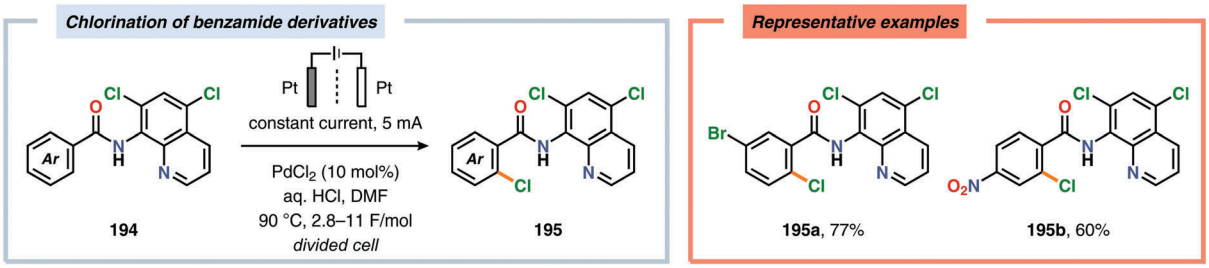
utilized this concept for site-selective functionalization of semiconducting chips.[511–516]

A variety of approaches have been reported for the C–H functionalization of arylpyridine derivatives *via* dual activation by transition-metal catalysis and electrochemical oxidation. In this regard, the Kakiuchi group has demonstrated that C(sp^2)–H halogenation of aromatics using hydrogen halides can be accomplished (Schemes 56 and 57). It was proposed that anodic oxidation of the halide ion produces a halonium ion, which subsequently reacts with the generated palladacycle to

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 56 Regioselective halogenation of arenes through dual Pd catalysis and electrochemical oxidation.

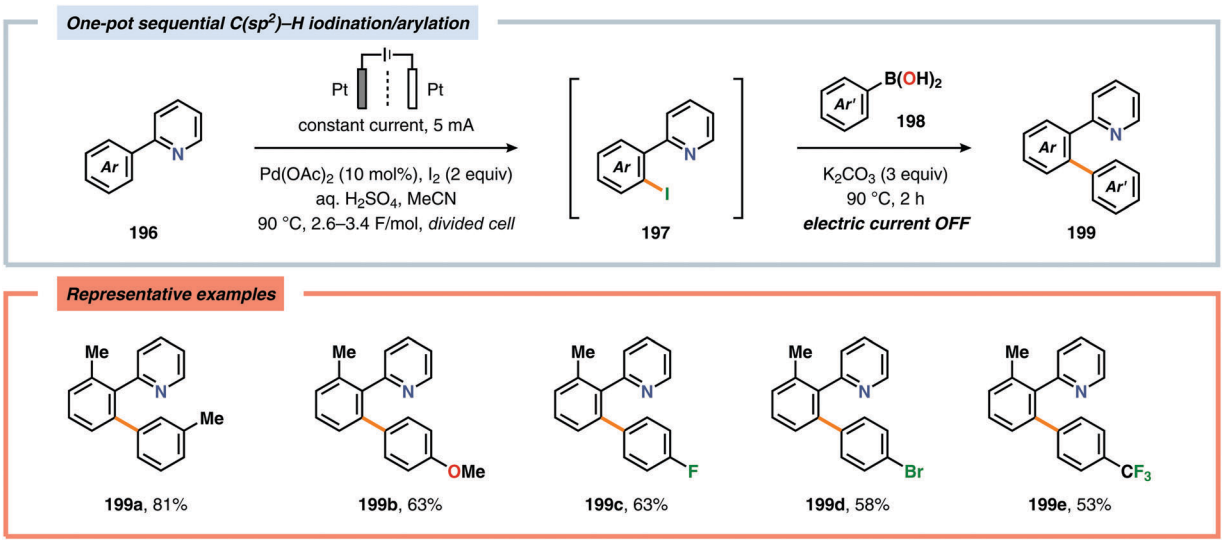


Scheme 57 Electrochemical C(sp^2)–H chlorination of benzamide derivatives.

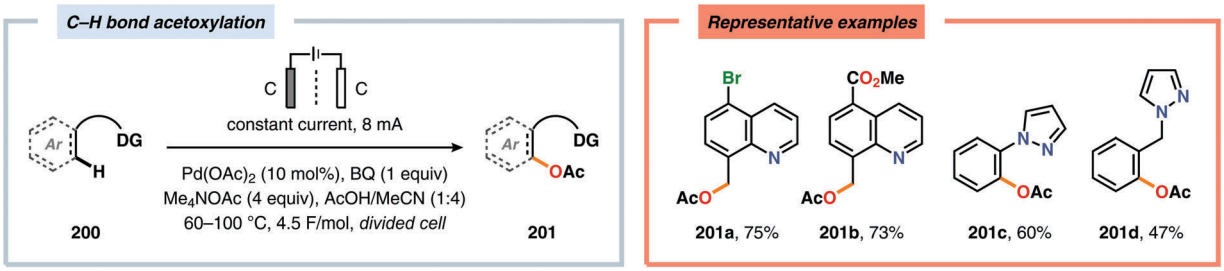
provide the halogenated product while regenerating the Pd^{II} catalyst.[517–520] Expanding on this concept, the Kakiuchi group disclosed a method for regioselective homocoupling of arylpyridines in the presence of I_2 employing electrooxidative Pd catalysis.[521] Additionally, protocols for perfluoroacetoxylation,[522] perfluoroalkylation[523,524] and phosphorylation[525,526] of arylpyridines have been established by the Budnikova group. A procedure for sequential electrochemical C–H iodination followed by a Suzuki–Miyaura coupling has also been developed by Kakiuchi and coworkers. Here, I_2 was employed as the iodine source for the electrochemical C–H iodination. Upon completion of the iodination reaction, the electric current was simply switched off whereupon the arylboronic acid was introduced to the reaction mixture to furnish the corresponding cross-coupled product (Scheme 58).[527] Along similar lines, Suga and coworkers reported a dual Pd-mediated electrooxidative homocoupling of *p*-bromophenylacetylene with a subsequent Suzuki–Miyaura coupling. Discrimination between the two reaction sites, the terminal alkyne and the Br group, was realized by the mere on/off application of electricity.[528] Furthermore, the Sanford group recently disclosed a Pd-catalyzed electrochemical method for acetoxylation of C(sp^3)–H and C(sp^2)–H bonds (Scheme 59). It was noted that differences in the relative rates of competing oxidation reactions could prevail under chemical *versus* electrochemical oxidation conditions, thereby resulting in the formation of complementary products. Additionally, the differences in reaction conditions for the chemical and electrochemical reactions, such as solvents as well as the addition of a supporting electrolyte, can translate to fundamental changes in catalyst performance and/or reaction outcomes, and will be imperative when designing new electrochemical platforms.[529]

Merging Pd catalysis and electrochemical oxidation has also proven to be a successful concept for carrying out C–H functionalization on oxime-based motifs. In this regard, Mei and coworkers have examined the Pd-catalyzed oxime-directed C(sp^2)–H acetoxylation employing an anodic oxidation strategy (Scheme 60). In this transformation, it was proposed that the rate-determining oxime-assisted C–H activation generates the desired palladacycle. This Pd^{II} species is subsequently oxidized at the anode to produce the corresponding Pd^{IV} intermediate, which can undergo reductive elimination to deliver the acetoxylation product, thus regenerating Pd^{II} to close the catalytic cycle.[530] Shortly thereafter, the authors reported a dual Pd/electrochemical strategy to achieve C(sp^2)–H methylation and acylation (Scheme 61). Here, reductive elimination was suggested to occur from a high-valent Pd intermediate, presumably Pd^{III} or Pd^{IV}.[531] Finally, functionalization of unactivated C(sp^3)–H bonds has also been reported using oximes as directing groups. Along similar lines, C–H activation furnishes a Pd^{II} palladacycle, which subsequently is oxidized at the anode to

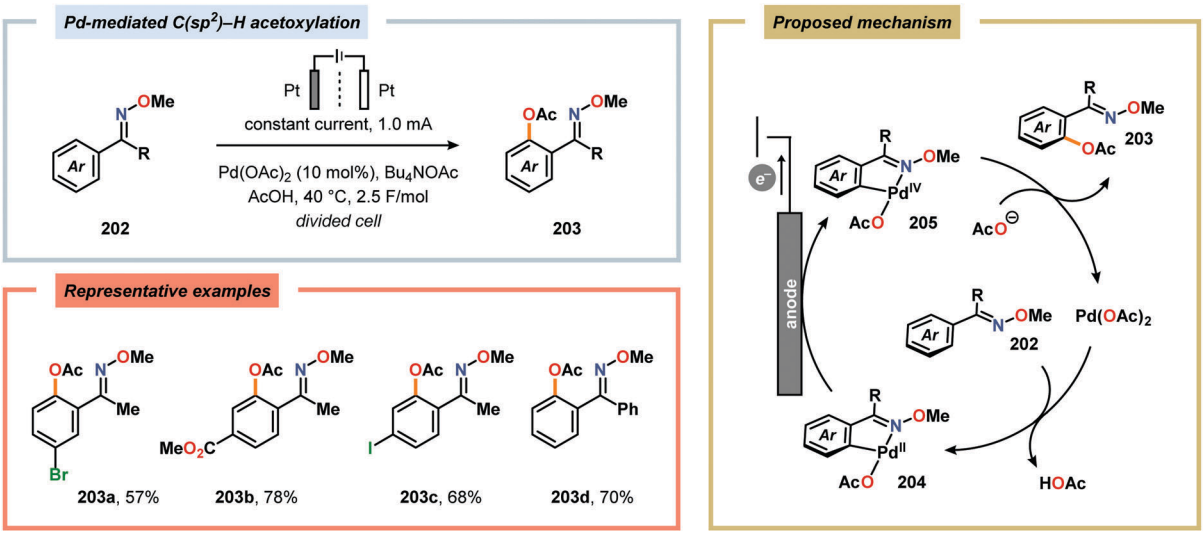
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 58 One-pot C–H iodination/arylation of arylpyridines.



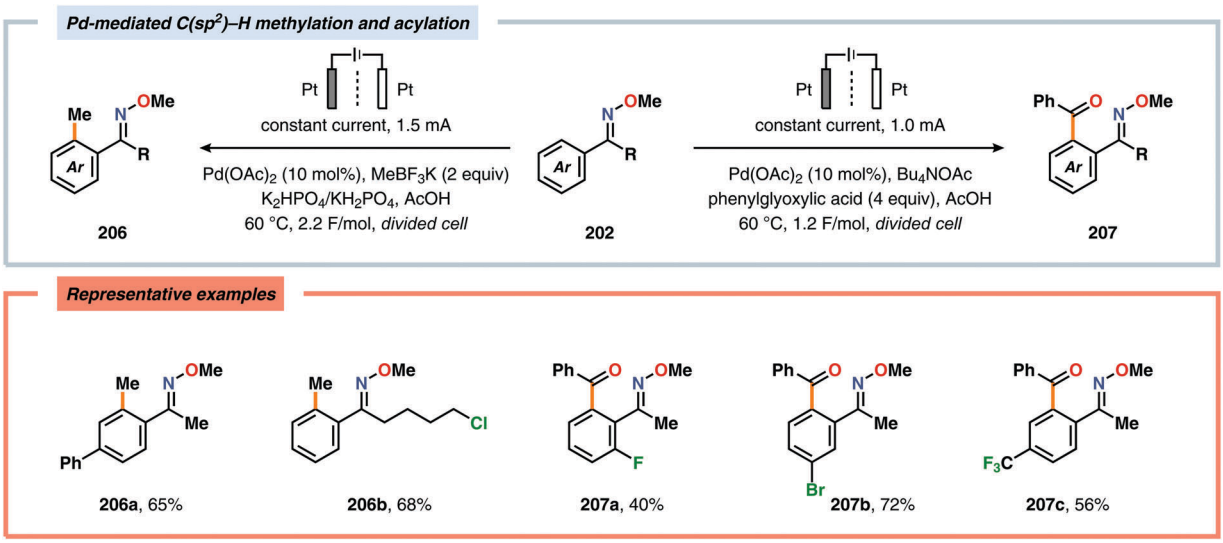
Scheme 59 Pd-catalyzed electrochemical method for acetoxylation of C(sp^3)–H and C(sp^2)–H bonds.



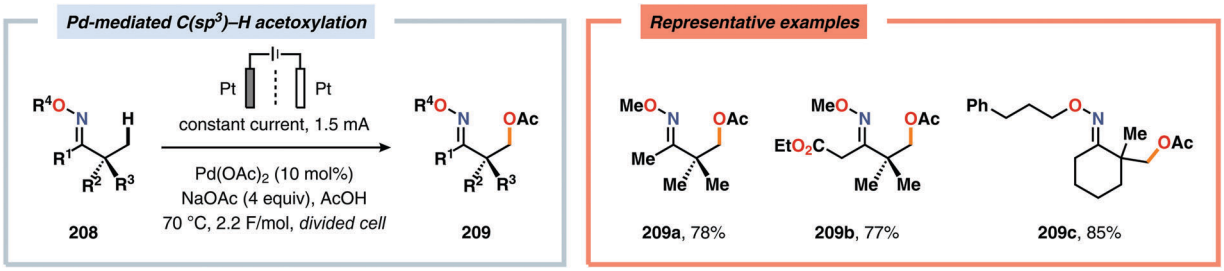
Scheme 60 Pd-catalyzed C(sp^2)–H acetoxylation *via* anodic oxidation.

give a high-valent Pd species, presumably a Pd^{IV} intermediate. Generation of the high-valent Pd center promotes the otherwise challenging C–O reductive elimination event to deliver the oxygenated product while regenerating Pd^{II}. In addition to carrying out C(sp^3)–H acetoxylation (Scheme 62), the dual Pd/electrochemical strategy could also be conducted with a variety of oxyanion coupling partners, such as methoxide, tosylate and trifluoroacetate.[532] The catalytic protocols described here

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 61 Pd-mediated C(sp^2)–H methylation and acylation.



Scheme 62 Dual Pd/electrochemical C(sp^3)–H functionalization.

proceed under mild reaction conditions and in the absence of exogenous ligands, representing sustainable alternatives to conventional platforms for C–H functionalization that rely on stoichiometric chemical oxidants.

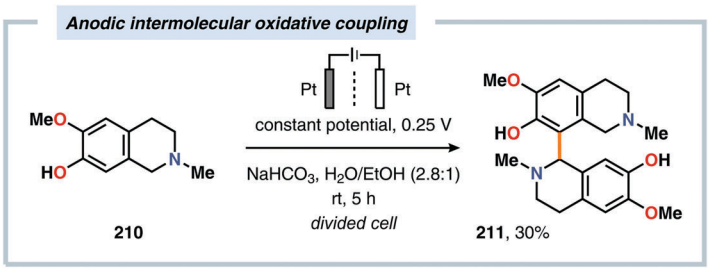
2.5. Electrochemical oxidative cross-couplings

Transition metal-catalyzed cross-coupling reactions serve as an integral part of contemporary synthetic chemistry for the rapid and selective construction of complex architectures from simple building blocks. However, these catalytic bond forming platforms require the use of prefunctionalized substrates, which adds additional preceding synthetic steps. This inevitably culminates in the generation of undesired stoichiometric amounts of waste. In recent years, constructing C–C bonds through the direct coupling of two C–H coupling fragments has emerged as a highly appealing concept. This effective and straightforward strategy not only circumvents the need for using prefunctionalized reagents but also considerably enhances the atom- and step-economy. In principle, dihydrogen (H_2) is the sole (by)product resulting from forging the two C–H bonds; however, typically a sacrificial oxidant is required to accept the H_2, hence the name oxidative cross-coupling or cross-dehydrogenative coupling (CDC). Oxidative cross-coupling is undoubtedly an attractive catalytic platform, yet conceptually and practically challenging due to, for example, the low reactivity of the C–H bonds in combination with overcoming the limitations in site-selective functionalization.[533–536] Moreover, the development of radical-mediated oxidative coupling strategies has also gained significant attention from the scientific community.[537]

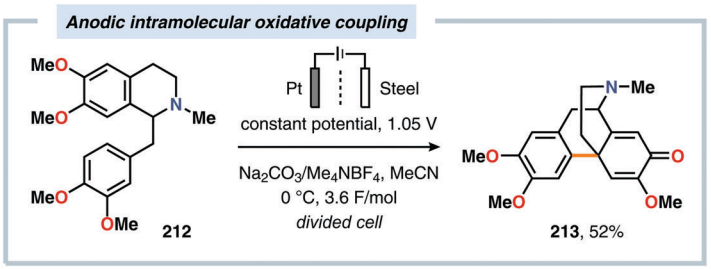
Despite the relatively simple motif, the synthesis of biphenols through oxidative cross-coupling of unprotected phenols has proven to be challenging and often relies on protocols involving expensive catalyst systems or reagents. In addition to producing the desired cross-coupled biphenolic product,[538] the reaction is also accompanied by products derived from oxidative dimerization, higher molecular weight polymers as well as compounds housing C–O coupling motifs.[539–543] Seminal work on the intermolecular anodic oxidative coupling of phenols demonstrated that corypalline (**210**) could be electrochemically oxidized to the corresponding dimer (**211**, Scheme 63).[544–548] However, attempts to carry out the related intramolecular C–C coupling on laudanosine (**212**) resulted in oxidative dearomatization of one of the phenolic moieties and furnished *O*-methylflavinantine (**213**, Scheme 64).[549–553] Similar reactivity has also been observed for the related alkaloids amurine, oxocrinine, oxomaritidine and pallidine.[554–556]

Pioneering studies by the Waldvogel group demonstrated that the direct anodic oxidation of 2,4-dimethylphenol (**214a**) produced the *ortho*,*ortho*-coupled biphenol **215a**, albeit in

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

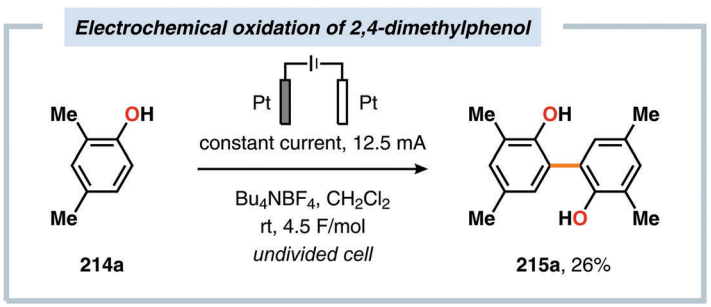


Scheme 63 Anodic intermolecular oxidative coupling of corypalline.

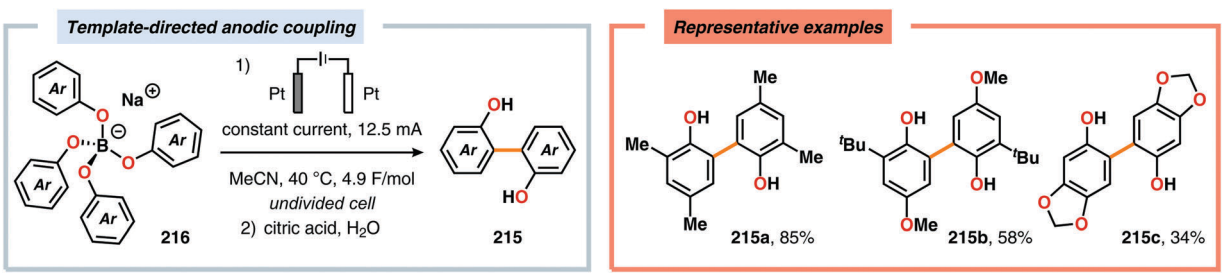


Scheme 64 Anodic intramolecular coupling of laudanosine.

rather poor yield (Scheme 65).[557] The authors subsequently devised a template-directed method for the anodic coupling of substituted phenols to the corresponding 2,2′-biphenols using tetraphenoxy borate derivatives (Scheme 66). Although the developed protocol was limited to electron-rich substrates, the electrochemical synthesis could be conducted on a kilogram scale.[558,559] In addition to generating the desired biphenol product, several additional coupling motifs can also be produced.[560–565] To circumvent the lack of selectivity, Waldvogel and coworkers realized that boron-doped diamond (BDD) electrodes[22] gave almost exclusive selectivity for the *ortho*-coupling product (Scheme 67).[566,567] It was rationalized that the observed selectivity originated from the high chemical and electrochemical stability of the BDD electrodes in combination with the high overpotential for oxygen evolution in aqueous media and

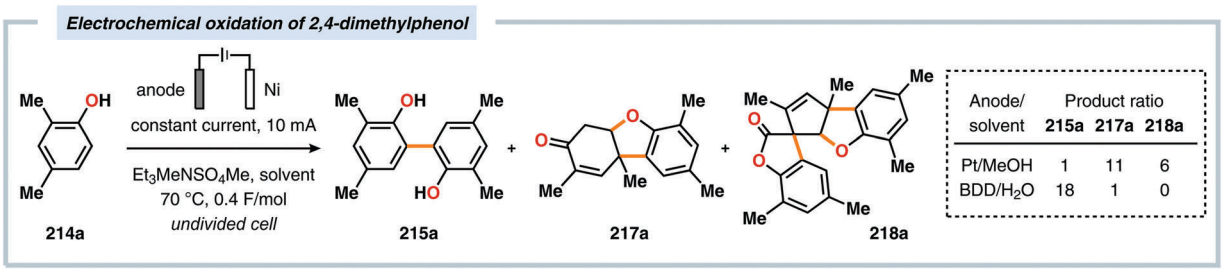


Scheme 65 Electrochemical oxidation of 2,4-dimethylphenol.

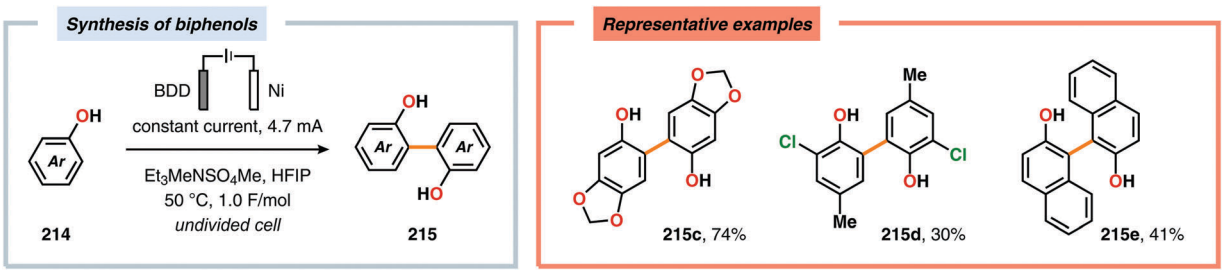


Scheme 66 Template-directed anodic coupling of phenols.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



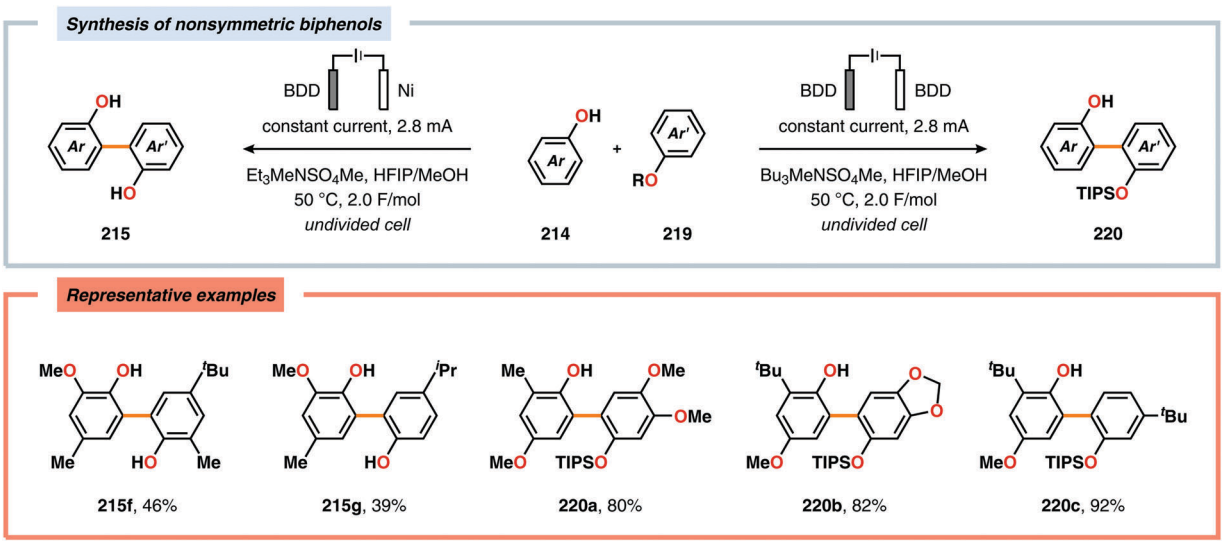
Scheme 67 Product diversity from the electrochemical oxidation of 2,4-dimethylphenol.



Scheme 68 Electrochemical synthesis of biphenols using a BDD anode.

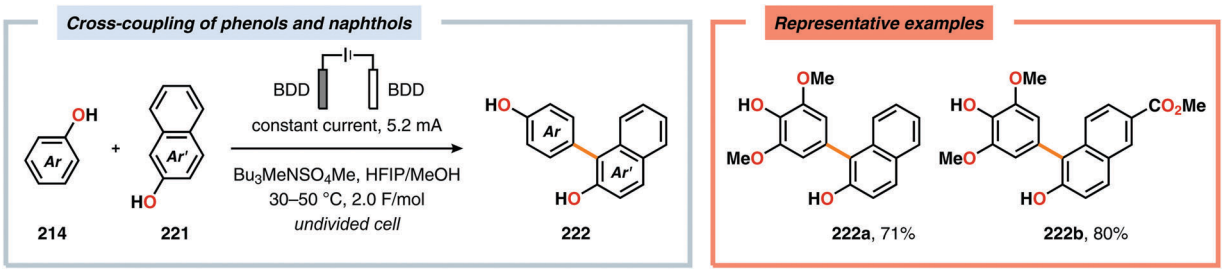
efficient formation of reactive alkoxy or hydroxyl radicals at the anode that can serve as mediators.[568–570] Interestingly, by simply switching to fluorinated solvents/mediators, such as hexafluoroisopropanol (HFIP), allowed for the creation of a powerful electrolyte system and significantly expanded the scope and efficiency for the anodic coupling of simple phenols[571] (Scheme 68) as well as guaiacol derivatives[572–574] on BDD electrodes. Protocols for synthesis of nonsymmetric biphenols (Scheme 69),[575–577] and anodic cross-coupling of phenols with naphthols[578] (Scheme 70) as well as aniline derivatives[579] (Scheme 71) have also been addressed.[580,581]

Metal-free electrochemical methodologies that facilitate anodic cross-coupling between phenols and electron-rich arenes have also been developed. The Waldvogel group discovered that this reaction could be accomplished employing a BDD anode in the presence of HFIP. It was established that HFIP had a crucial role in the reaction as no conversion was observed in the absence of HFIP but were able to rule out the possibility of HFIP acting as a redox mediator.[55,582] Since the phenol and arene components have similar oxidation potentials, the high level of chemoselectivity was rationalized by oxyl–HFIP hydrogen bonding interactions, which stabilize the produced spin centers.[583–585] The hydrogen-bond stabilized spin center undergoes HAT with the phenol (**214**), which produces the more easily oxidized phenoxyl radical **227**. Radical **227** is subsequently trapped by the electron-rich arene (**118**), which gives intermediate **228** upon deprotonation. This is followed by a second HAT event, resulting

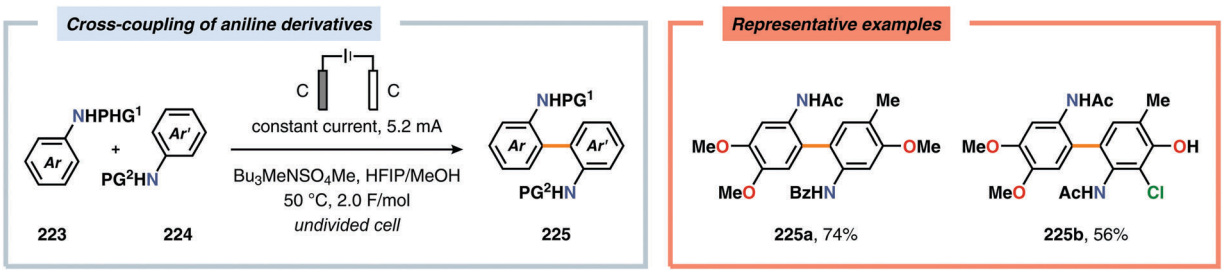


Scheme 69 Synthesis of nonsymmetric biphenols through anodic cross-coupling.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



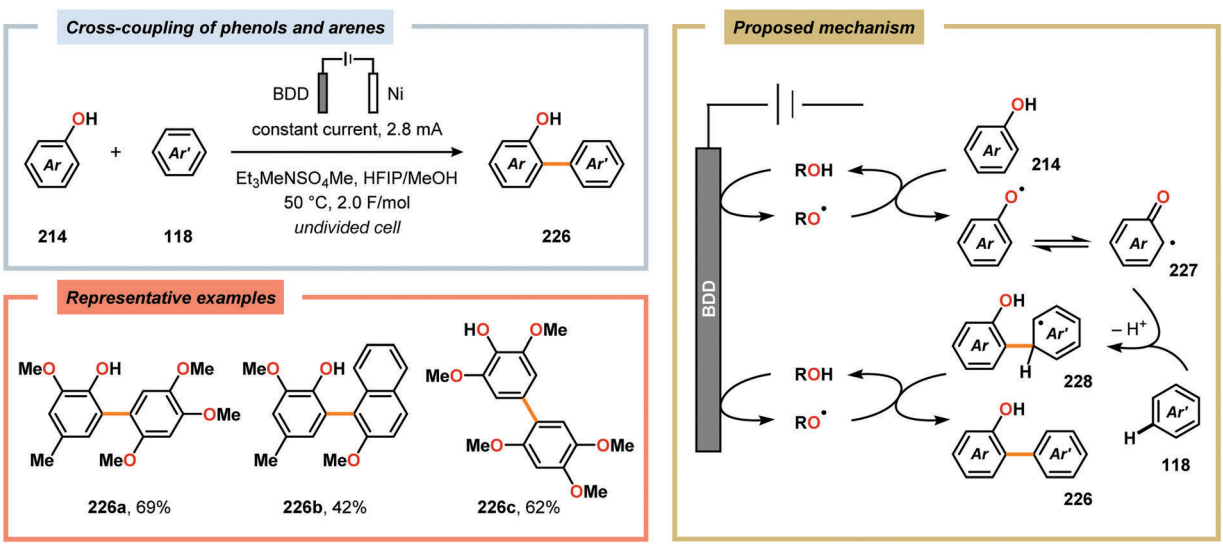
Scheme 70 Anodic cross-coupling of phenols and naphthols.



Scheme 71 Anodic cross-coupling of aniline derivatives. PG = protecting group.

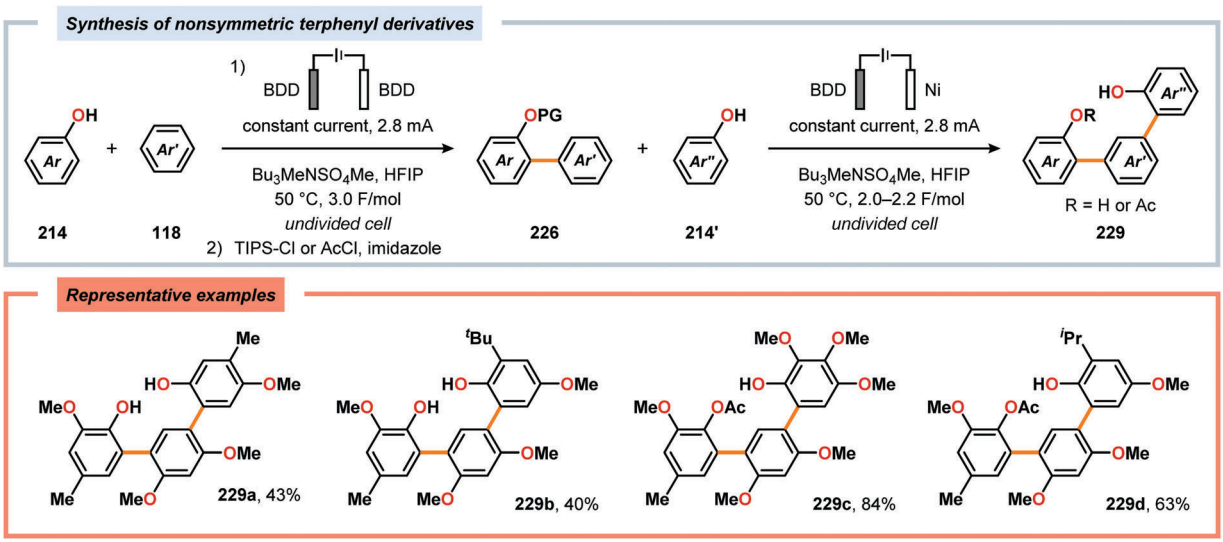
in rearomatization and furnishes the cross-coupled biaryl **226** (Scheme 72).[55,582] This strategy was subsequently extended to enable the development of anodic cross-coupling protocols for synthesis of *meta*-terphenyl-2,2″-diols (Scheme 73),[586,587] 2,5-bis(2-hydroxyphenyl)thiophenes[588] (Scheme 74) and dibenzofurans.[589,590]

Since the high oxidation potential of electron-deficient arenes, such as nitroarenes, limits direct anodic oxidation of these species, anodic cross-coupling reactions between electron-deficient arenes and nucleophiles engage in a nucleophilic aromatic substitution (S_NAr) type pathway. This two-step process can follow two mechanistically discrete pathways (Scheme 75). The first step involves nucleophilic addition to the electron-deficient arene unit, resulting in either a σ^H or σ^X adduct (Meisenheimer complex). For the σ^H intermediate (**233**), the second step follows a two-electron mechanism in which formal loss of a hydride occurs and is referred to as nucleophilic aromatic substitution of hydrogen (NASH, S_N^H). However, rearomatization of the σ^X adduct (**235**), generated from attack of the *ipso* carbon, involves an initial single-electron oxidation, which triggers radical elimination of the heteroatom ($X^\bullet$) and is hence called nucleophilic aromatic substitution of a heteroatom (NASX, S_N^X).[591–593] A range of nucleophiles, including acetate,[594] alkoxides,[594,595] alkyl lithium or Grignard reagents,[596] amides,[597] amines,[594,595,597] enolates,[594,598] fluoride,[595] cyanide,[594,599] organophosphorus compounds,[600] tetraalkylborate salts[601] and thiolates,[595] have been utilized to affect nucleophilic aromatic substitution through electrochemical

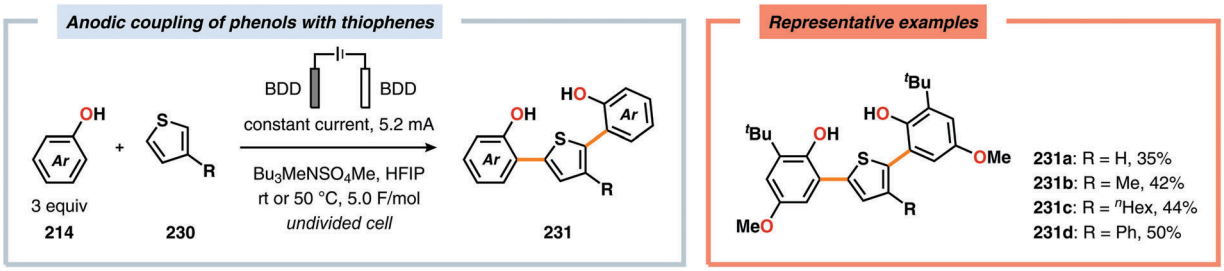


Scheme 72 Direct anodic cross-coupling of phenols and electron-rich arenes.

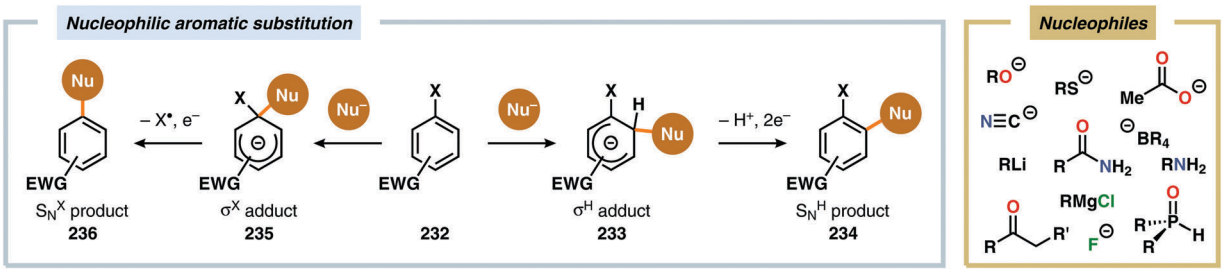
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 73 Synthesis of nonsymmetric *meta*-terphenyl-2,2″-diols. PG = protecting group.



Scheme 74 Electrochemical synthesis of 2,5-bis(2-hydroxyphenyl)thiophenes.

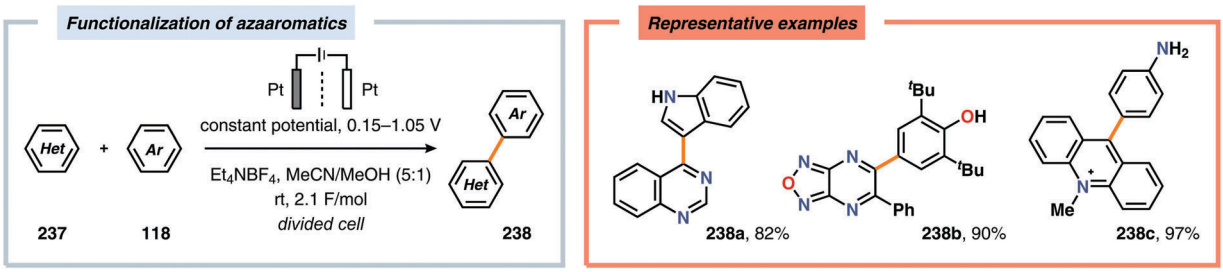


Scheme 75 Nucleophilic aromatic substitution *via* electrochemical oxidation.

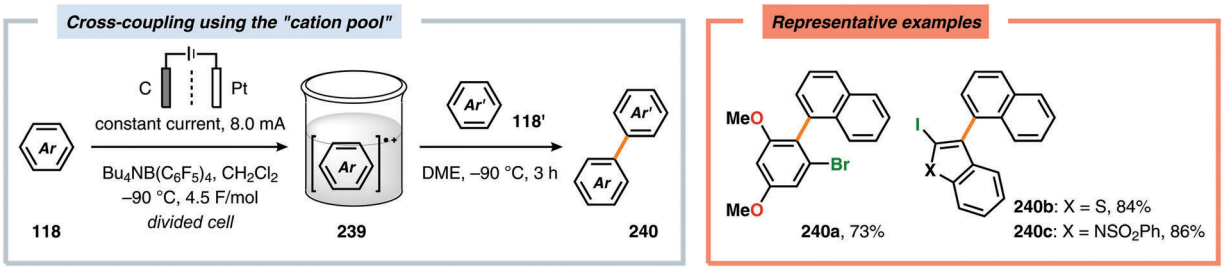
oxidation of the intermediate σ adducts. This concept was recently applied on azaaromatic scaffolds for the synthesis of asymmetrical bi(hetero)aryl motifs (Scheme 76).[602,603]

Although oxidative coupling between electron-deficient and electron-rich arenes, two moieties with distinctly different π-electronic properties, generally provides high chemoselectivity of the cross-coupled products over homocoupling as a result of the "electronic differential principle", anodic cross-coupling of unactivated electron-rich arenes has been achieved. Recently, the Yoshida group highlighted that employing the "cation pool" method allowed the merging of two unactivated aromatic compounds in a straightforward fashion. Here, initial anodic oxidation at −78 °C of the less electron-rich arene component (naphthalene) resulted in accumulation of the corresponding radical cation species. Subsequent introduction of the other aromatic moiety under non-oxidative conditions at −90 °C avoided nonselective oxidation of the starting material and overoxidation of the biaryl product (Scheme 77). Notably, the high regioselectivity of the developed cross-coupling method can be estimated based on the spin density of the radical cation and the highest occupied molecular orbital (HOMO) coefficients of the nucleophilic aromatic partner.[604,605] The use of parallel

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 76 Electrochemical functionalization of azaaromatics through nucleophilic aromatic substitution.

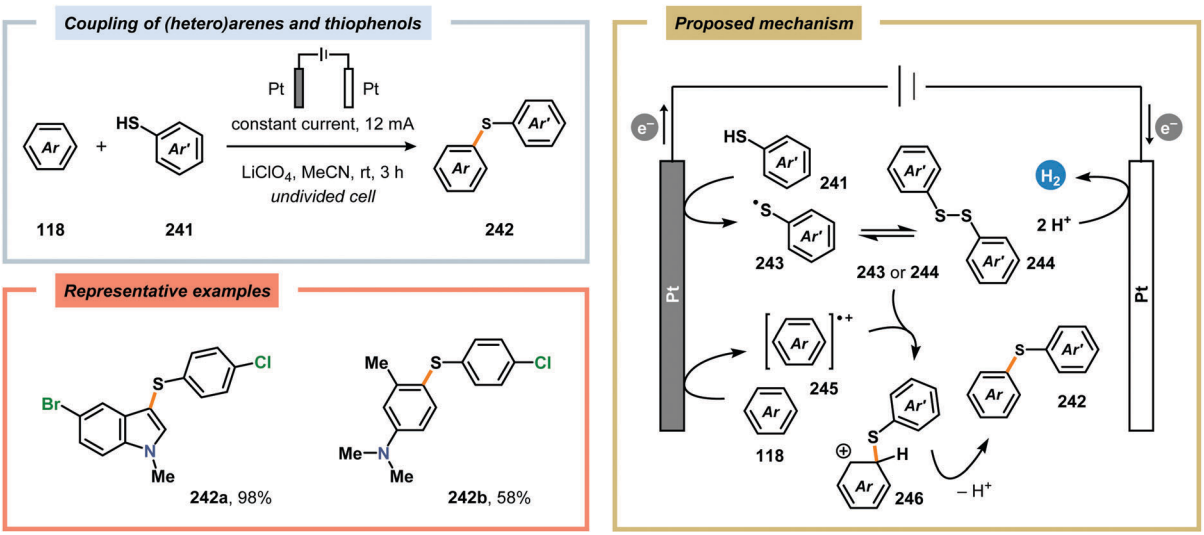


Scheme 77 Cross-coupling of aromatic compounds using the "cation pool" method.

laminar flow also serves as a convenient method for preparing nonsymmetrical biaryl scaffolds as demonstrated by Atobe and coworkers.[606] An electrochemical method for constructing carbon–sulfur (C–S) bonds was recently reported by the Lei group. The electrochemical reactions were conducted in an undivided cell and allowed a wide range of electron-rich arenes and heteroarenes, such as indole, pyrrole and thiophene, to be coupled with various substituted thiophenols. Mechanistic studies indicated that performing the reaction in the presence of the persistent radical TEMPO did not produce any of the desired C–S coupling product. Furthermore, the involvement of disulfides, generated from dimerization of the thiophenols, was also observed. Therefore, two mechanistically distinct pathways could explain the noted reaction outcome. Thus, product formation can occur either through direct radical–radical coupling between the arene cation radical intermediate (**245**) and the thiyl radical (**243**). Alternatively, C–S bond formation can proceed *via* arene radical addition to the produced disulfide (**244**) as depicted in Scheme 78.[607,608]

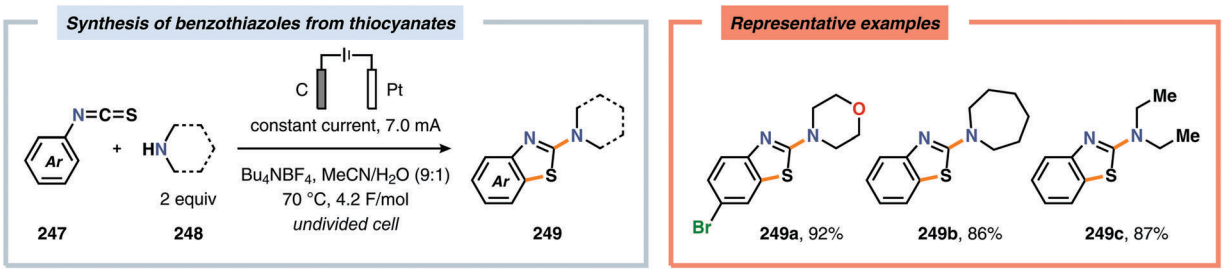
2.6. Oxidative annulations involving (hetero)arenes

Electrochemical approaches for intramolecular C–H thiolations have been reported by the Lei[609] and Xu[610] groups.[611] Lei and coworkers' protocol utilized aryl isothiocyanates, which were

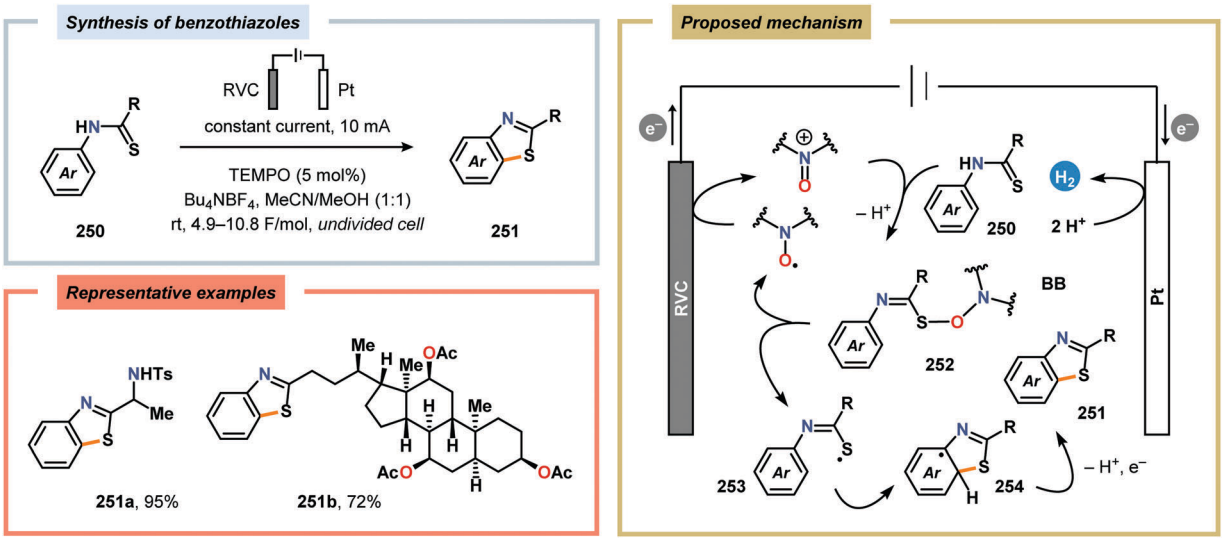


Scheme 78 Direct anodic cross-dehydrogenative coupling of (hetero)arenes and thiophenols.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 79 Electrochemical synthesis of benzothiazoles from aryl isothiocyanates and amines.

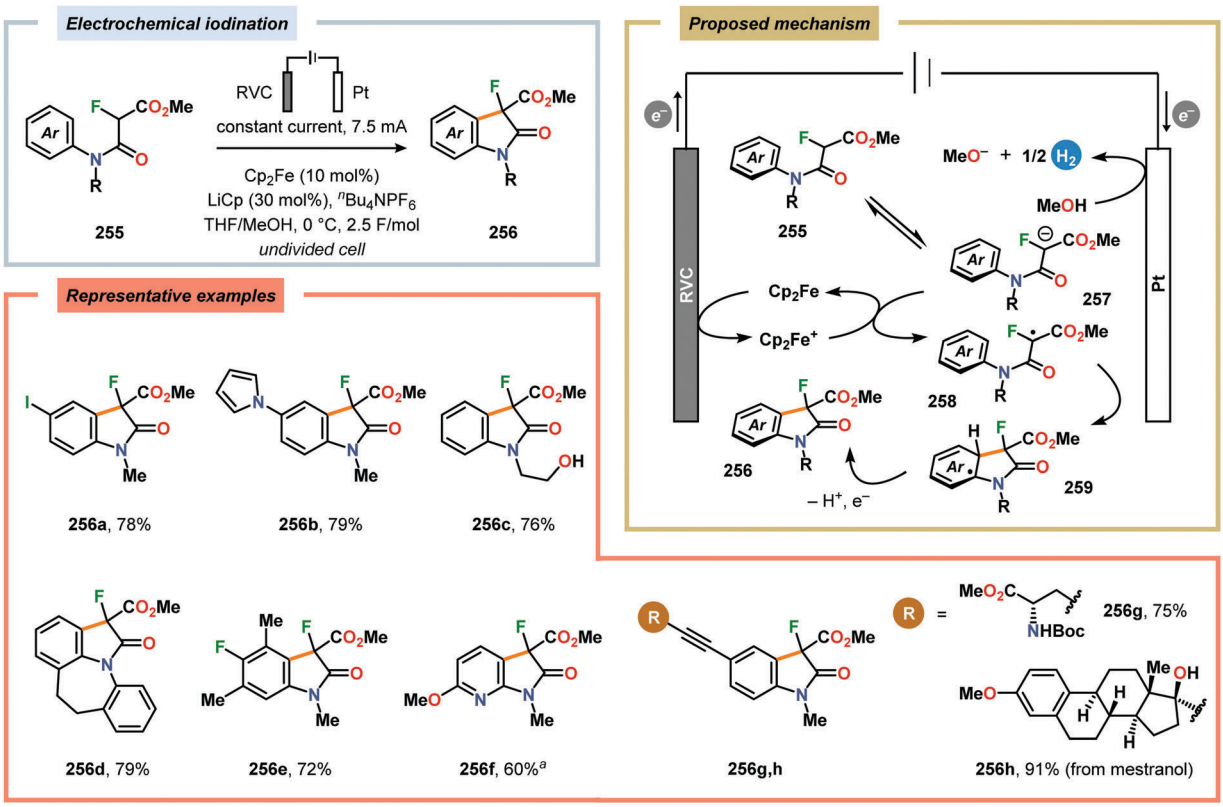


Scheme 80 Electrochemical synthesis of benzothiazoles using TEMPO as redox mediator.

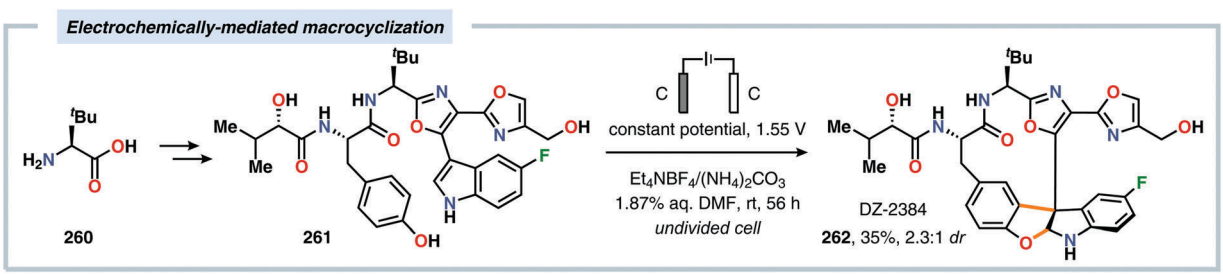
treated with amines to produce the corresponding thioamides. The *in situ* generated thioamides were subsequently electrolyzed at 70 °C in an undivided cell to furnish the desired benzothiazoles (Scheme 79).[609] On the contrary, Xu and coworkers strategy employed TEMPO as a mild redox-mediator to enable selective C–H thiolation of a variety of N-(hetero)arylthioamides with diverse electronic properties (Scheme 80).[610,612] Both C–H thiolation strategies are believed to proceed *via* thioamidyl radicals with concomitant formation of H_2 at the cathode. Expanding on the C–H thiolation strategy, the Xu group has recently demonstrated that 3-fluorooxindoles could be accessed through cross-dehydrogenative coupling.[613,614] Here, ferrocene (Cp_2Fe) was selected as a redox mediator due to earlier literature precedents highlighting its capability of promoting oxidative radical cyclizations.[615–618] The reactions were carried out in an undivided cell and tolerated a multitude of functional groups, including alcohols, alkynes, aminoesters, halogens, silylethers as well as redox-sensitive carbazole, *N*-phenyl carbamate and pyrrole groups (Scheme 81). However, products containing electron-withdrawing groups, such as CF_3, OCF_3 and CO_2Me, were prone to base-induced decomposition and therefore had to be performed at −30 °C for optimal yields. The mechanism is presumed to involve formation of H_2 at the cathode from MeOH, which results in production of MeO^-. The electrochemically generated base facilitates deprotonation of the substrate (**255**) to furnish the corresponding enolate (**257**), which is subsequently oxidized by the anodically generated Cp_2Fe^+ to produce α-carbon radical intermediate **258**. Finally, radical cyclization and rearomatization afford oxindole **256**.[613,619,620]

Harran and coworkers recently showcased electrochemistry as a powerful strategy in organic synthesis, enabling a scalable preparation of DZ-2384 (**262**), a diazonamide-based drug candidate for treatment of cancer.[621] Initial attempts were aimed at using $PhI(OAc)_2$ for mediating the dehydrogenative macrocyclization between the phenol and indole components. However, this approach afforded comparable amounts of a spirocyclohexadienone, which severely limited the yield of the desired macrocyclized product and complicated its purification.[622] To circumvent formation of this byproduct, an alternative method was pursued. In this regard, the use of an electrochemical method was explored as it would employ substrates derived from common amino acids. In the refined route towards DZ-2384 (**262**), the anodic oxidation could be performed in the final step at ambient temperature in aqueous DMF in an undivided cell open to air (Scheme 82), highlighting the practicality of the developed electrochemical macrocyclization approach.[621] A related macrocyclization approach was also explored in the synthesis of azonazine.[623] Subsequently, the Lei group reported a related

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 81 Synthesis of 3-fluorooxindoles though electrochemical cross-dehydrogenative coupling using ferrocene (Cp_2Fe) as redox mediator. [a] Reaction was performed at −30 °C.

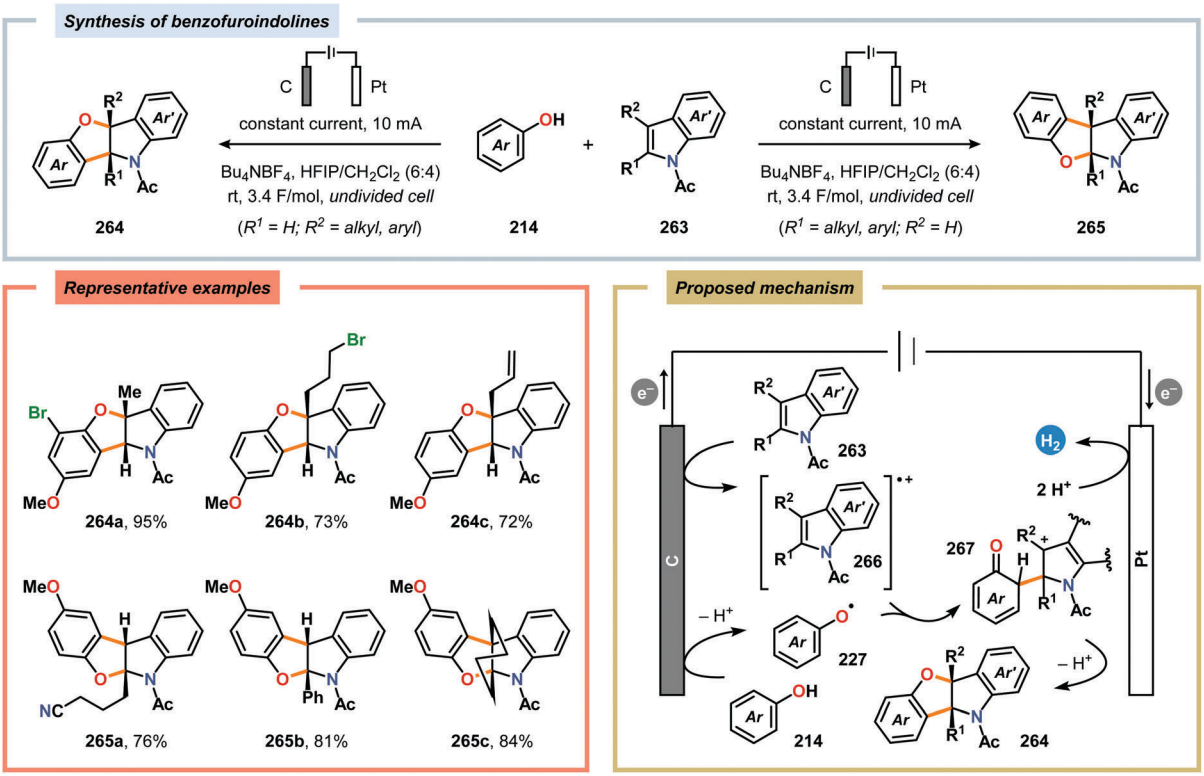


Scheme 82 Preparation of a diazonamide-based drug development candidate by employing a dehydrogenative macrocyclization approach.

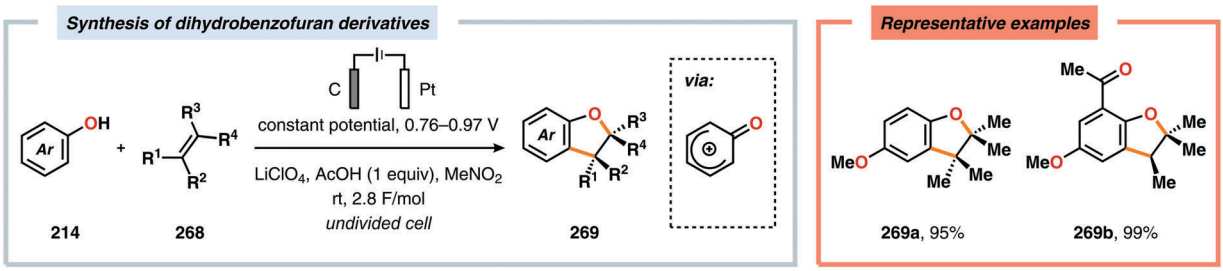
dehydrogenative [3+2] annulation between phenol and indole derivatives for accessing benzofuroindolines using an undivided cell (Scheme 83). Here, various *N*-acetylindoles bearing different C-3 substituents were compatible with the electrochemical annulation and afforded the corresponding benzofuro[3,2-*b*]-indolines (**264**) in high to excellent yields. However, applying 2-substituted *N*-acetylindoles resulted in a switch in selectivity and instead produced the corresponding benzofuro[2,3-*b*]indoline core (**265**). CV experiments of *p*-methoxyphenol and various *N*-acetylindoles revealed that the oxidation potentials were quite similar, suggesting that anodic oxidation of both coupling components might occur under the electrolytic conditions. Radical coupling between the indolyl cation radical (**266**) and phenoxyl radical (**227**) furnishes cation **267**. Finally, intramolecular cyclization and deprotonation gives the target benzofuroindoline with concomitant formation of H_2 at the cathode. However, a mechanism involving nucleophilic attack of the phenol moiety onto the indolyl cation radical cannot be excluded.[624]

Anodic oxidation of phenols can result in generation of phenoxonium ions upon removal of two electrons and a proton, which can react with simple nucleophiles, such as MeOH or H_2O, to afford 4-substituted cyclohexa-2,5-dienones.[625–630] Chiba and coworkers highlighted that the anodically generated phenoxonium ions could engage in intermolecular [3+2] cycloaddition with unactivated alkenes to afford dihydrobenzofuran derivatives (Scheme 84).[631–635] The reaction has also been conducted using a temperature-controlled multiphase solution consisting of cyclohexane and $LiClO_4/MeNO_2$,[636] which was

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 83 Electrochemical synthesis of benzofuro[2,3-*b*]indolines and benzofuro[3,2-*b*]indolines.

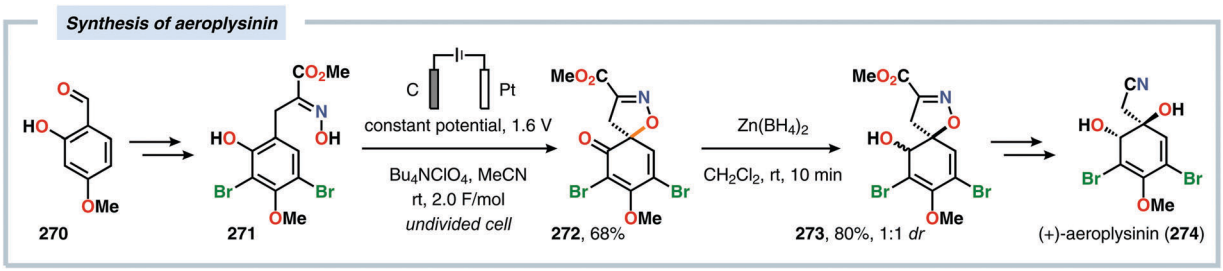


Scheme 84 Synthesis of dihydrobenzofuran derivatives *via* anodic intermolecular cycloaddition of phenols and unactivated alkenes.

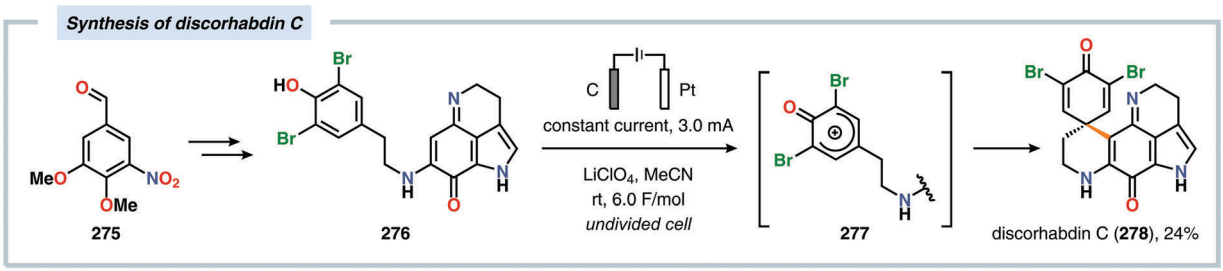
shown to significantly enhance the interaction between the hydrophobic aliphatic alkenes and the polar unstable cation intermediates, thereby accelerating the desired [3+2] cycloaddition while avoiding overoxidation of the products.[637] Some of these dihydrobenzofuran derivatives display fluorescent properties,[638,639] a feature which has been exploited for fluorescent labeling of amino acid derivatives[640] as well as homonucleosides.[641] In addition, the electrochemically produced cyclohexadienone derivatives have also been reported to engage in [4+2][642–648] and [5+2][649–652] cycloadditions.[653] Alternatively, incorporation of a pending nucleophile into the phenol unit allows the design of various types of intramolecular reaction manifolds for construction of spirodienone derivatives.[654–661] This feature has been leveraged in the synthesis of aeroplysinin (Scheme 85),[662–665] discorhabdin C (Scheme 86),[666–669] gymnastatin A (Scheme 87)[670] as well as heliannuol E (Scheme 88)[671–673] and derivatives thereof.[674]

Catechols and related derivatives constitute another class of compounds that can undergo two-electron anodic oxidation to afford, for example, 1,2-benzoquinones.[675–681] Considering their relatively low oxidation potentials, it is perhaps unsurprising that an assortment of nucleophiles are compatible with these processes without being subjected to oxidation. Here, amines,[682–689] azide,[690,691] enolates and other carbon nucleophiles,[692–711] nitrate,[712] sulfinates[713–727] as well as thiols[728–744] have been shown to engage in Michael additions with the anodically generated (imino)quinones. A variety of transformations for delivering annulated scaffolds have also been developed (Scheme 89). In the pursuit of this goal, a range of different benzofurans (**291** and **293**) have been accessed using 1,3-diketones,[745–756] α-cyanoketones[757] or chloranilic acid[758] as

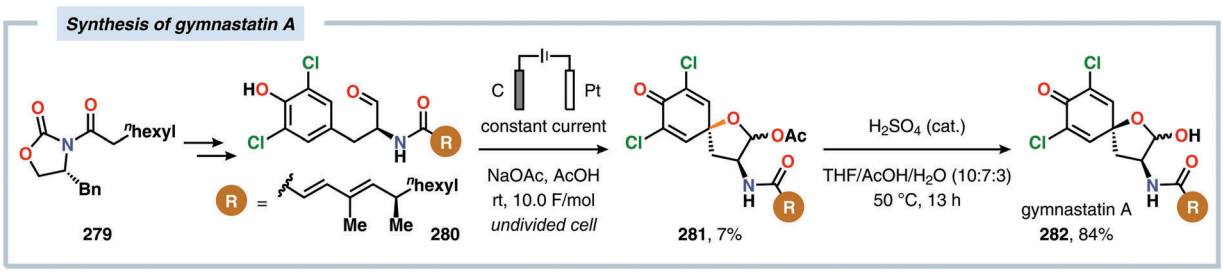
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



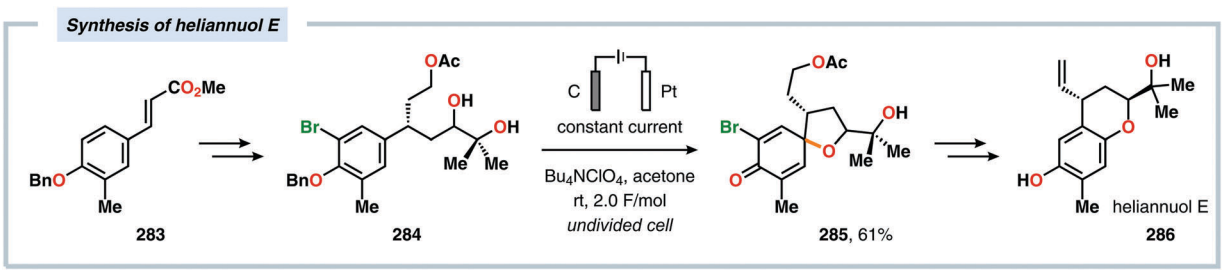
Scheme 85 Utilization of anodically generated phenoxonium ions in the synthesis of aeroplysinin.



Scheme 86 Synthesis of discorhabdin C using a phenoxonium ion-based strategy.



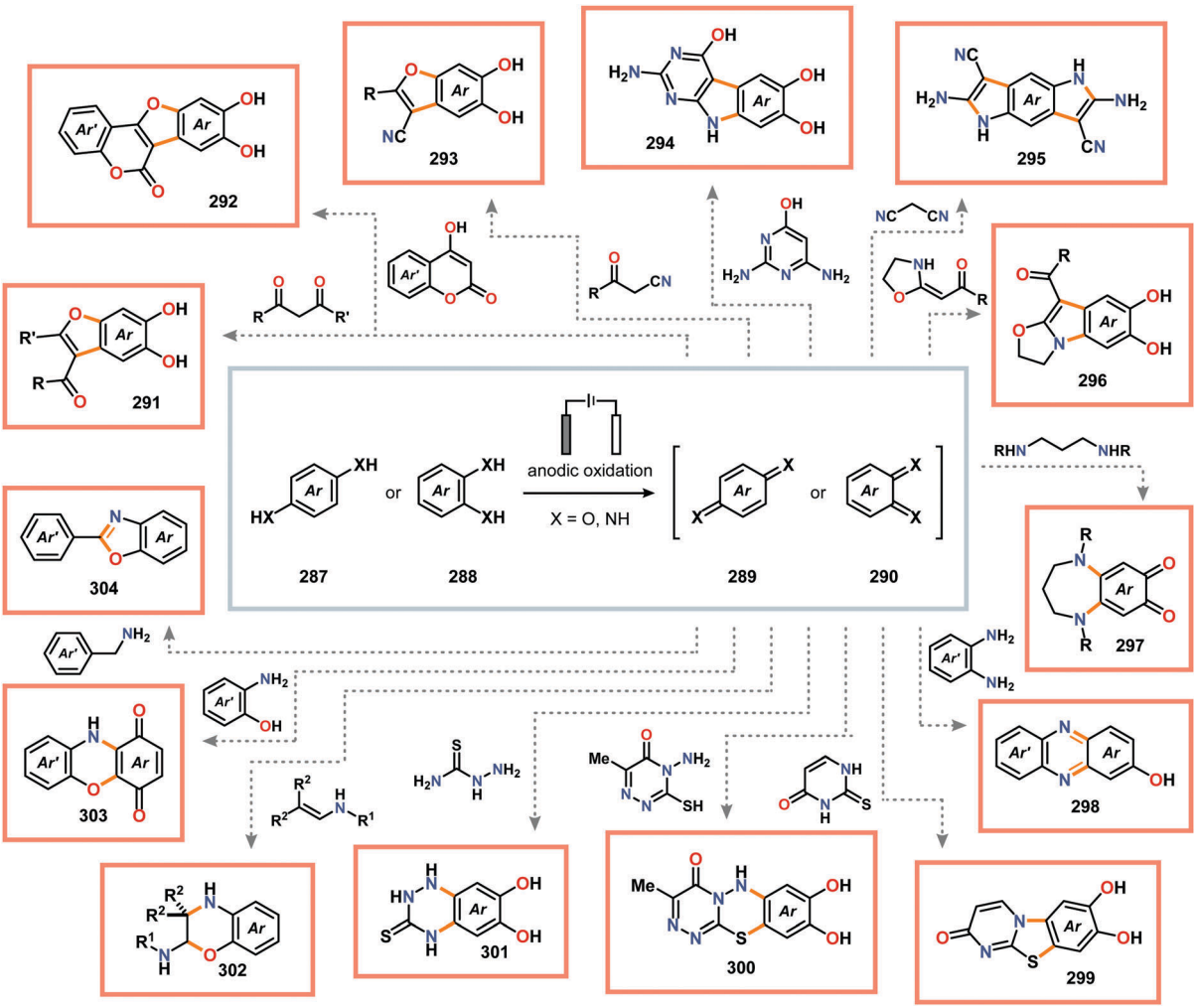
Scheme 87 Electrochemically-mediated spiro hemiacetal formation in the synthesis of gymnastatin A.



Scheme 88 Electrochemical production of a spirodienone in the synthesis of heliannuol E.

the nucleophile. Additional scaffolds that have been synthesized are benzo[*e*][1,2,4]triazino[3,4-*b*][1,3,4]thiadiazine (**300**),[759] benzoxazine (**302**),[760,761] benzoxazole (**304**),[762,763] coumestan (**292**),[764,765] dihydrobenzo[*e*][1,2,4]triazine-3(2*H*)-thione (**301**),[766] indole (**294–296**),[767–769] phenazine (**298**),[770,771] phenoxazine (**303**)[772] tetrahydro-1*H*-benzo[*b*][1,4]diazepine (**297**)[773] and thia-1,4*a*-diaza-fluorenone (**299**)[774] derivatives. Furthermore, the electrochemically generated benzoquinone derivatives can also engage in [4+2] cycloaddition in the presence of cyclopentadiene,[775,776] produce trimerization products[777–779] or engage in transfer hydrogenation catalysis.[780–782] A dual electro- and organocatalysis strategy was outlined by Jørgensen and coworkers, which enabled the

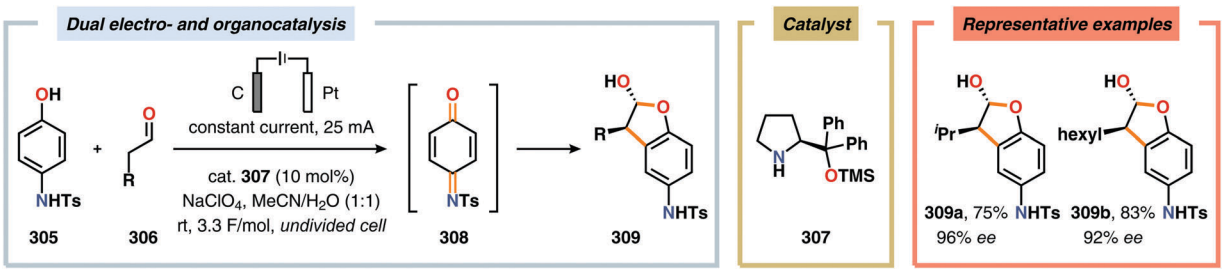
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 89 Annulation of electrochemically generated benzoquinones and related derivatives.

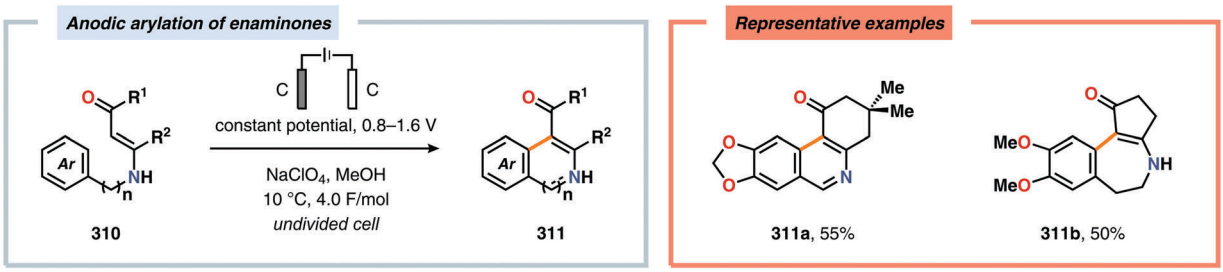
enantioselective intermolecular α-arylation of aldehydes.[783,784] Here, the combination of anodic oxidation and asymmetric enamine catalysis provided access to *meta*-alkylated anilines in good yields and excellent enantiomeric excess (Scheme 90).[783]

Several research groups have targeted the development of oxidative annulation protocols involving alkenes, and enol ethers or derivatives thereof. Based on earlier work that showed that enaminones[785] can be anodically dimerized to yield 3,4-diketopyrroles,[786,787] Schäfer and Eilenberg reported that *N*-benzyl- and β-phenethyl enaminones undergo intramolecular arylation to produce isoquinolines and benzazepines upon anodic oxidation using an undivided cell (Scheme 91).[788] Lei and coworkers explored a related strategy for synthesis of indoles **313**. The protocol relied on intramolecular dehydrogenative annulation of *N*-aryl enamines **312** and utilized iodide as a redox mediator.[789] Mechanistic insight revealed that carrying out the annulation reaction in the absence of electricity but using *N*-iodosuccinimide (NIS) as an oxidant resulted in a high yield of

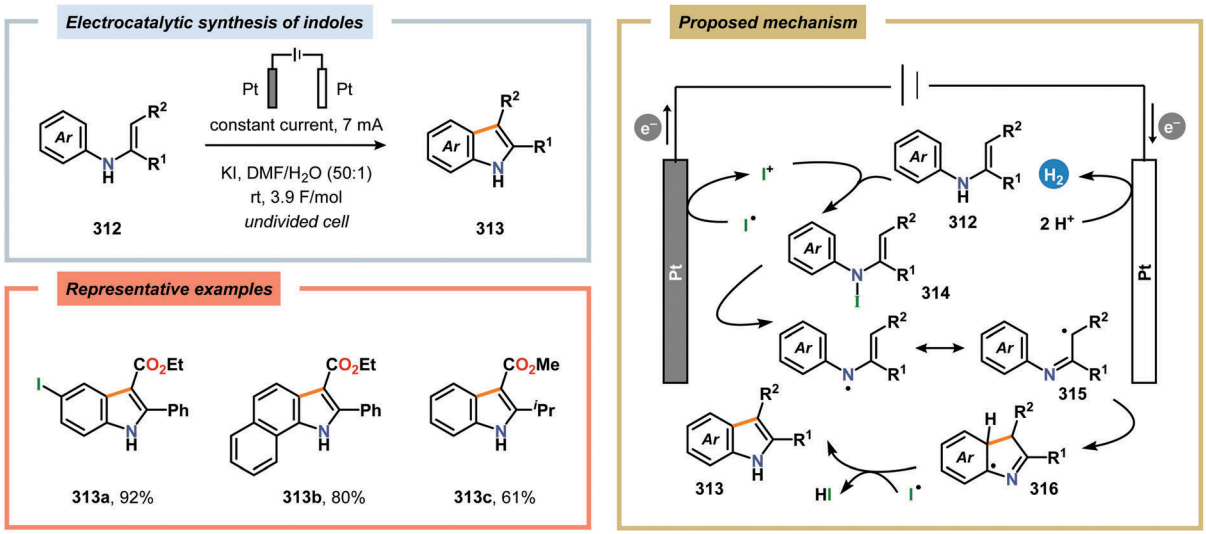


Scheme 90 α-Arylation of aldehydes through the merger of electrochemical oxidation and organocatalysis.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 91 Anodic intramolecular arylation of enaminones.

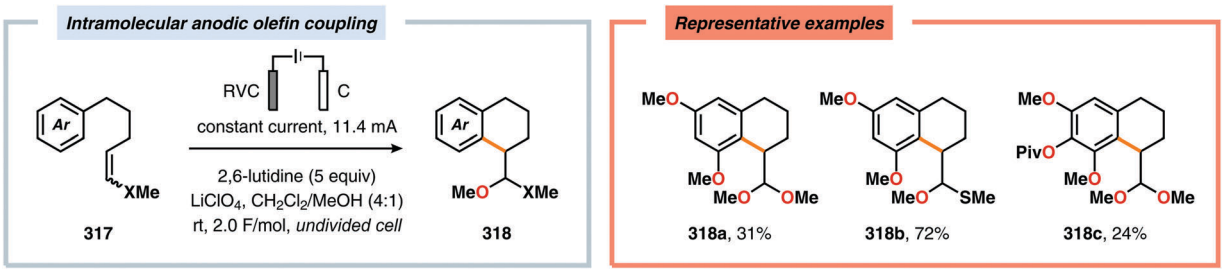


Scheme 92 Electrocatalytic synthesis of indoles using iodide as a redox mediator.

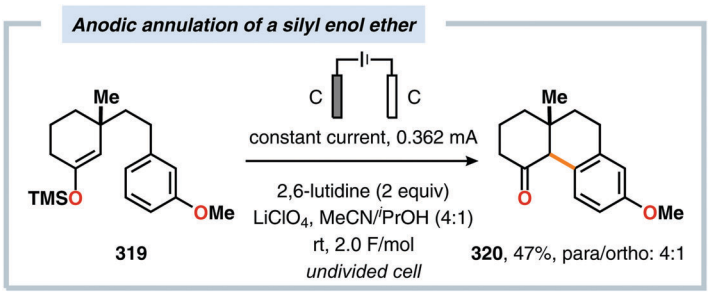
the desired indole. Furthermore, radical inhibition experiments with TEMPO did not produce any product. Based on these results and previous observations,[790] a mechanism involving a hyperiodide intermediate (I^+) generated *via* anodic oxidation was proposed. The *in situ* produced I^+ subsequently reacts with the *N*-aryl enamine to furnish an *N*-iodo intermediate (**314**), which upon N–I bond homolysis provides radical intermediate **315**. Finally, intramolecular radical addition, oxidation and deprotonation afford the target product (Scheme 92).[789,791,792]

Pioneering studies by the Moeller group have demonstrated that alkenes, such as enol ethers, and electron-rich (hetero)-arenes can be oxidatively coupled under electrochemical conditions. In these transformations, an alkene that is normally nucleophilic is initially oxidized at the anode to form a radical cation intermediate that is subsequently trapped by a second nucleophile to deliver anti-Markovnikov addition products (*vide infra*). Here, electron transfer confers umpolung reactivity of a functional group, thus enabling alternative retrosynthetic disconnections of complex structures through previously challenging bond disconnections.[63,81] The Moeller[793–795] and Wright[796] groups have exploited the intramolecular anodic coupling of enol ethers or vinyl sulfides and electron-rich aromatic systems to deliver the fused, bicyclic or tricyclic ring skeletons in moderate to high yields (Schemes 93 and 94). A related approach was also utilized by the Wright group for the synthesis of the hamigeran skeleton (Scheme 95).[797] Heteroaromatic rings are also compatible coupling partners in these intramolecular anodic olefin coupling reactions. Here, different tethers can be employed to afford, for example, six- (Scheme 96)[794,796,798,799] or seven-membered[800] (Scheme 97) annulated furan systems, which can be readily transformed into various carbo- and oxacyclic motifs.[801–807] These electrochemical annulations are presumed to involve initial oxidation of the silyl enol ether moiety rather than the furyl group.[800] Additional heterocycles that have been utilized consist of imidazoles,[81] pyrroles[794] as well as thiophenes.[808–810] The applicability of the anodic olefin coupling reaction between silyl enol ethers and furans has also been exploited in syntheses of natural products. Moeller and coworkers employed the methodology as the key step in their synthesis of alliacol A[65,66] (Scheme 98) while an anodic coupling strategy enabled the construction of the central seven-membered ring in Trauner's synthesis of guanacastepene E (Scheme 99).[67,68] Furthermore, the arteannuin ring skeleton has been assembled through the use of a pair of anodic coupling reactions (Scheme 100). Here, both coupling reactions exploited the furan moiety as the coupling partner in which the first cyclization made use of a chiral *N*,*O*-ketene acetal initiating group to provide the desired bicyclic product in high yield and the second

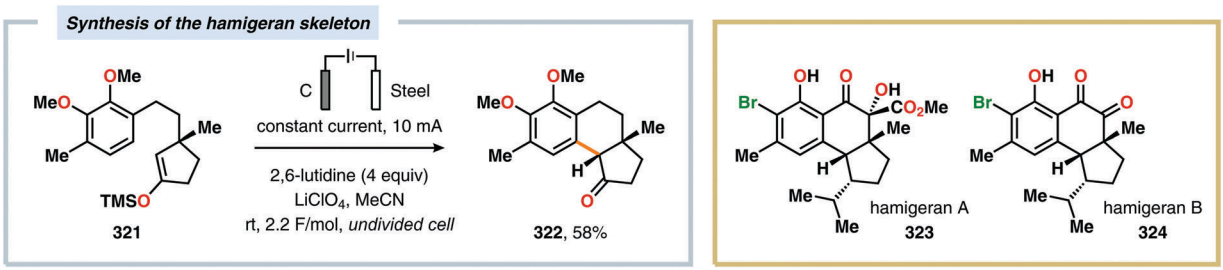
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



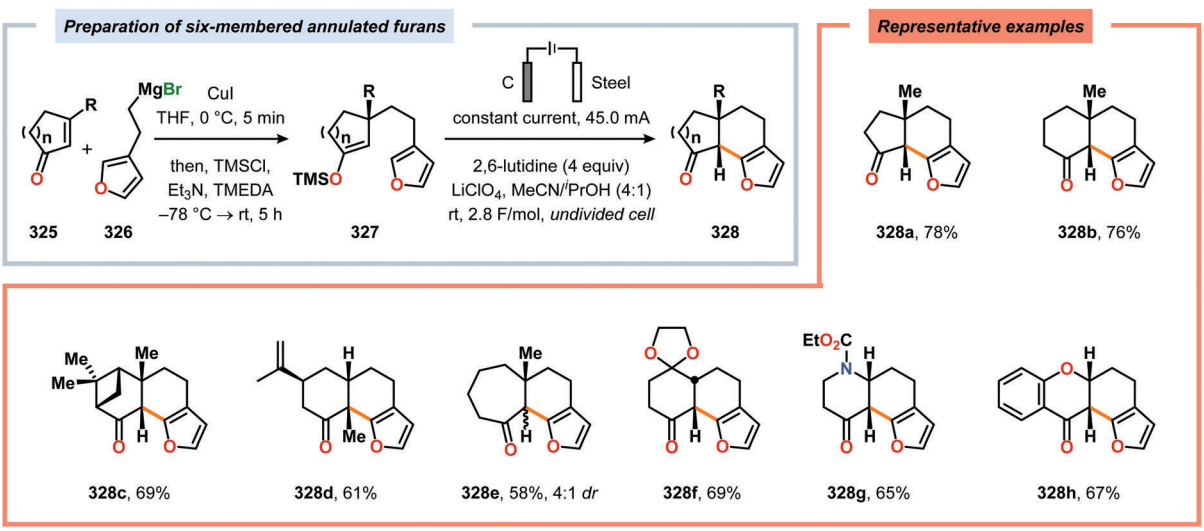
Scheme 93 Intramolecular anodic olefin coupling.



Scheme 94 Anodic annulation of a silyl enol ether and an electron-rich arene.

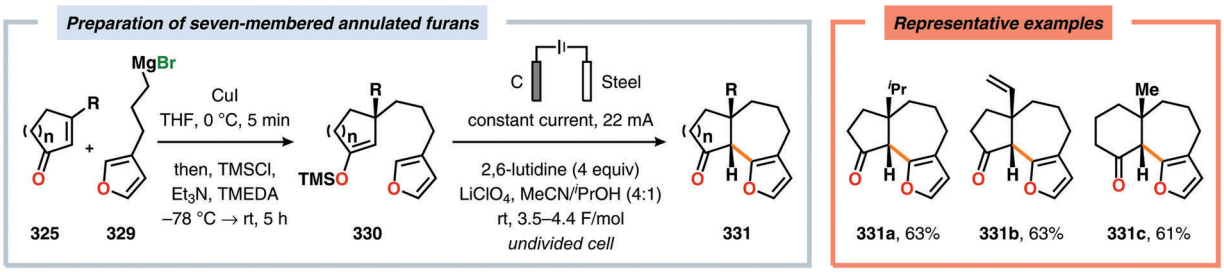


Scheme 95 Synthesis of the hamigeran skeleton through anodic coupling with an electron-rich arene.

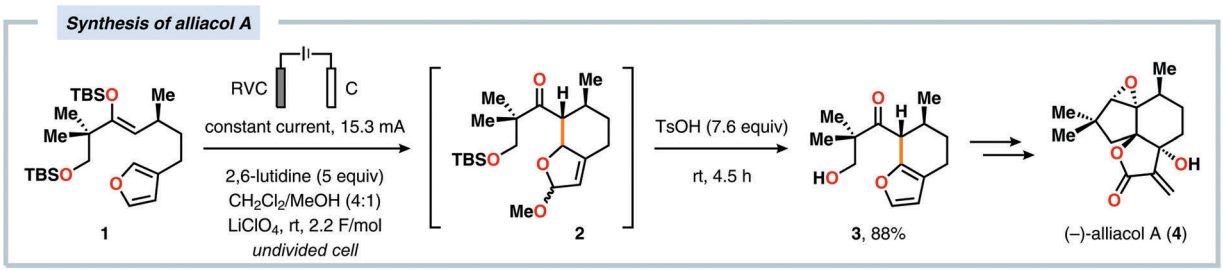


Scheme 96 Electrochemical coupling of silyl enol ethers and furans for synthesis of six-membered annulated furans. Yields are reported for two steps based on the starting enone as the limiting agent.

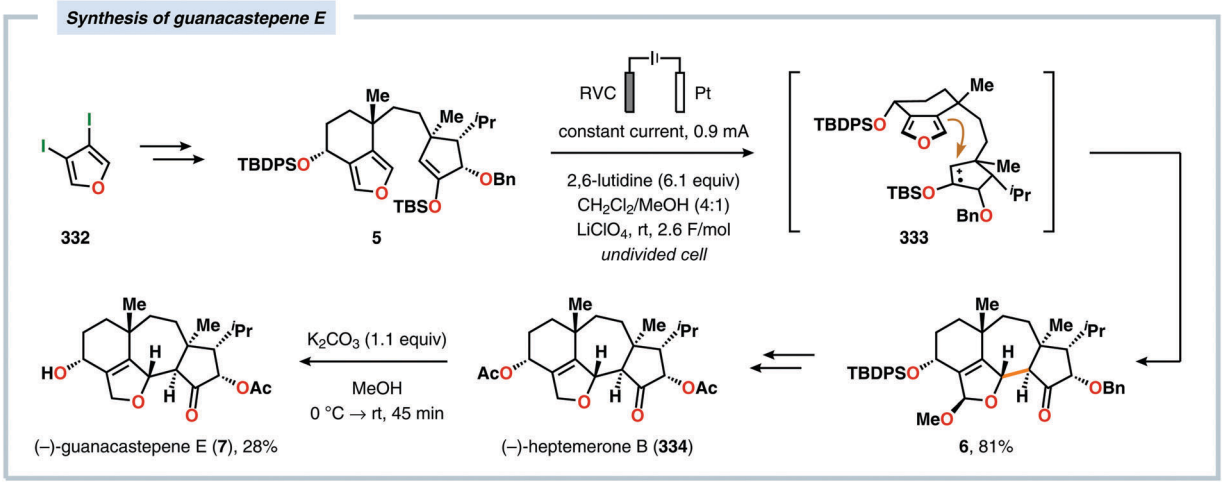
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 97 Anodic coupling for synthesis of seven-membered annulated furans. Yields are reported for two steps based on the starting enone as the limiting agent.



Scheme 98 Application of the anodic olefin coupling in the synthesis of (−)-alliacol A.



Scheme 99 Utilization of the anodic olefin coupling in the synthesis of (−)-guanacastepene E.

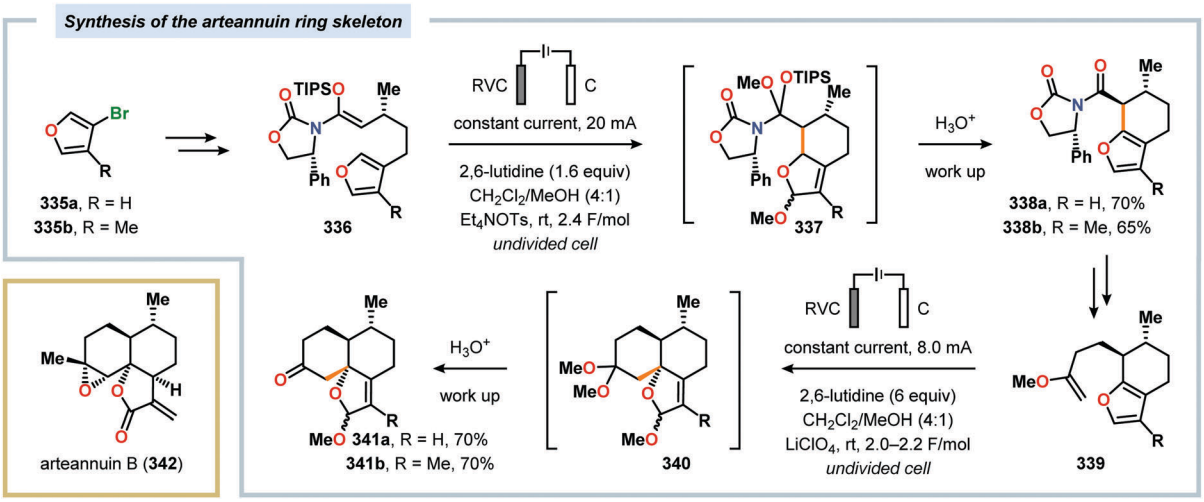
anodic coupling afforded the tricyclic core while generating the required quaternary stereocenter.[811]

Enol ether radical cations are also prone to undergo intermolecular [2+2] and [4+2] cycloaddition with another alkene or diene. Upon intermolecular C–C bond formation, the inclusion of an electron-rich arene ring in either the enol ether or alkene moiety facilitates intramolecular electron transfer from the aryl ring to form the final cycloadduct.[812–822] The arene radical cation subsequently generates a second equivalent of the enol ether radical cation through intermolecular electron transfer.[823] A characteristic feature of these "redox-tag"[824] mediated cycloaddition reactions is that only small amounts of electricity are needed to initiate the radical chain process.[825]

3. Electrochemical approaches to carbon–nitrogen (C–N) bond formation

Nitrogen-containing compounds constitute essential motifs and are present in natural products, pharmaceutical agents, agrochemicals and material science. Chemists have actively

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 100 Sequential anodic couplings for synthesis of the arteannuin ring skeleton.

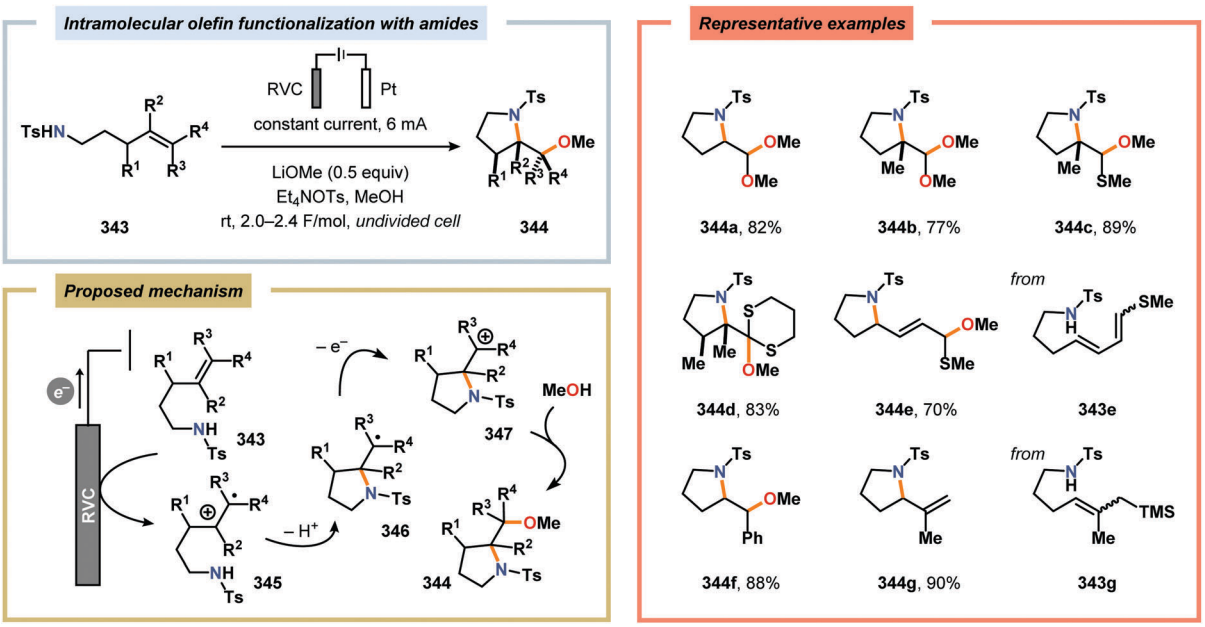
targeted the development of facile and selective C–N bond formation platforms. However, despite the recent advances in C–N cross-coupling reactions, these approaches normally require elevated temperatures, prefunctionalized starting materials, such as aryl halides or pseudohalides, stoichiometric oxidants or the use of expensive catalysts. In this context, methods for direct C–H amination have emerged as efficient, step- and atom-economical alternatives, and have expanded the repertoire of reaction manifolds.[826,827] In the following sections, the development of electrochemical methodologies for construction of C–N bonds and nitrogen-containing motifs is examined.

3.1. Amination of alkenes and alkynes

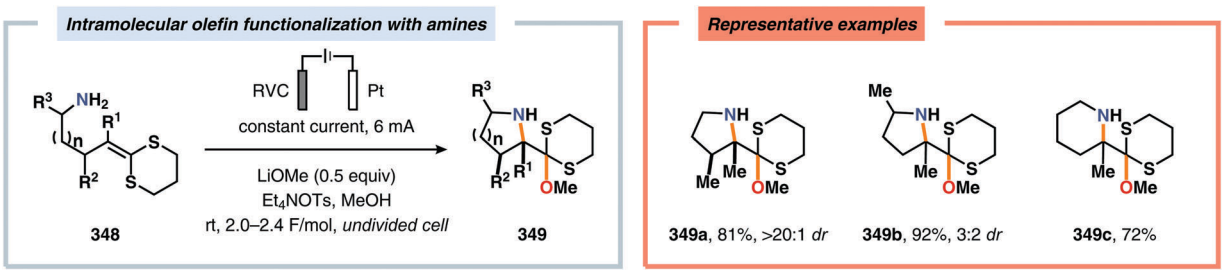
Considerable efforts have been devoted to the development of methods that facilitate the addition of, for example, N–H (hydroamination) and arene C–H bonds (hydroarylation) across alkenes and alkynes. Here, atom-economical and regioselective platforms for direct functionalization of double or triple bonds allow access to a range of functionalized products, thereby offering considerable opportunities for strategic chemical synthesis.[828–848]

In addition to the intramolecular anodic olefin coupling reactions discussed in the previous section, other carbon-[849–862] as well as oxygen-based[863–875] nucleophiles have also been utilized in radical cation-initiated cyclization protocols for constructing C–C and C–O bonds, respectively. Furthermore, Moeller and coworkers have designed systems for direct functionalization of alkenes using nitrogen functionalities as trapping groups. It was demonstrated that the nature of the substituents on the generated radical cation intermediate had a profound influence on its ability to react with various trapping groups.[876] Thus, more polarized radical cations tend to favor C–C bond forming reactions, while less polarized radical cations are beneficial when using heteroatomic trapping groups. Employing more basic reaction conditions also had a dramatic influence on the cyclization reactions. Here, the use of a strong base, such as LiOMe, was considered to increase the nucleophilicity of the nitrogen trapping group by facilitating formation of the sulfonamide anion (Scheme 101). Although five-membered ring systems were accessible in a relatively straightforward fashion, coupling reactions leading to six-membered rings presented problems due to competing elimination of a proton from the carbon-center next to the radical cation intermediate.[877,878] Ensuing mechanistic studies aimed at examining the relative reactivity of olefin-based radical cations toward various intramolecular nucleophiles. To address this, an electron-rich olefin tethered to two different nucleophiles/trapping groups, an alcohol and a sulfonamide, was exploited. Trapping of the formed radical cation with the alcohol moiety was shown to be facile and reversible, affording the kinetically favored product. In contrast, sulfonamide cyclization was demonstrated to furnish the thermodynamically favored product and can be enhanced with nonpolar reaction conditions, elevated temperatures, and low current. In these reactions, the normally nucleophilic olefin functionality is initially oxidized to a radical cation intermediate. This intermediate is subsequently trapped by the sulfonamide anion to form the desired C–N bond. However, a mechanistic portrayal involving initial anodic oxidation of the sulfonamide anion to produce a nitrogen-centered radical that is subsequently reduced through intramolecular electron transfer from the olefin unit to afford an olefinic radical cation cannot be ruled out for certain substrates.[879–881] The Moeller group has also employed photovoltaic cells in which sunlight is harnessed as the power source for driving anodic olefin cyclization reactions.[882] Following on from these reports, the Moeller group disclosed a potentially more practical approach that took advantage of unprotected amines as the trapping group (Scheme 102). Although the method employed a dithioketene acetal group as the electron-rich olefin, the oxidation potential of the secondary amine moiety in the product is lower than that of either functional group in the starting material, suggesting that overoxidation of the product should be of concern. However, this reasoning does not take into consideration that rapid cyclization reactions can lower the oxidation potential of the substrate. Hence, in the developed protocol the intramolecular cyclization is so rapid that it causes a 460 mV drop in the oxidation potential of substrate **348**.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 101 Intramolecular anodic olefin coupling reactions with sulfonamides.

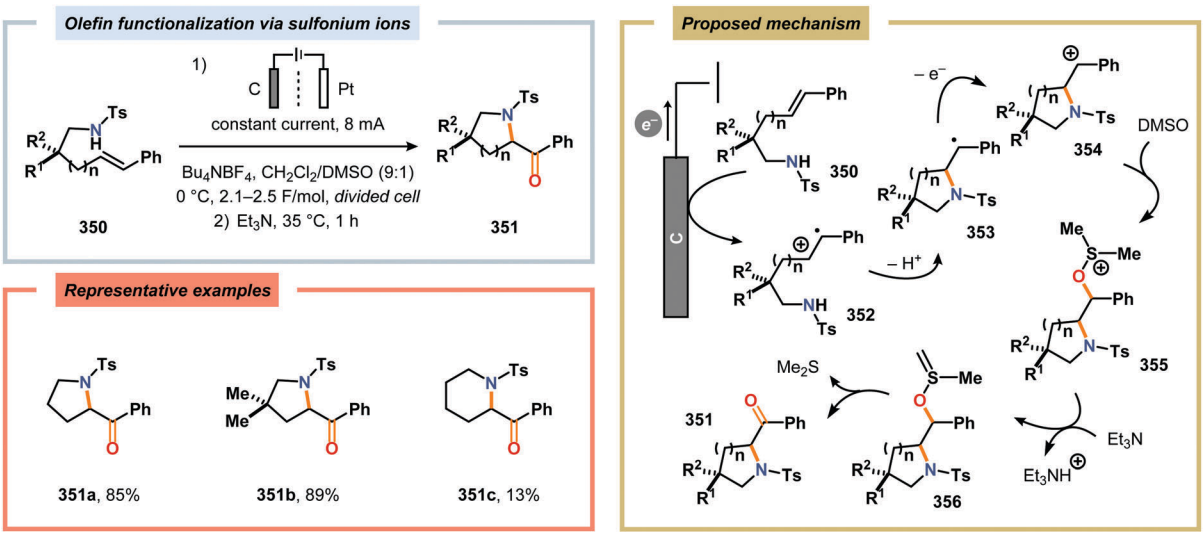


Scheme 102 Anodic coupling of amines and dithioketene acetals.

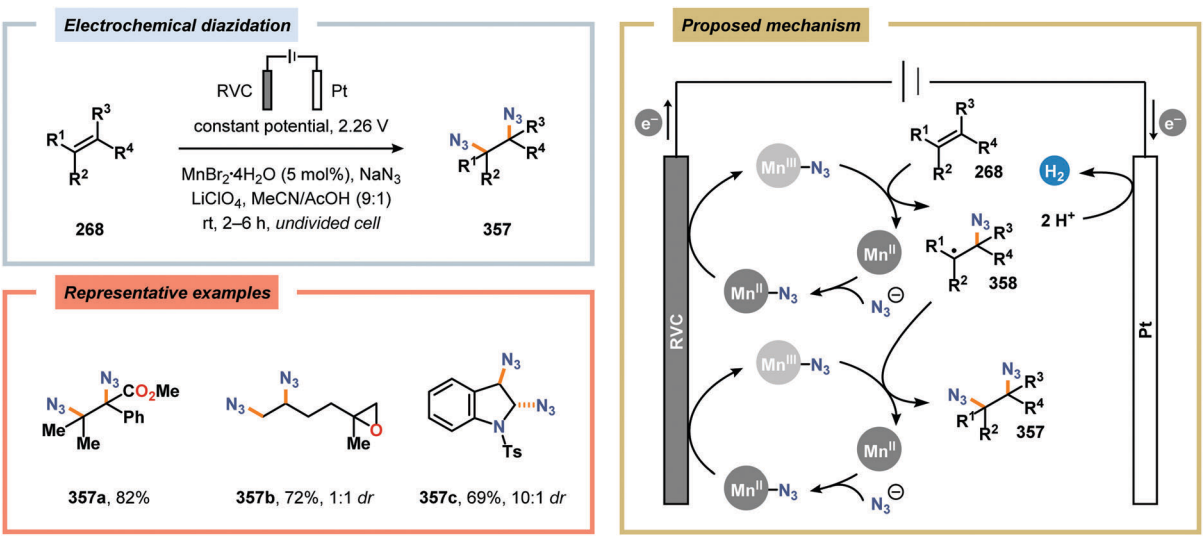
This effect is so dramatic that the substrate potential is now significantly lower than that of the product, highlighting that a simple analysis of oxidation potentials for isolated functional groups can be deceiving when attempting to predict the success of an oxidative cyclization reaction.[883] Yoshida and coworkers have demonstrated that anodic cyclization and chemical oxidation manifolds can be successfully integrated in a one-pot, sequential manner (Scheme 103). Here, anodic oxidation of the olefinic bond is followed by intramolecular cyclization by the tethered nucleophilic moiety to ultimately afford a carbocation **354** upon a subsequent one-electron oxidation. In the presence of DMSO as an external nucleophile, efficient trapping of intermediate **354** occurs to provide the corresponding alkoxysulfonium ion **355**. This intermediate is sufficiently stable at low temperatures and subsequent treatment with Et_3N gives a sulfur ylide (**356**) that can undergo an intramolecular proton transfer event, resulting in elimination of dimethylsulfide and the exocyclic ketone **351**.[467,884]

Recently, the Lin laboratory presented an electrochemical protocol for the diazidation of alkenes. The reported operationally simple and environmentally friendly method converted alkenes and NaN_3 to 1,2-diazides (Scheme 104). The merger of electrochemical oxidation and an earth-abundant manganese catalyst enabled the transformation to proceed under mild reaction conditions. The system exhibited a broad substrate scope and high functional group compatibility. For example, substituents typically susceptible to nucleophilic displacement by N_3^-, such as epoxides, esters, alkyl halides, were tolerated by the metal-catalyzed electrochemical diazidation strategy. Furthermore, the authors showed that several 1,2-diazides, including those that contain reductively labile groups, could be chemoselectively converted to the corresponding 1,2-diamines. This can be performed consecutively and circumvents elaborate isolation of any intermediates, providing a general and operationally simple alternative for synthesis of vicinal diamine. Finally, mechanistic insight through radical clock experiments confirmed the intermediacy of radical adduct **358**.[885–888] Based on this platform, the Lin group subsequently reported a manganese-catalyzed electrochemical protocol for dichlorination[889] as well as chlorotrifluoromethylation[890] of alkenes.[891]

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 103 Integrated electrooxidative cyclization followed by chemical oxidation of alkenes *via* alkoxysulfonium ions.



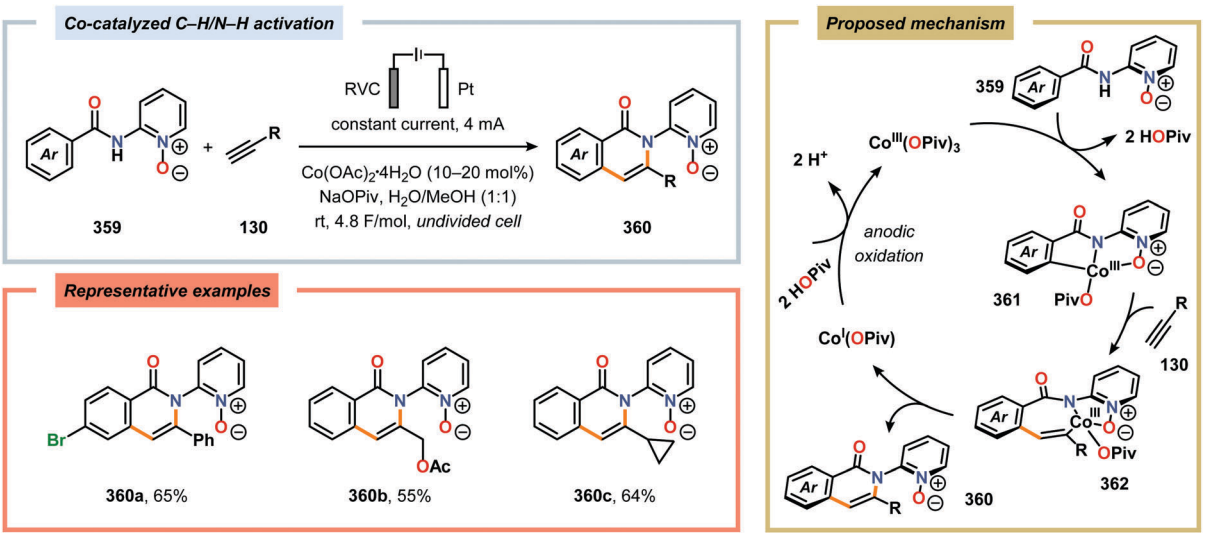
Scheme 104 Metal-catalyzed electrochemical diazidation of alkenes.

Alkyne annulations are typically limited to high reaction temperatures and/or the use of stoichiometric amounts of toxic metals as sacrificial oxidants. Recently, the Ackermann[892] and Lei[893] groups independently reported methods for cobalt-catalyzed electrochemical C–H/N–H annulation of alkynes, such as ethyne, as well as ethylene (Schemes 105 and 106). The cobalt-catalyzed electrochemical strategies allowed for efficient C–H functionalization of *ortho*-, *meta*- and *para*-substituted amides, and aryl halides while obviating undesired coupling reactions. Subsequently, the two groups independently disclosed methods for electrochemical cobalt-catalyzed C–H amination of arenes with alkylamines.[894,895] Furthermore, a ruthenium-catalyzed electrochemical dehydrogenative annulation reaction of aniline derivatives and alkynes for the synthesis of indoles was recently described.[896] Here, the electric current was proposed to promote reoxidation of the active ruthenium-based catalyst and facilitate H_2 evolution.

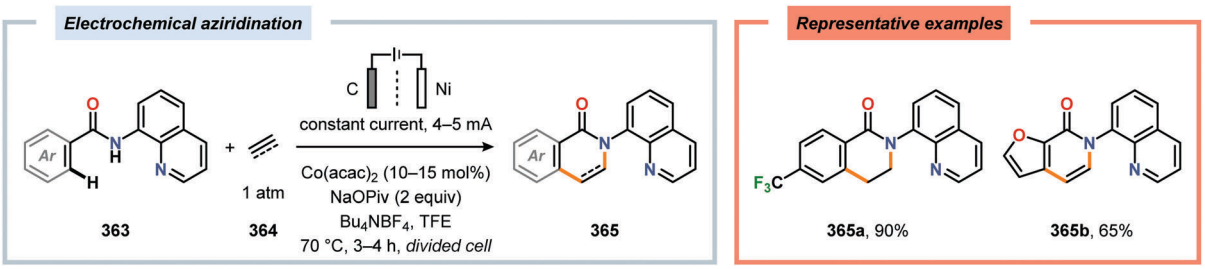
3.2. Electrochemical generation of nitrogen-centered radicals

The term nitrogen-centered radical refers to species where the generated radical is localized on a nitrogen atom. In contrast to carbon-centered radicals, nitrogen-centered radicals have historically been relatively underutilized from a synthetic viewpoint. This was mainly due to a lack of mild and reliable methods for generating nitrogen-centered radicals, which constrained their applicability in academic and industrial settings.[897–900] However, strategies based on nitrogen-centered radicals have recently been shown to expedite C–N bond construction and have therefore received considerable attention from the scientific community. Nitrogen-centered radicals have, for example, been demonstrated to add to alkenes, alkynes, dienes

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 105 Cobalt-catalyzed electrochemical C–H/N–H annulation of alkynes.



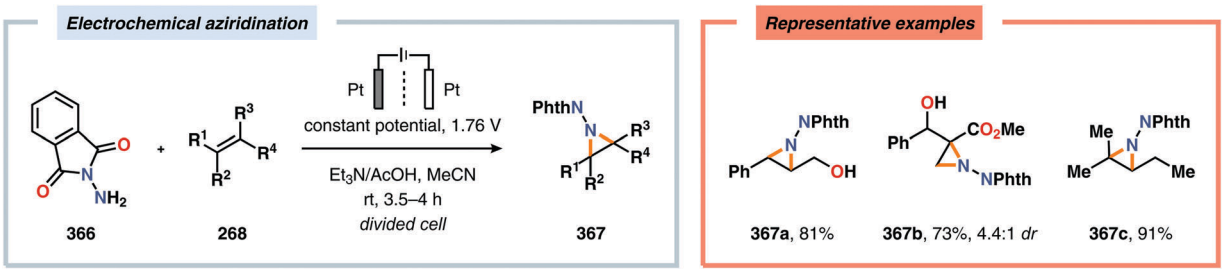
Scheme 106 Cobalt-catalyzed electrochemical C–H/N–H annulation of ethylene or ethyne.

and engage in C–H functionalization, making them versatile intermediates in contemporary organic synthesis.[13m,901–908]

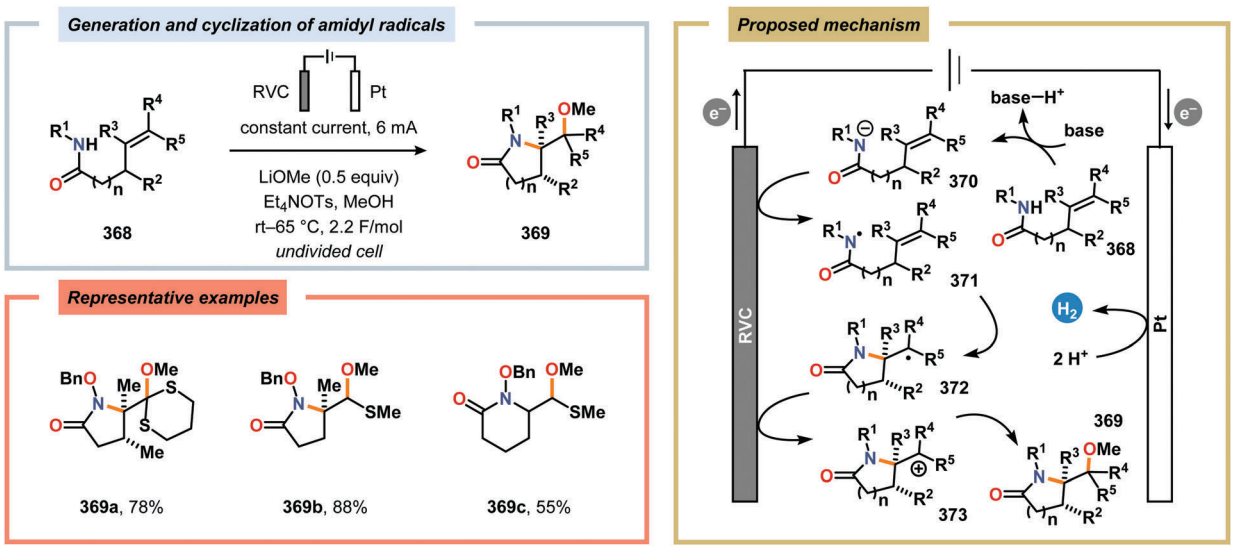
Early work from the Yudin group demonstrated that electrochemical aziridination of olefins could be achieved in the presence of *N*-aminophthalimide (**366**), a reaction that typically is hampered by the use of stoichiometric amounts of toxic Pb^{IV} oxidants. In this reaction, nitrene-transfer from *N*-aminophthalimide to olefins occurs under mild conditions on platinum electrodes (Scheme 107). Using the developed approach, both electron-rich and electron-poor olefins were effectively converted to aziridines in good to excellent yields. Although some olefins displayed similar oxidation potentials as the nitrene-transfer reagent, the crucial factor that accounts for the high levels of chemoselectivity in the transformations is the overpotential. It was noted that the nature of the electrode material was found to be critical for the reaction to proceed. For example, replacing the platinum electrodes with carbon-based ones resulted in termination of the nitrene-transfer reactions. Thus, under particular conditions various substrates possess different overpotentials depending on, for example, the electrode material. This highlights that reactive species can be selectively generated by maximizing the difference in overpotentials between the substrate and the reagent, thereby avoiding detrimental background reactions.[909–911] Likewise, under similar reaction conditions a variety of sulfoxides could be chemoselectively converted into the corresponding sulfoximines.[912]

The Moeller group developed a protocol for synthesis of γ- and δ-lactams using *O*-benzyl hydroxamates or *N*-phenyl amides.[913] Here, the anodically generated nitrogen-centered amidyl radicals **371** underwent cyclization reactions with electron-rich olefins to form carbon-centered radicals **372** that are subsequently oxidized to furnish carbocations **373**. These carbocations can then be trapped with nucleophiles, such as MeOH, to provide the target lactams **369** (Scheme 108). In general, ketene dithioacetal motifs were found to be the most effective coupling partners for the developed cyclization strategy. While the *O*-benzyl hydroxamate derivatives efficiently underwent 6-*exo*-trig radical cyclization to produce the six-membered ring products, the related *N*-phenyl amides produced a complex mixture of products. Furthermore, attempts to induce 7-*exo*-trig radical cyclizations were unsuccessful and instead resulted in dimerization to afford the corresponding hydrazide. Thus, the success of these amidyl radical cyclization reactions is influenced by the energetics of the cyclization relative to the competing pathways, such as radical–radical dimerization and hydrogen atom abstraction. Further advances in this field were made by Xu and coworkers and enabled the intramolecular oxidative amination of tri- and tetrasubstituted alkenes (Scheme 109).[914]

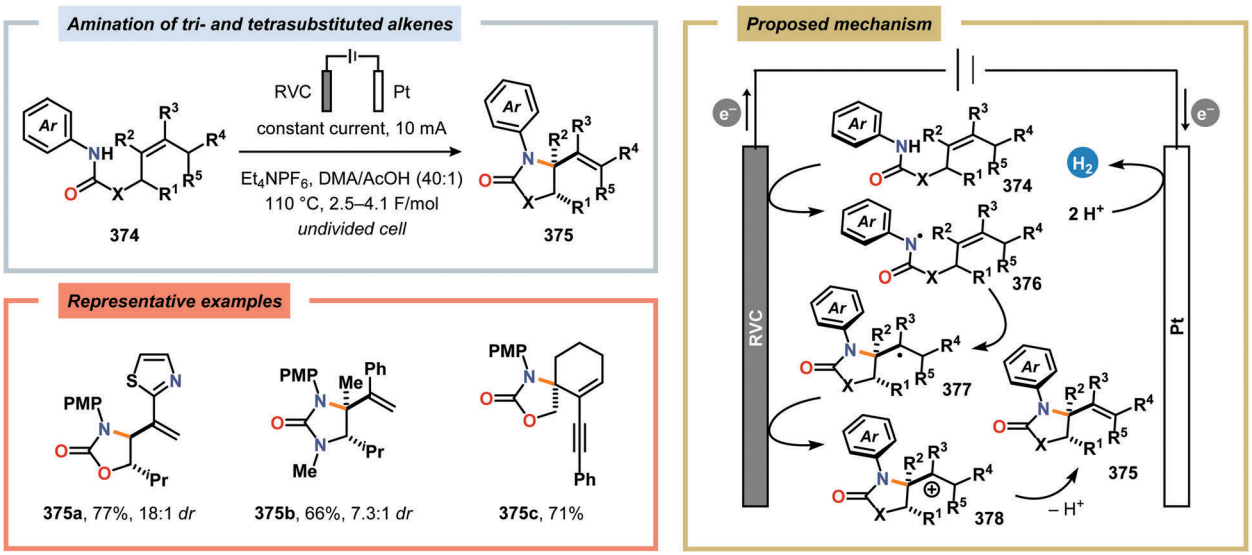
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 107 Electrochemical aziridination of olefins.

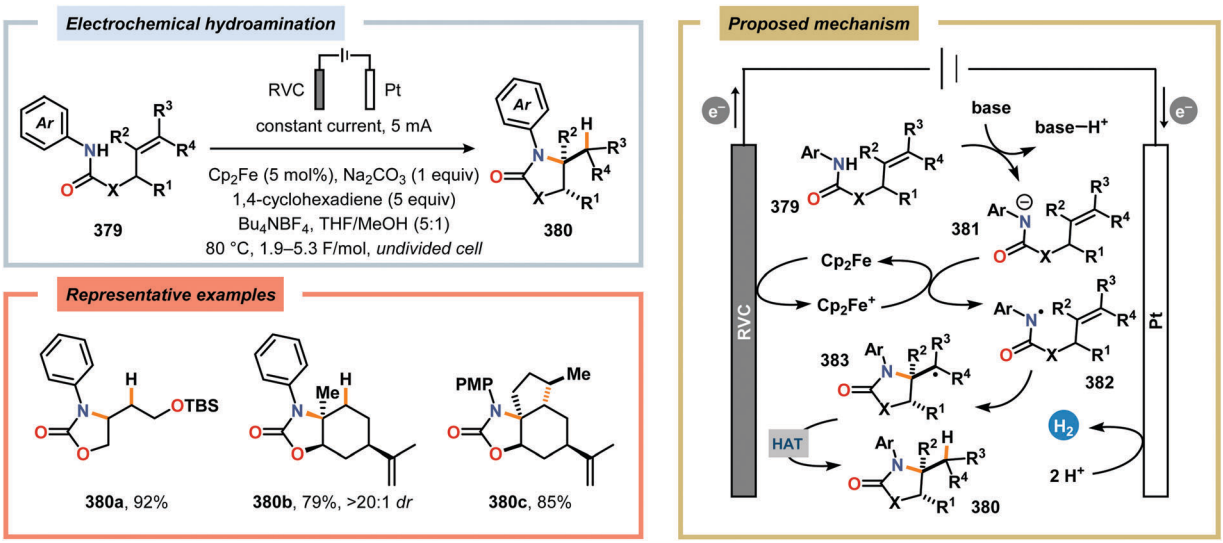


Scheme 108 Electrochemical generation of amidyl radicals for synthesis of γ- and δ-lactams.



Scheme 109 Oxidative amination of tri- and tetrasubstituted alkenes.

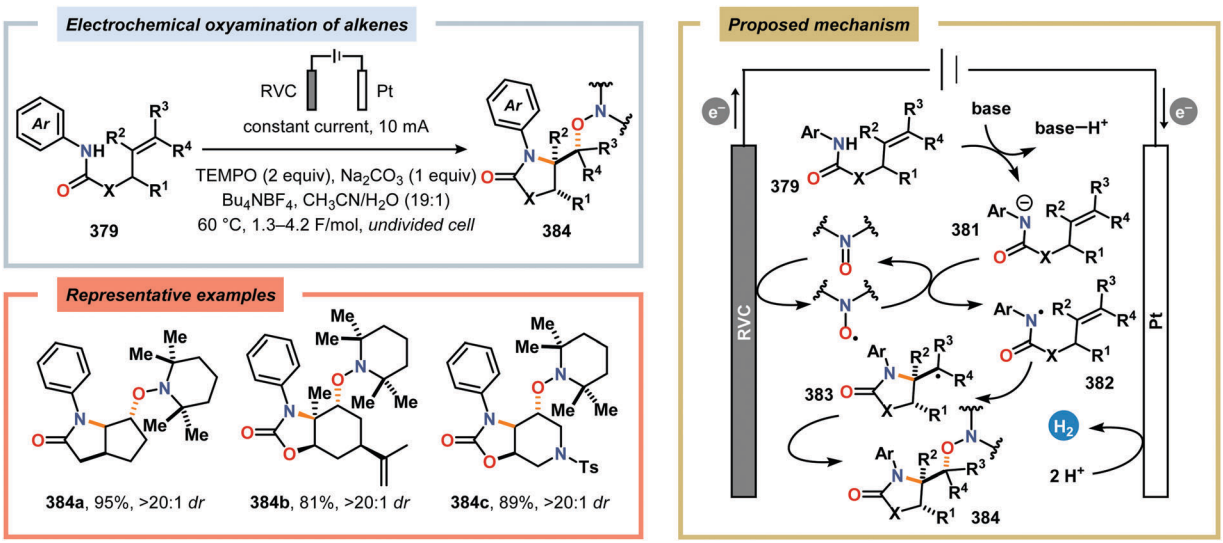
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 110 Electrochemical hydroamination of alkenes *via* ferrocene-mediated generation of amidyl radicals.

A wide range of amide, carbamate and urea derivatives carrying various trisubstituted or sterically demanding tetrasubstituted alkenyl moieties were all viable substrates. The protocol was compatible with a variety of (hetero)aryl-, alkyl- and alkynyl-substituted olefins and a range of functional groups were tolerated, including alcohols, alkynes, aryl bromides, esters, pyridines, silyl ethers, thiazoles and thiophenes. Upon radical cyclization, proton elimination occurred regioselectively at the distal carbon center, providing the cyclized products **375** in good to high diastereoselectivity. Alternatively, the generated carbon-centered radical can also engage in hydrogen atom abstraction to provide hydroamination products[915] *via* ferrocene-mediated generation of amidyl radicals (Scheme 110) or be trapped with *N*-oxyl radicals, such as TEMPO, to give oxyamination products (Scheme 111).[916] The latter reaction has also been conducted using an electrochemical flow microreactor (Scheme 112), making the process less costly and enabling easier purification.[917] Here, TEMPO has a dual function, being the mediator as well as the oxygen source.

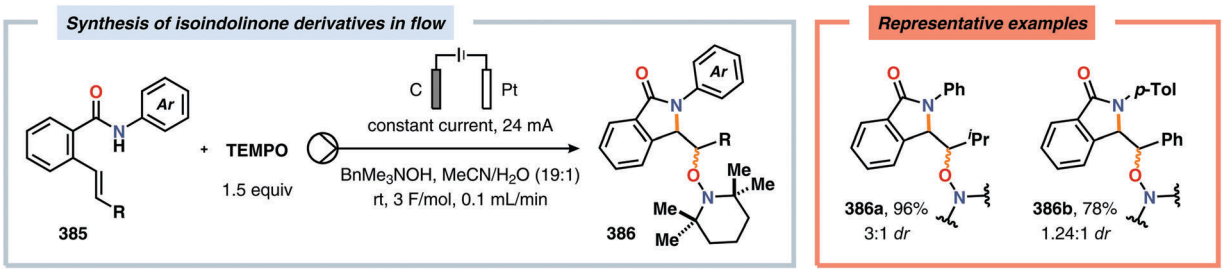
The Waldvogel and Moeller groups have exploited the electrochemical generation of amidyl radicals from anilides, enabling the direct synthesis of benzoxazoles (Scheme 113).[918] Although methanol has been commonly applied in electrochemical transformations involving amidyl radicals, this proved to be incompatible, leading to degradation of anilides **387**. This observation suggested that the intramolecular cyclization process was slow. Of the screened solvents, only HFIP was found to efficiently stabilize the electrochemically generated radical intermediates, affording the desired benzoxazoles **389** in good to high yields. The authors also demonstrated that the aromatic moiety connected to the produced amidyl radicals was of substantial radical nature as the



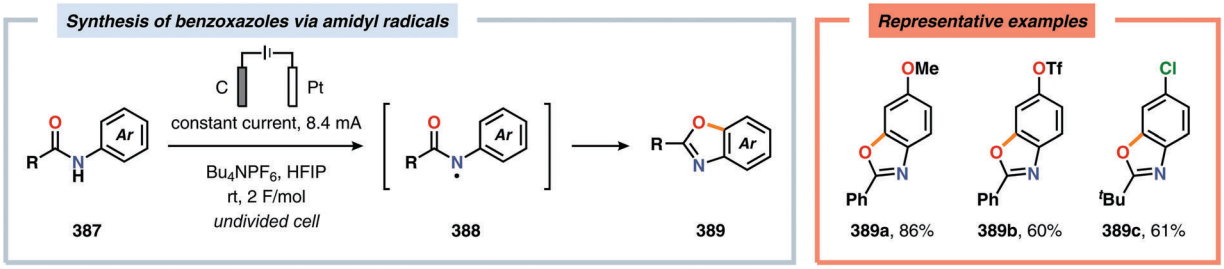
Scheme 111 Electrochemical oxyamination of alkenes *via* TEMPO-mediated generation of amidyl radicals.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



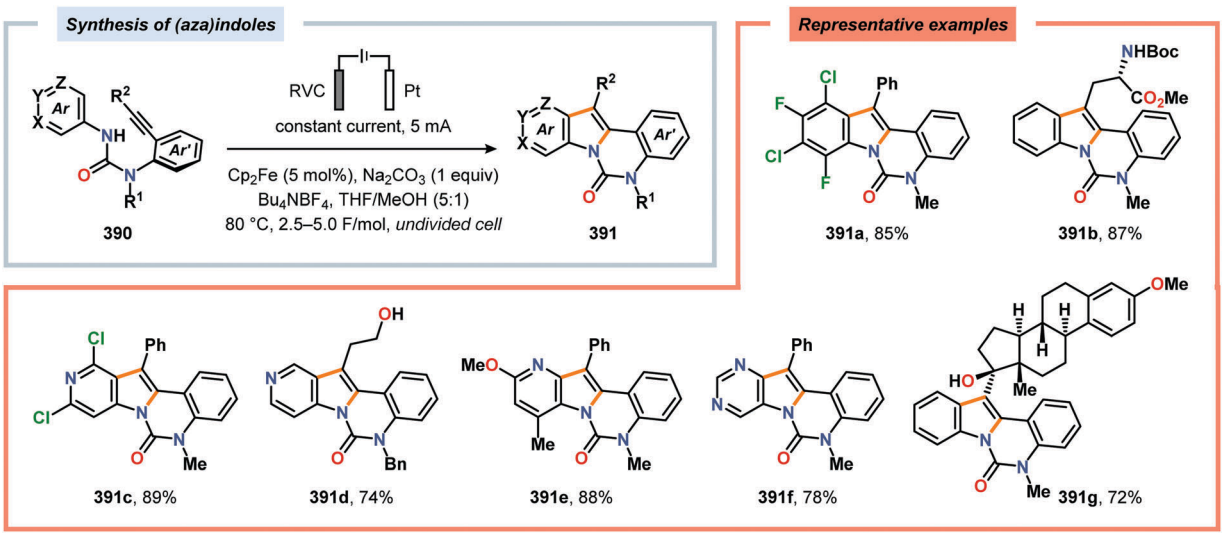
Scheme 112 Synthesis of isoindolinones using an electrochemical flow microreactor.



Scheme 113 Electrochemical synthesis of benzoxazoles from anilides.

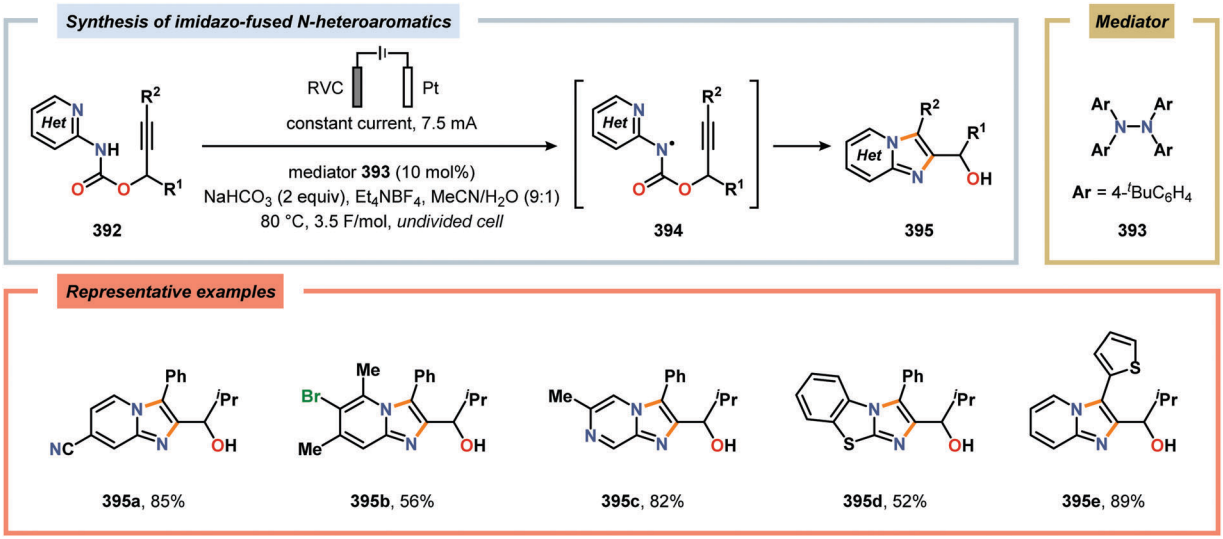
para-position of the *N*-substituted arene ring had to be substituted in order to prevent coupling reactions at this position.

The Xu group has recently made significant contributions through the design of distinct reaction manifolds that capitalize on nitrogen-centered radicals. Various mediator-based radical cyclization manifolds using (hetero)arylamines containing tethered alkynes have been developed. In this context, functionalized (aza)indoles **391** were accessed through an oxidative alkyne annulation reaction (Scheme 114).[919,920] The broad functional group tolerance of the electrochemical protocol was illustrated by the preparation of (aza)indoles housing assorted substituents, such as acetals, alcohols, aldehydes, *N*-aryl carbamates, dipeptides, halides, ketones, (*ortho*)esters and sulfonamides. The reaction is believed to be initiated with the anodic oxidation of $[Cp_2Fe]$ to $[Cp_2Fe]^+$. Subsequent SET between $[Cp_2Fe]^+$ and the deprotonated urea derivative results in regeneration of $[Cp_2Fe]$ and furnishes a nitrogen-centered radical. This radical is suggested to engage in 6-*exo*-dig cyclization, which is followed by a second cyclization with the aryl moiety. Finally, rearomatization of the generated arene radical through oxidation and loss of a proton provides (aza)indoles **391**. A related mediator-based radical cyclization cascade approach was applied to the preparation of imidazo-fused *N*-heteroaromatic compounds (**395**), including imidazo[1,2-*a*]pyridines, imidazo[1,2-*b*]pyridazines and



Scheme 114 Synthesis of (aza)indoles using ferrocene as mediator.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

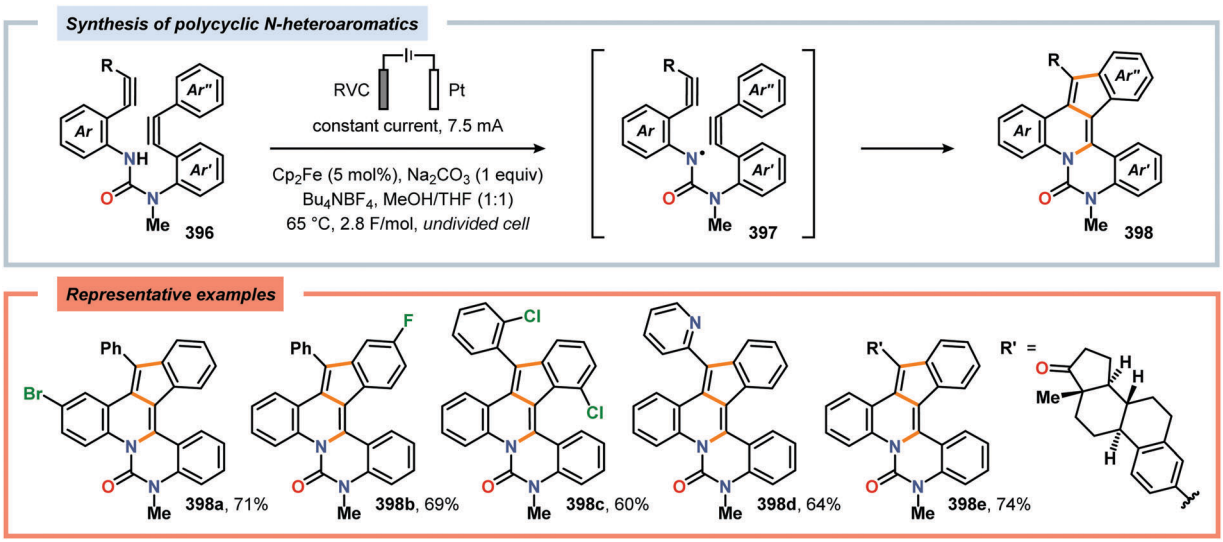


Scheme 115 Synthesis of imidazo-fused N-heteroaromatic compounds through a radical cyclization cascade.

imidazo[1,2-*a*]pyrazines (Scheme 115).[921,922] However, here tetraarylhydrazine **393** was found to be the optimal redox catalyst. Nitrogen-centered radicals have also been harnessed for synthesis of nitrogen-doped polycyclic aromatic hydrocarbons (Scheme 116).[923–925] In the presence of ferrocene, the urea-tethered diynes **396** underwent efficient radical cyclization to afford a variety of electron-rich polycyclic aromatic hydrocarbons (**398**). The use of a mild redox catalyst to facilitate the cascade cyclization instead of carrying out direct electrolysis was crucial to avoid overoxidation of the products as the diynes **396** are oxidized at a higher potential than that of products **398**. Strategies relying on direct electrolysis for generation of nitrogen-centered radicals have also been achieved. Based on this concept, amidinyl and iminyl radicals have been exploited for preparing polycyclic imidazole derivatives[926,927] (Scheme 117) and assorted pyridine-fused aromatic motifs[928] (Scheme 118), respectively. In addition to using nitrogen-centered radicals in C–N bond coupling manifolds, nitrogen–nitrogen (N–N) bond formation has also been accomplished. These intermediates have, for example, been explored for N–N dimerization[929] in the synthesis of dixiamycin B,[930,931] phthalazin-1,4-diones[932] and pyrazolidin-3,5-diones.[933,934]

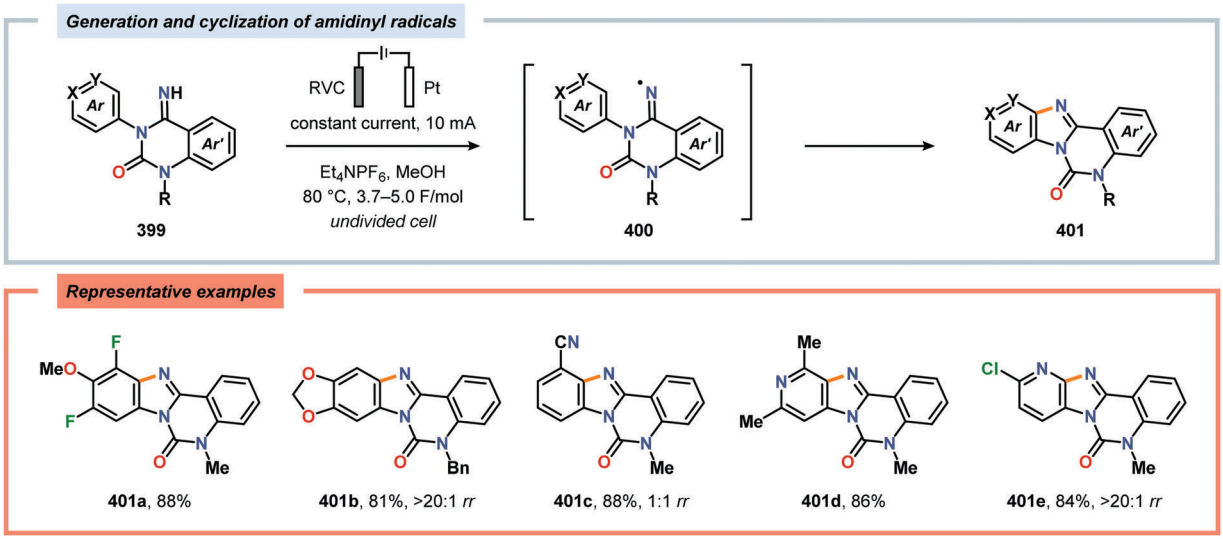
3.3. Pyridine and related nitrogen-containing heterocycles as nitrogen sources

Seminal work by Lund during the 1950s highlighted that aromatic hydrocarbons could be oxidized at a Pt electrode in acetonitrile solutions containing pyridine and $NaClO_4$ to afford the corresponding pyridinium perchlorate species.[935] Although related electrochemical amination reactions were studied by other research groups,[936–943] it took more than five decades until Yoshida and coworkers disclosed an efficient and practical method for C–H amination of aromatic compounds based on electrochemical

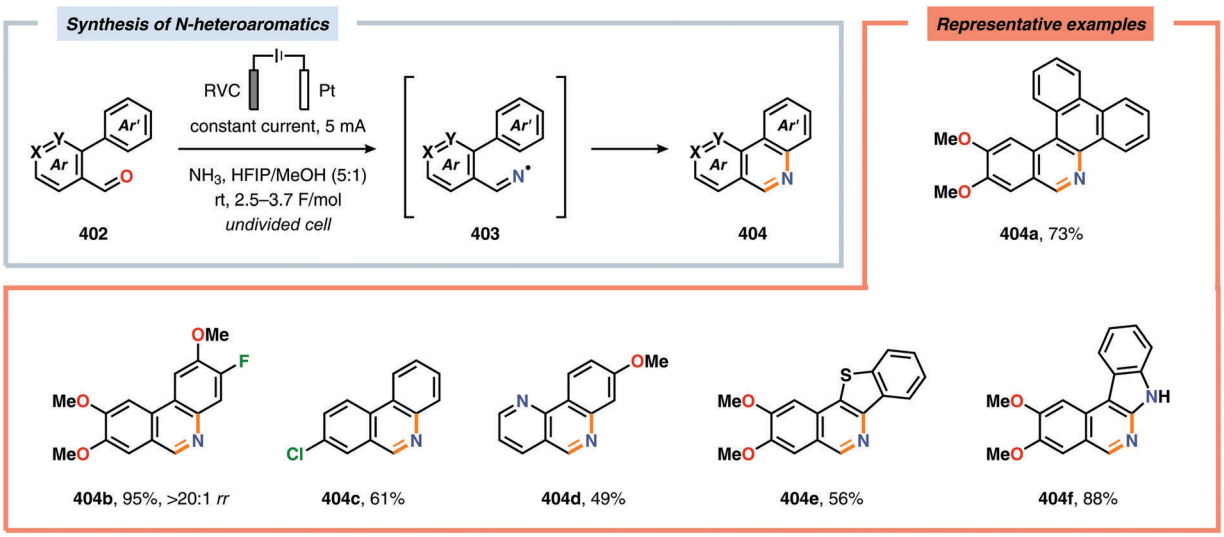


Scheme 116 Electrochemical synthesis of nitrogen-doped polycyclic aromatic hydrocarbons through radical cyclization of diynes.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 117 Electrochemical generation of amidinyl radicals for synthesis of tetracyclic benzimidazoles and pyridoimidazoles.

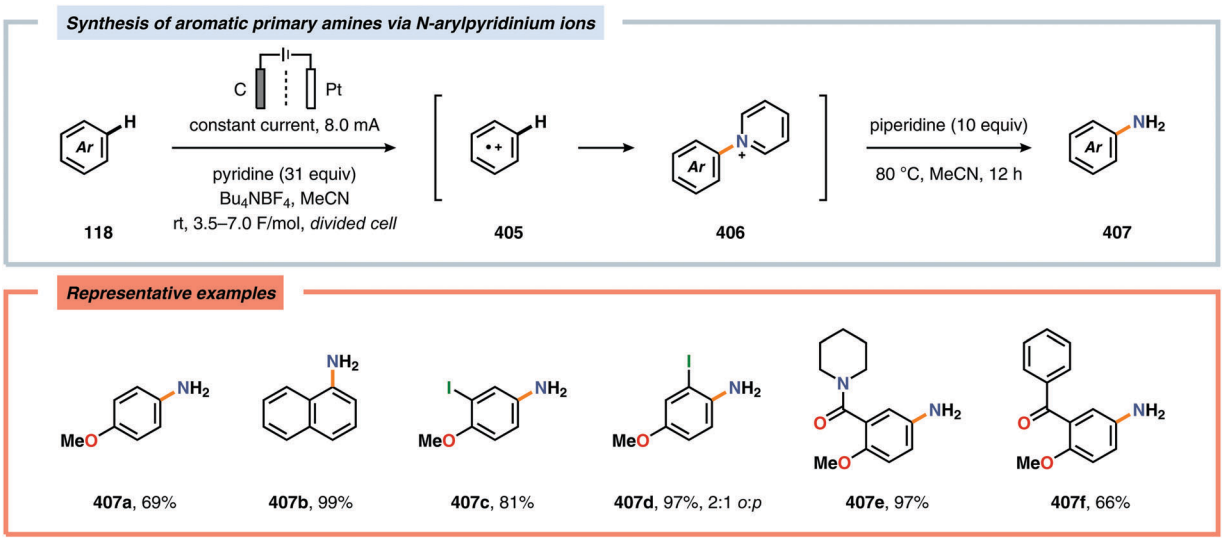


Scheme 118 Synthesis of pyridine-fused polycyclic aromatic motifs *via* iminyl radicals.

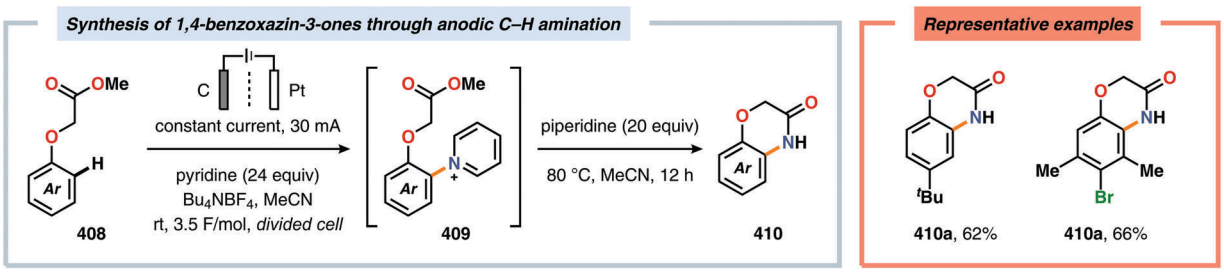
oxidation of aromatic compounds in the presence of pyridine (Scheme 119).[56,944,945] Conducting the electrochemical oxidation in an H-type divided cell at ambient temperature furnished *N*-arylpyridinium ions (**406**), so called Zincke intermediates,[946–948] which upon treatment with an alkylamine resulted in primary amines **407**.[56] Here, the high oxidation potential of pyridine enables selective oxidation of the aromatic compound in the presence of pyridine. Furthermore, the high nucleophilicity of pyridine allows for efficient trapping of the generated arene radical cation and the introduction of multiple amino groups is avoided due to the strong electron-withdrawing effect of the pyridinium moiety in **406**, thereby suppressing overoxidation. However, one drawback was that only electron-rich and activated substrates, such as anisole derivatives, were viable substrates. Subsequently, the Waldvogel group demonstrated that the use of BDD anodes allowed for amination of less activated substrates[949] as well as diamination.[950] Furthermore, a protocol for the anodic C–H amination of phenoxy acetates was shown to provide efficient access to 1,4-benzoxazin-3-ones (Scheme 120).[951] Alternatively, rendering the C–H amination intramolecular through the use of arenes containing a tethered 2-pyrimidyl moiety provided access to 2-aminobenzoxazoles and 2-aminobenzothiazoles (Scheme 121).[952,953]

Work on expanding the scope of the nitrogen nucleophiles revealed that *N*-protected imidazole derivatives were suitable trapping agents. The choice of an appropriate protecting group was crucial to the success of the developed C–H amination method. After an initial examination of various protecting groups, the use of *N*-methylsulfonyl (Ms) protected imidazole afforded the C–N coupled product in the highest yield. The developed protocol

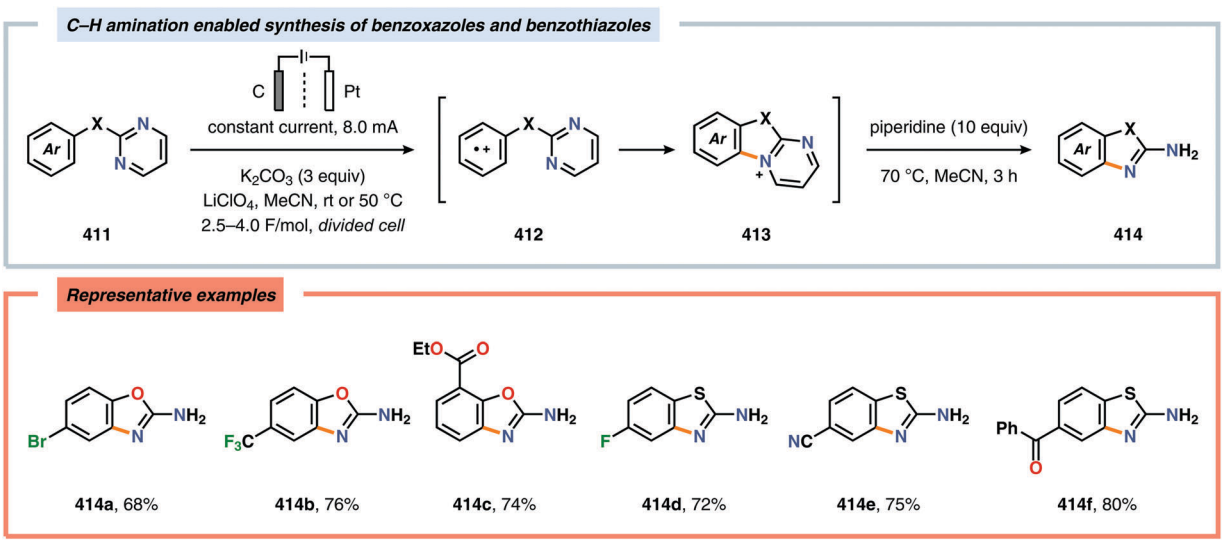
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 119 Electrochemical synthesis of aromatic primary amines *via* *N*-arylpyridinium ions.



Scheme 120 Synthesis of 1,4-benzoxazin-3-ones *via* anodic C–H amination.

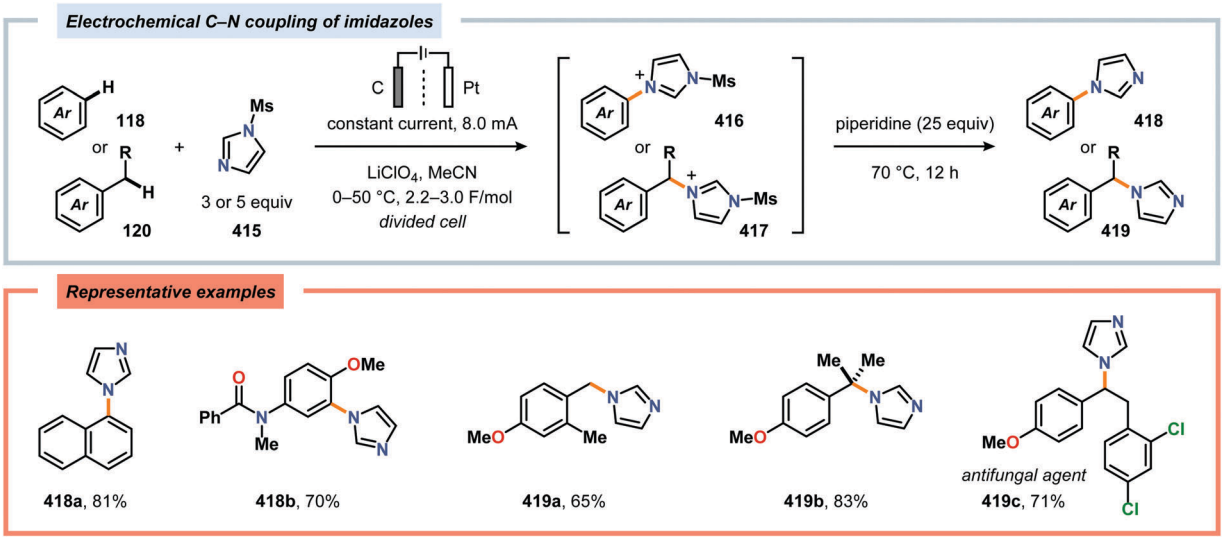


Scheme 121 Preparation of 2-aminobenzoxazoles and 2-aminobenzothiazoles *via* electrochemical intramolecular C–H amination.

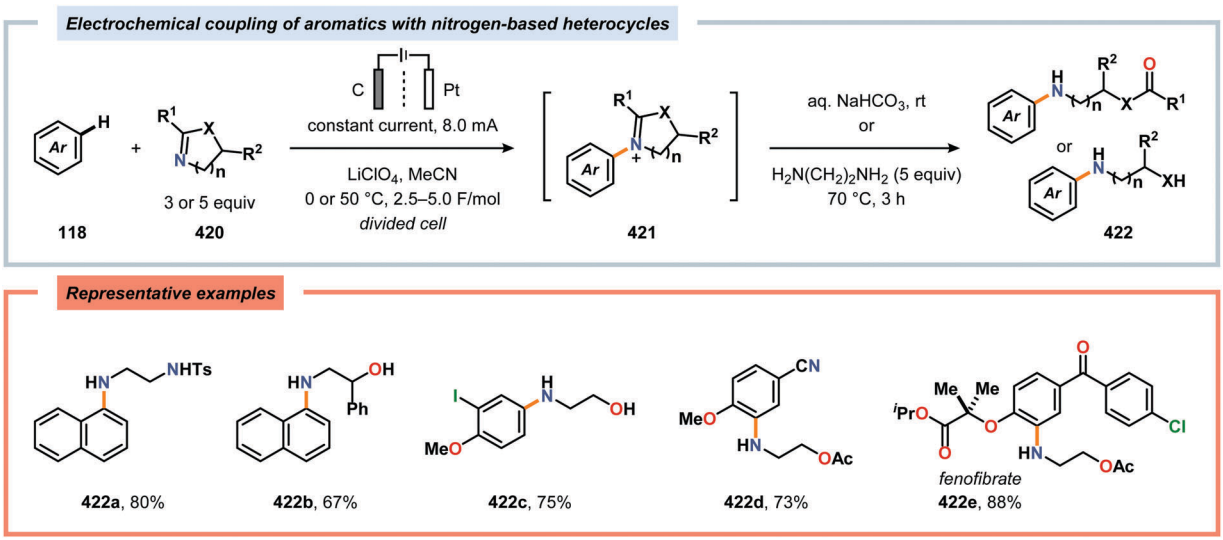
was applicable not only to aromatic but also to benzylic C–H amination. Here, the initial *N*-aryl or *N*-benzylimidazolium cations can be subjected to non-oxidative removal of the protecting group, resulting in cleavage of the sulfonyl protecting group to produce the corresponding *N*-aryl or *N*-benzylimidazoles (Scheme 122). Similar to *N*-arylpyridinium ions **406**, the intermediate imidazolium ions

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



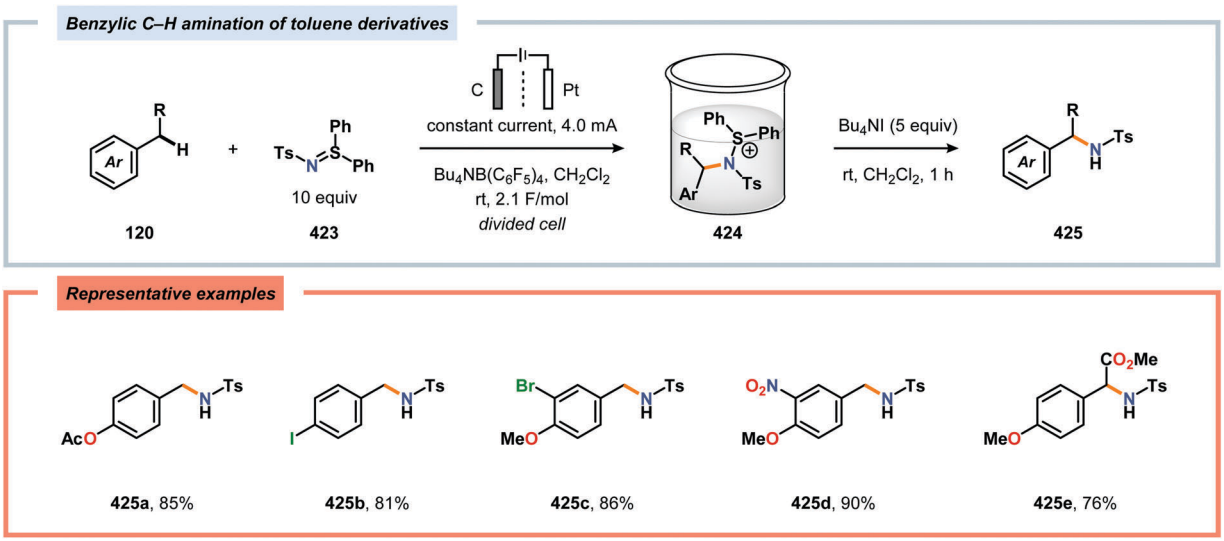
Scheme 122 Direct C–N coupling of imidazoles with aromatic and benzylic compounds.



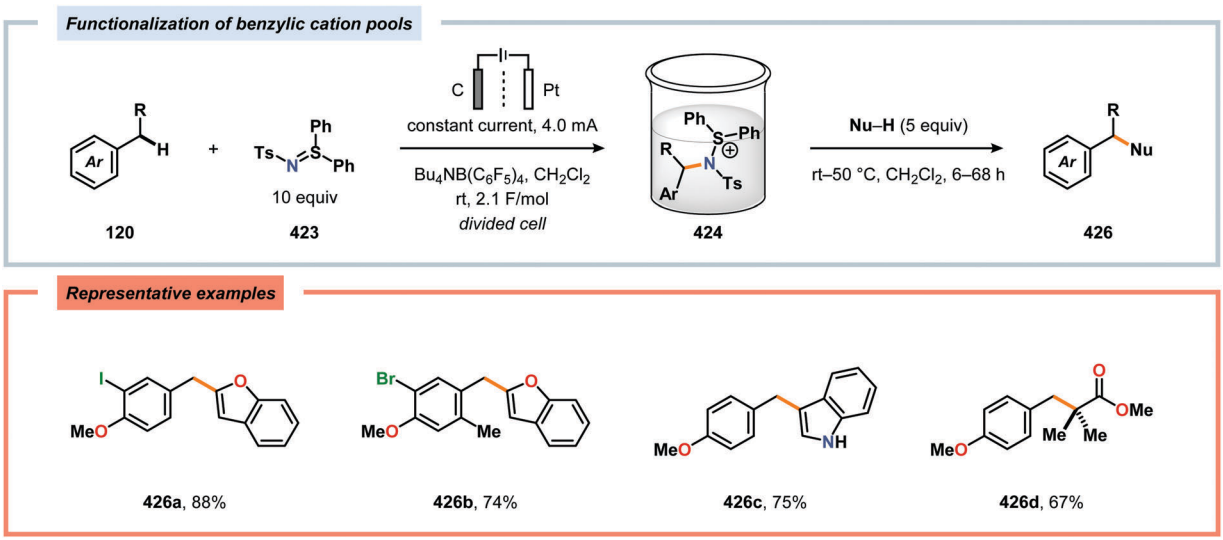
Scheme 123 C–N bond coupling of aromatic compounds and nitrogen-containing heterocycles.

416 and **417** are electrochemically inactive, thus preventing over-oxidation and allow these intermediates to be converted to the corresponding imidazole products **418** and **419**.[954–956] Subsequently, the Yoshida group reported a mechanistically similar reaction for the coupling of aromatics and masked primary alkyl amines bearing additional oxygen or nitrogen functionalities (Scheme 123). The masked primary amines **420** were prepared by treatment of the amines with nitriles or derivatives thereof to afford the target five- or six-membered heterocycle. Under electrochemical conditions, these heterocycles undergo coupling with arene radical cations to give cationic intermediates **421**, which can be chemically converted to the desired cross-coupling products **422** under non-oxidative conditions.[957] The same group also developed an electrochemical approach for C–H amination of toluene derivatives employing *N*-tosyldiphenylsulfilimine **423** as the amination reagent (Scheme 124). This enabled the preparation of *N*-tosylbenzylaminosulfonium ions **424**, which upon treatment with an iodide source, such as Bu_4NI, under non-electrolytic conditions results in N–S bond cleavage to yield the corresponding *N*-tosylbenzylamines **425**.[958] Alternatively, the generated benzylaminosulfonium ions **424** can react with added aromatic nucleophiles to give the corresponding cross-coupling products **426** (Scheme 125).[959] In these transformations, the high oxidation potential of *N*-tosyldiphenylsulfilimine **423** allows a wide range of toluene derivatives to be oxidized and functionalized. Finally, Lei and coworkers have developed an electrochemical strategy for amination of C(sp^3)–H bonds. Here, C(sp^3)–H bonds adjacent to heteroatoms, such as nitrogen, oxygen and sulfur, could

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



Scheme 124 C–H amination *via* electrochemically generated benzylaminosulfonium ions.



Scheme 125 Benzylic cross-coupling *via* electrochemically generated benzylaminosulfonium ions.

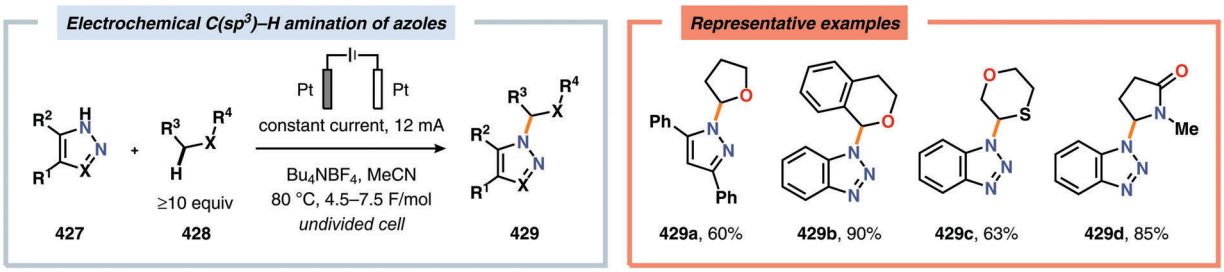
be functionalized with various amines to give the corresponding C–N coupled products **429** in moderate to good yields (Scheme 126).[960,961]

3.4. Halide-mediated approaches to C–N bond coupling

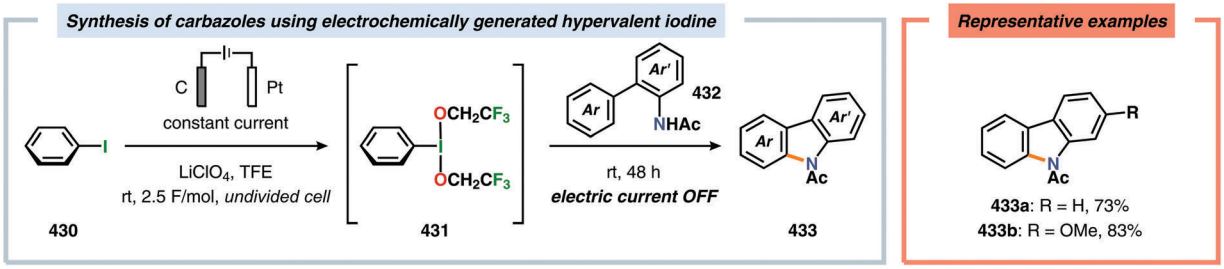
Hypervalent iodine compounds have been extensively employed in organic synthesis as selective metal-free oxidants and environmentally friendly reagents. Synthetic applications of these versatile reagents include, for example, aminations, C–C bond-forming reaction manifolds, halogenations, oxidations, rearrangements, and various dual catalytic activation modes.[962–969] In this context, hypervalent iodine species electrochemically generated from the corresponding iodoarenes have been applied for the synthesis of various heterocyclic frameworks, such as carbazoles[970] (Scheme 127), pyrroloindoles,[971] quinolinone derivatives[972] (Scheme 128) and spirocycles (Scheme 129).[973,974] The utility of electrochemically generated hypervalent iodine reagents has also been demonstrated in the synthesis of glycozoline[970] and tetrahydropyrroloiminoquinone alkaloids.[975,976] The supporting electrolyte has also been merged with the iodine mediator through tethering of the redox-active iodophenyl moiety to an alkylammonium moiety. The ionically tagged iodophenyl motif was applied to several oxidative C–N bond coupling scenarios and allowed for straightforward recovery and reuse of the ionic mediator.[977,978] A related approach was also employed for the synthesis of benzoxazoles.[979] Additionally, various oxidative amination strategies for preparing 2-substituted benzoxazoles using iodide as a redox mediator have also been reported.[980–982]

An electrochemical protocol for α-amination of ketones was recently reported using iodide as a redox catalyst (Scheme 130).[983] Here, NH_4I was found to be the optimal redox catalyst for facilitating

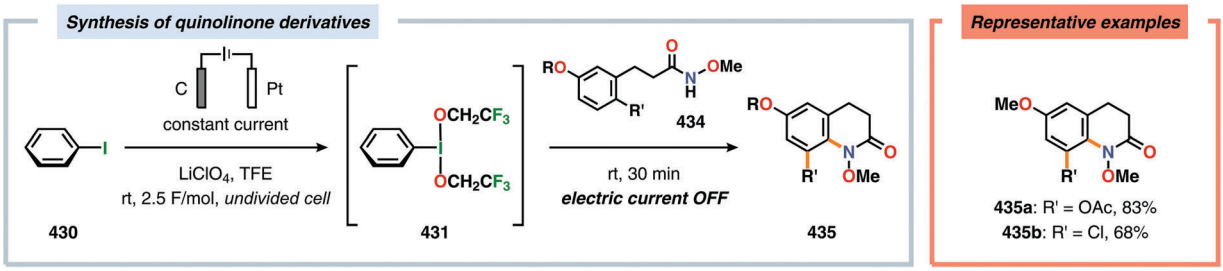
Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.



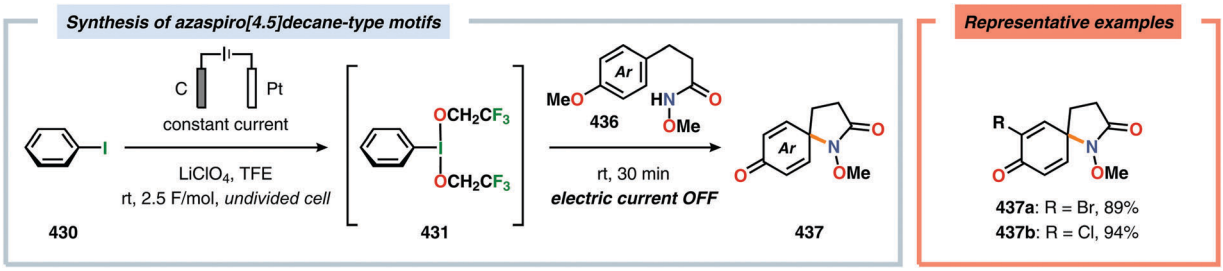
Scheme 126 Electrochemical C(sp^3)–H amination of azoles with heterocycles.



Scheme 127 Synthesis of carbazoles using an electrochemically generated hypervalent iodine oxidant.



Scheme 128 Preparation of quinolinone derivatives through oxidative cyclization and concomitant rearrangement of the functional group.

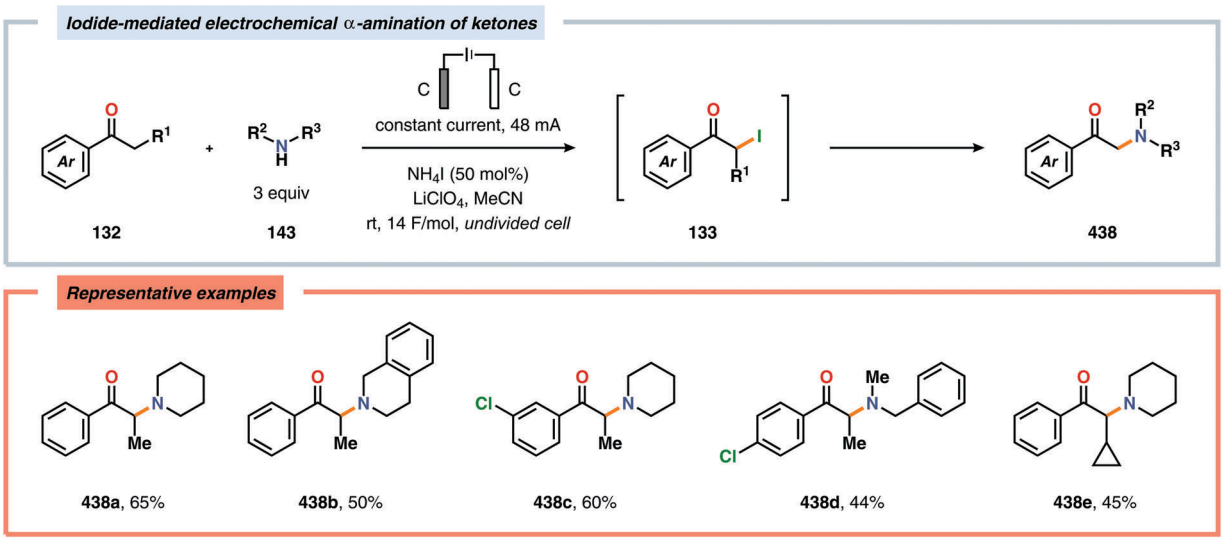


Scheme 129 Hypervalent iodine-mediated synthesis of azaspiro[4.5]decane-type motifs.

the coupling of ketones and secondary amines at ambient temperature. The reactions were conveniently carried out in a simple undivided beaker-type cell and were believed to proceed *via* initial α-iodination of the ketones (**132**) with the anodically generated I_2, followed by nucleophilic substitution of the amines (**143**), thus avoiding isolation of the key α-iodo ketone intermediate **133**. It was noted that the addition of radical scavangers, such as TEMPO, terminated the α-amination process and did not yield any product. Related halide-mediated procedures for cleavage of β-O-4 lignin model compounds,[984–986] thiocyanation[987] as well as synthesis of 3-amino-2-thiocyanato-α,β-unsaturated carbonyl derivatives[988] and β-keto sulfones[989] have also been developed.[990]

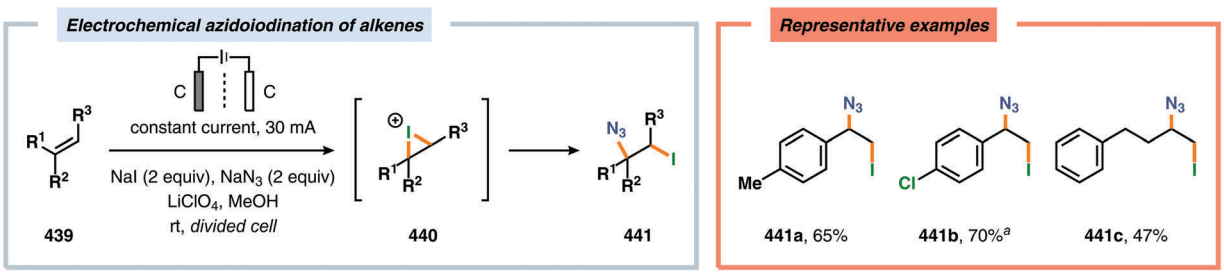
Halide-mediated protocols for C–N bond coupling with alkenes have also been targeted. Zeng, Little and coworkers

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

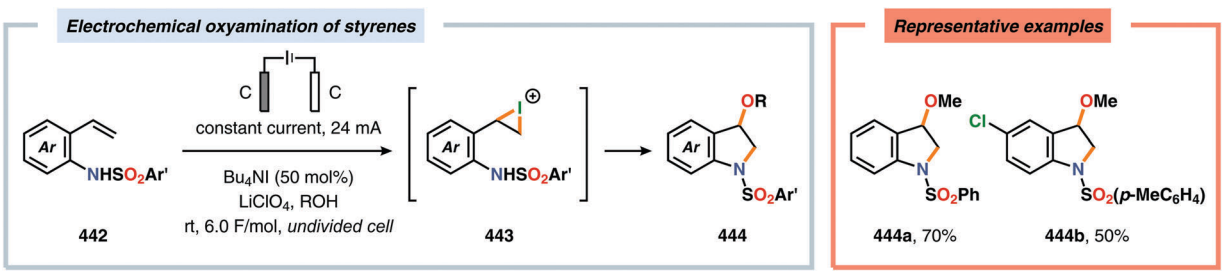


Scheme 130 Electrochemical α-amination of ketones using NH_4I as redox catalyst.

demonstrated an efficient approach for aziridination of alkenes Bu_4NI as a redox catalyst and *N*-aminophthalimide (**366**) as the source of electrophilic nitrogen (*cf.* Scheme 107). The process was carried out at a constant current in an undivided cell and was proposed to follow a radical mechanism where hydrogen atom abstraction from *N*-aminophthalimide by $I^{\bullet}$ generates a nitrogen-centered radical that is subsequently trapped by the alkene.[991] Based on earlier precendent,[992] the Zeng group presented an indirect electrochemical method for synthesis of vicinal iodoazides (Scheme 131). In the developed protocol, IN_3 is electrochemically generated *in situ*, thus avoiding the use of external oxidants or corrosive I_2. Furthermore, the reaction proceeds *via* a cyclic iodonium intermediate to afford regioselective azidoiodination products in a Markovnikov-type fashion.[993] Finally, an electrochemical strategy for oxyamination of styrenes for accessing indoline derivatives was recently achieved (Scheme 132).[994] The electrochemical reactions were conducted under constant current conditions in a simple undivided cell using Bu_4NI as a redox catalyst. This paired electrolysis process avoids the use of external bases and oxidants, representing an appealing and environmentally benign route for synthesis of these heterocyclic compounds. Mechanistic insight suggested that the reaction



Scheme 131 Electrochemical regioselective azidoiodination of alkenes *via in situ* generation of IN_3. [a] Carried out in $MeOH/H_2O$ (5 : 1).



Scheme 132 Electrochemical oxyamination of styrenes for synthesis of indolines.

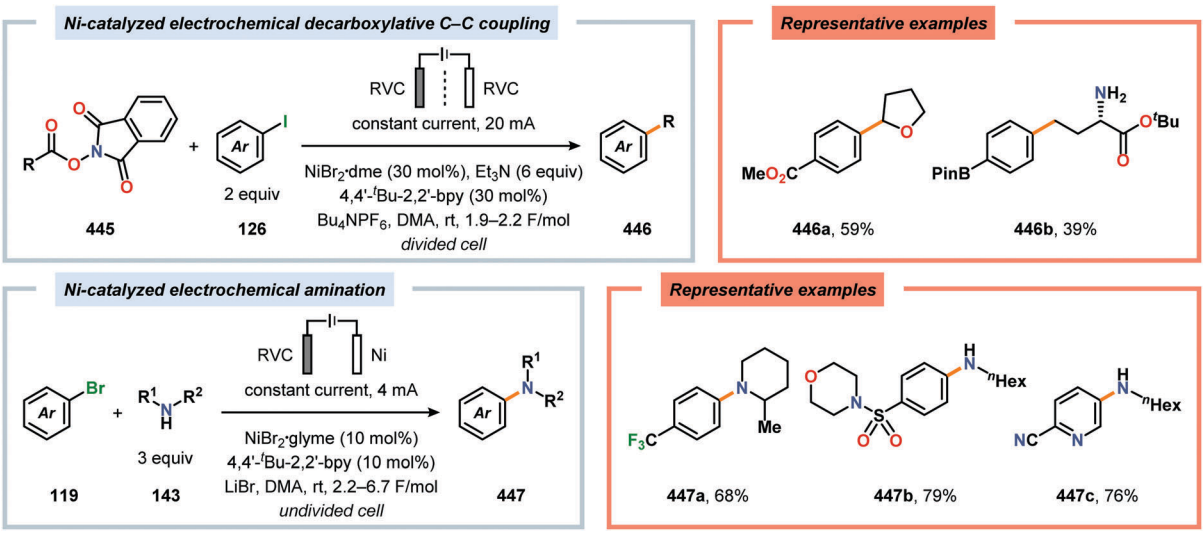
 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

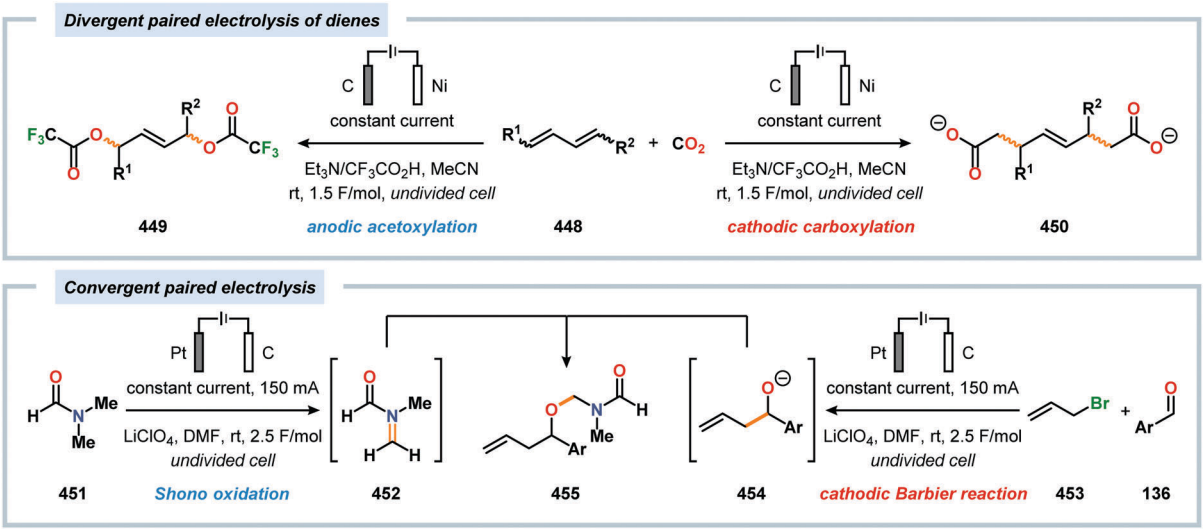
commences with the anodic oxidation of iodide to generate I_2, which subsequently reacts with alkene **442** to afford the corresponding iodonium intermediate **443**. Finally, intramolecular cyclization with the pendant sulfonamide followed by nucleophilic substitution by alkoxide yields the indoline product **444**. The examples discussed here highlight that halide ions are versatile mediators capable of promoting a collection of different transformations.

4. Conclusions and outlook

During the last few decades, the imperative to develop and adopt more sustainable and atom economical methodologies for synthetic purposes has not been overlooked by the scientific community. In this regard, electrochemistry serves as an appealing platform since only electrons serve as reagents,[995] thereby minimizing production of reagent waste while allowing the reactions to be carried out at milder conditions.[996,997] Organic electrochemistry can be classified into two categories, direct and indirect electrolysis. Here, the initial electron transfer between the electrode and a substrate to produce a radical intermediate does not generally pose a problem. Instead, it is crucial to control the reactive intermediates that are subsequently generated throughout the process. Although related activated species can be accessed using traditional organic reagents, there is a dramatic difference in the nature of these species.[998] While the activated species are uniformly distributed throughout the solution in conventional organic reactions, these species are only produced at the surface of the electrode in electrochemical reactions. This difference in distribution of the reactive species will thus affect the chemical behavior as well as reactivity and selectivity.



Scheme 133 Ni-catalyzed electrochemical coupling platforms.



Scheme 134 Divergent and convergent paired electrolysis.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

For an unexperienced electrochemist it might seem that there are a wide range of experimental variables that need to be optimized, such as cell type, electrode composition and electrolyte.[999] However, this should be seen as an opportunity for accessing unique selectivity and expediting unconventional reaction pathways for challenging bond construction, thus enabling novel and strategic bond disconnections in synthetic endeavors. Since organic chemists are rather unfamiliar with the concept of using electric current to facilitate chemical reactions, the purpose of the present review is to highlight the potential of applying organic electrochemistry to organic synthesis. Here, an overview of the development of electrosynthetic reaction manifolds for C–H bond functionalization and C–N bond formation is provided. The electrochemical strategies and methods discussed in this review clearly showcase the diversity of transformations that can be facilitated through organic electrosynthesis. Furthermore, the refinement and expansion of reaction engineering methods and wider commercial availability of electrochemical reactors provide a multifaceted and powerful toolbox for efficient generation of reactive intermediates from an ensemble of organic substrates. These include, among others, the development of dual catalytic platforms (Scheme 133),[1000–1007] paired electrolysis[1008–1012] (Scheme 134) and electrochemical flow cells,[1013–1019] which have contributed to advancing the field of electrosynthesis. Based on these unique features and the current renaissance in synthetic electrochemistry, it is anticipated that this field will soon be considered as a reliable and versatile platform by the broader scientific community for nontraditional bond construction in organic chemistry.

Conflicts of interest

There are no conflicts of interest to declare.

Acknowledgements

Financial support from the Swedish Research Council (637-2013-7314) is gratefully acknowledged. The author would also like to thank Dr Bryan S. Matsuura (New York University, New York) for helpful discussions and feedback on this manuscript.

References

1 G. M. Whitesides, *Isr. J. Chem.*, 2018, **58**, 142–150.
2 R. Noyori, *Nat. Chem.*, 2009, **1**, 5–6.
3 G. M. Whitesides, *Angew. Chem., Int. Ed.*, 2015, **54**, 3196–3209.
4 *Organic chemistry – Breakthroughs and perspectives*, ed. K. Ding and L.-X. Dai, Wiley-VCH Verlag & Co. KGaA, Weinheim, Germany, 2012.
5 P. Wyatt and S. Warren, *Organic synthesis: Strategy and control*, John Wiley and Sons Ltd., Chichester, England, 2007.
6 For selected reviews, see: (*a*) K. C. Nicolaou, D. Vourloumis, N. Winssinger and P. S. Baran, *Angew. Chem., Int. Ed.*, 2000, **39**, 44–122; (*b*) P. A. Wender, V. A. Verma, T. J. Paxton and T. H. Pillow, *Acc. Chem. Res.*, 2008, **41**, 40–49; (*c*) I. S. Young and P. S. Baran, *Nat. Chem.*, 2009, **1**, 193–205; (*d*) K. C. Morrison and P. J. Hergenrother, *Nat. Prod. Rep.*, 2014, **31**, 6–14; (*e*) X.-W. Li and B. Nay, *Nat. Prod. Rep.*, 2014, **31**, 533–549; (*f*) A. M. Armaly, Y. C. DePorre, E. J. Groso, P. S. Riehl and C. S. Schindler, *Chem. Rev.*, 2015, **115**, 9232–9276; (*g*) R. Ardkhean, D. F. J. Caputo, S. M. Morrow, H. Shi, Y. Xiong and E. A. Anderson, *Chem. Soc. Rev.*, 2016, **45**, 1557–1569; (*h*) C. Cabrele and O. Reiser, *J. Org. Chem.*, 2016, **81**, 10109–10125.
7 For selected reviews on the use of renewable chemical feedstocks, see: (*a*) C.-J. Li and B. M. Trost, *Proc. Natl. Acad. Sci. U. S. A.*, 2008, **105**, 13197–13202; (*b*) Q. Liu, L. Wu, R. Jackstell and M. Beller, *Nat. Commun.*, 2015, **6**, 5933; (*c*) G. Fiorani, W. Guo and A. W. Kleij, *Green Chem.*, 2015, **17**, 1375–1389; (*d*) A. Otto, T. Grube, S. Schiebahn and D. Stolten, *Energy Environ. Sci.*, 2015, **8**, 3283–3297; (*e*) Q.-W. Song, Z.-H. Zhou and L.-N. He, *Green Chem.*, 2017, **19**, 3707–3728.
8 For selected reviews on biocatalysis, see: (*a*) U. T. Bornscheuer, G. W. Huisman, R. J. Kazlauskas, S. Lutz, J. C. Moore and K. Robins, *Nature*, 2012, **485**, 185–194; (*b*) M. T. Reetz, *J. Am. Chem. Soc.*, 2013, **135**, 12480–12496; (*c*) M. Hönig, P. Sondermann, N. J. Turner and E. M. Carreira, *Angew. Chem., Int. Ed.*, 2017, **56**, 8942–8973; (*d*) R. A. Sheldon and P. C. Pereira, *Chem. Soc. Rev.*, 2017, **46**, 2678–2691; (*e*) F. Schwizer, Y. Okamoto, T. Heinisch, Y. Gu, M. M. Pellizzoni, V. Lebrun, R. Reuter, V. Köhler, J. C. Lewis and T. R. Ward, *Chem. Rev.*, 2018, **118**, 142–231; (*f*) R. A. Sheldon and J. M. Woodley, *Chem. Rev.*, 2018, **118**, 801–838.
9 For discussions and applications of base-metal catalysis, see: (*a*) J. R. Ludwig and C. S. Schindler, *Chem*, 2017, **2**, 313–316; (*b*) A. Fürstner, *ACS Cent. Sci.*, 2016, **2**, 778–789; (*c*) J. E. Zweig, D. E. Kim and T. R. Newhouse, *Chem. Rev.*, 2017, **117**, 11680–11752; (*d*) P. J. Chirik, *Angew. Chem., Int. Ed.*, 2017, **56**, 5170–5181.
10 For selected reviews on cascade/tandem reactions in organic synthesis, see: (*a*) K. C. Nicolaou, D. J. Edmonds and P. G. Bulger, *Angew. Chem., Int. Ed.*, 2006, **45**, 7134–7186; (*b*) K. C. Nicolaou and J. S. Chen, *Chem. Soc. Rev.*, 2009, **38**, 2993–3009; (*c*) C. Grondal, M. Jeanty and D. Enders, *Nat. Chem.*, 2010, **2**, 167–178; (*d*) Y. Hayashi, *Chem. Sci.*, 2016, **7**, 866–880.
11 Q.-L. Zhou, *Angew. Chem., Int. Ed.*, 2016, **55**, 5352–5353.
12 B. König, *Eur. J. Org. Chem.*, 2017, 1979–1981.
13 For recent reviews on visible light photocatalysis, see: (*a*) M. H. Shaw, J. Twilton and D. W. C. MacMillan, *J. Org. Chem.*, 2016, **81**, 6898–6926; (*b*) D. Staveness, I. Bosque and C. R. J. Stephenson, *Acc. Chem. Res.*, 2016, **49**, 2295–2306; (*c*) M. D. Levin, S. Kim and F. D. Toste, *ACS Cent. Sci.*, 2016, **2**, 293–301; (*d*) J. C. Tellis, C. B. Kelly, D. N. Primer, M. Jouffroy, N. R. Patel and G. A. Molander, *Acc. Chem. Res.*, 2016, **49**, 1429–1439; (*e*) E. C. Gentry and R. R. Knowles, *Acc. Chem. Res.*, 2016, **49**, 1546–1556; (*f*) T. Koike and M. Akita, *Acc. Chem. Res.*, 2016, **49**,

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

1937–1945; (*g*) M. D. Kärkäs, J. A. Porco, Jr. and C. R. J. Stephenson, *Chem. Rev.*, 2016, **116**, 9683–9747; (*h*) K. L. Skubi, T. R. Blum and T. P. Yoon, *Chem. Rev.*, 2016, **116**, 10035–10074; (*i*) N. A. Romero and D. A. Nicewicz, *Chem. Rev.*, 2016, **116**, 10075–10166; (*j*) N. Corrigan, S. Shanmugam, J. Xu and C. Boyer, *Chem. Soc. Rev.*, 2016, **45**, 6165–6212; (*k*) J.-R. Chen, D.-M. Yan, Q. Wei and W.-J. Xiao, *ChemPhotoChem*, 2017, **1**, 148–158; (*l*) J. Twilton, C. Le, P. Zhang, M. H. Shaw, R. W. Evans and D. W. C. MacMillan, *Nat. Rev. Chem.*, 2017, **1**, 0052; (*m*) M. D. Kärkäs, *ACS Catal.*, 2017, **7**, 4999–5022.

14 K. D. Moeller, *Electrochemistry*, 2006, **74**, 583.

15 A. V. Shchepochkin, O. N. Chupakhin, V. N. Charushin and V. A. Petrosyan, *Russ. Chem. Rev.*, 2013, **82**, 747–771.

16 V. V. Turygin and A. P. Tomilov, *Russ. J. Electrochem.*, 2015, **51**, 999–1020.

17 K. Chiba and Y. Okada, *Curr. Opin. Electrochem.*, 2017, **2**, 53–59.

18 The auxiliary electrode is the anode for a reduction and cathode for an oxidation reaction.

19 O. R. Luca, J. L. Gustafson, S. M. Maddox, A. Q. Fenwick and D. C. Smith, *Org. Chem. Front.*, 2015, **2**, 823–848.

20 C. Gütz, B. Klöckner and S. R. Waldvogel, *Org. Process Res. Dev.*, 2016, **20**, 26–32.

21 For a review on electrode materials, see: R. L. McCreery, *Chem. Rev.*, 2008, **108**, 2646–2687.

22 For a review on electrochemical synthesis on boron-doped diamond electrode materials, see: S. R. Waldvogel and B. Elsler, *Electrochim. Acta*, 2012, **82**, 434–443.

23 (*a*) J. Y. Becker, B. E. Smart and T. Fukunaga, *J. Org. Chem.*, 1988, **53**, 5714–5720; (*b*) Y. Matsumura, M. Yamada, N. Kise and M. Fujiwara, *Tetrahedron*, 1995, **51**, 6411–6418.

24 K. Fujimoto, Y. Tokuda, H. Maekawa, Y. Matsubara, T. Mizuno and I. Nishiguchi, *Tetrahedron*, 1996, **52**, 3889–3896.

25 For reviews, see: (*a*) H. Liu, Y. Liu and J. Li, *Phys. Chem. Chem. Phys.*, 2010, **12**, 1685–1697; (*b*) M. Kathiresan and D. Velayutham, *Chem. Commun.*, 2015, **51**, 17499–17516; (*c*) M. Alvarez-Guerra, J. Albo, E. Alvarez-Guerra and A. Irabien, *Energy Environ. Sci.*, 2015, **8**, 2574–2599. For selected examples, see; (*d*) M. Mellah, J. Zeitouny, S. Gmouh, M. Vaultier and V. Jouikov, *Electrochem. Commun.*, 2005, **7**, 869–874; (*e*) A. C. Herath and J. Y. Becker, *Electrochim. Acta*, 2008, **53**, 4324–4330.

26 For recent reviews, see: (*a*) K. E. Toghill, M. A. Méndez and P. Voyame, *Electrochem. Commun.*, 2014, **44**, 27–30; (*b*) J. A. Branch and P. N. Bartlett, *Philos. Trans. R. Soc., A*, 2015, **373**, 20150007.

27 L. Eberson and K. Nyberg, *Tetrahedron*, 1976, **32**, 2185–2206.

28 L. Eberson, *J. Mol. Catal.*, 1983, **20**, 27–52.

29 For reviews on radical cations, see: (*a*) A. Ledwith, *Acc. Chem. Res.*, 1972, **5**, 133–139; (*b*) H. D. Roth, *Acc. Chem. Res.*, 1987, **20**, 343–350; (*c*) N. L. Bauld, D. J. Bellville, B. Harirchian, K. T. Lorenz, R. A. Pabon, Jr., D. W. Reynolds, D. D. Wirth, H. S. Chiou and B. K. Marsh, *Acc. Chem. Res.*, 1987, **20**, 371–378; (*d*) N. L. Bauld, *Tetrahedron*, 1989, **45**, 5307–5363; (*e*) A. Albini, E. Fasani and N. d'Alessandro, *Coord. Chem. Rev.*, 1993, **125**, 269–281; (*f*) M. Schmittel and A. Burghart, *Angew. Chem., Int. Ed. Engl.*, 1997, **36**, 2550–2589; (*g*) A. A. Fokin and P. R. Schreiner, *Chem. Rev.*, 2002, **102**, 1551–1594.

30 For reviews on radical anions, see: (*a*) L. M. Dorfman, *Acc. Chem. Res.*, 1970, **3**, 224–230; (*b*) N. L. Holy and J. D. Marcum, *Angew. Chem., Int. Ed. Engl.*, 1971, **10**, 115–124; (*c*) N. L. Holy, *Chem. Rev.*, 1974, **74**, 243–277; (*d*) F. Gerson and W. Huber, *Acc. Chem. Res.*, 1987, **20**, 85–90.

31 For reviews on the applications of radical reactions in organic synthesis, see: (*a*) E. Godineau and Y. Landais, *Chem. – Eur. J.*, 2009, **15**, 3044–3055; (*b*) L. J. Sebren, J. J. Devery III and C. R. J. Stephenson, *ACS Catal.*, 2014, **4**, 703–716; (*c*) A. Studer and D. P. Curran, *Angew. Chem., Int. Ed.*, 2016, **55**, 58–102; (*d*) M. Yan, J. C. Lo, J. T. Edwards and P. S. Baran, *J. Am. Chem. Soc.*, 2016, **138**, 12692–12714; (*e*) A. J. Clark, *Eur. J. Org. Chem.*, 2016, 2231–2243; (*f*) H. Miyabe, A. Kawashima, E. Yoshioka and S. Kohtani, *Chem. – Eur. J.*, 2017, **23**, 6225–6236; (*g*) M. P. Plesniak, H.-M. Huang and D. J. Procter, *Nat. Rev. Chem.*, 2017, **1**, 0077; (*h*) H. Yi, G. Zhang, H. Wang, Z. Huang, J. Wang, A. K. Singh and A. Lei, *Chem. Rev.*, 2017, **117**, 9016–9085; (*i*) H. Miyabe, *Eur. J. Org. Chem.*, 2017, 3302–3310.

32 A. J. Fry, *Synthetic organic electrochemistry*, John Wiley & Sons, New York, 1989.

33 Throughout this review redox potentials are given in volts (V) *versus* the saturated calomel electrode (SCE). Occasionally the referenced potentials have been determined using different electrodes and have been converted to SCE for the purpose of comparison. For conversion constants, see: V. V. Pavlishchuk and A. W. Addison, *Inorg. Chim. Acta*, 2000, **298**, 97–102.

34 M. Faraday, *Ann. Phys.*, 1834, **109**, 433–451.

35 (*a*) H. Kolbe, *Ann. Chem. Pharm.*, 1848, **64**, 339–341; (*b*) H. Kolbe, *Ann. Chem. Pharm.*, 1849, **69**, 257–294.

36 For reviews on the Kolbe electrolysis, see: (*a*) A. K. Vijh and B. E. Conway, *Chem. Rev.*, 1967, **67**, 623–664; (*b*) H.-J. Schäfer, *Top. Curr. Chem.*, 1990, **152**, 91–151; (*c*) H. J. Schäfer, *Chem. Phys. Lipids*, 1979, **24**, 321–333.

37 For synthetic applications of the Kolbe electrolysis, see: (*a*) J. Knolle and H. J. Schäfer, *Angew. Chem., Int. Ed. Engl.*, 1975, **14**, 758; (*b*) H. Klünenberg and H. J. Schäfer, *Angew. Chem., Int. Ed. Engl.*, 1978, **17**, 47–48; (*c*) U. Jensen and H. J. Schäfer, *Chem. Ber.*, 1981, **114**, 292–297; (*d*) L. Becking and H. J. Schäfer, *Tetrahedron Lett.*, 1988, **29**, 2797–2800; (*e*) J. Weiguny and H. J. Schäfer, *Liebigs Ann. Chem.*, 1994, 235–242; (*f*) A. Matzeit, H. J. Schäfer and C. Amatore, *Synthesis*, 1995, 1432–1444.

38 D. Pletcher, *Rev. Port. Quim.*, 1985, **27**, 449–458.

39 C. A. C. Sequeira and D. M. F. Santos, *J. Braz. Chem. Soc.*, 2009, **20**, 387–406.

40 D. S. P. Cardoso, B. Šljukić, D. M. F. Santos and C. A. C. Sequeira, *Org. Process Res. Dev.*, 2017, **21**, 1213–1226.

41 For a discussion on the historical advances in organic electrochemistry, see: H. Lund, *J. Electrochem. Soc.*, 2002, **149**, S21–S33.

42 For reviews and applications of indirect electrolysis reactions, see: (*a*) E. Steckhan, *Angew. Chem., Int. Ed. Engl.*,

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

1986, **28**, 683–701; (*b*) E. Steckhan, *Top. Curr. Chem.*, 1987, **142**, 1–69; (*c*) Y. N. Ogibin, M. N. Elinson and G. I. Nikishin, *Russ. Chem. Rev.*, 2009, **78**, 89–140; (*d*) R. Francke and R. D. Little, *Chem. Soc. Rev.*, 2014, **43**, 2492–2521.

43 F. Fichter, *Organische Elektrochemie*, Steinkopf, Dresden, 1942, pp. 13 and 75.

44 C. Jackson and A. T. Kuhn, *Industrial electrochemical processes*, Elsevier, Amsterdam, 1971, pp. 513–519.

45 For early examples, see: (*a*) E. T. Seo, R. F. Nelson, J. M. Fritsch, L. S. Marcoux, D. W. Leedy and R. N. Adams, *J. Am. Chem. Soc.*, 1966, **88**, 3498–3503; (*b*) L. Papouchado, R. N. Adams and S. W. Feldberg, *J. Electroanal. Chem. Interfacial Electrochem.*, 1969, **21**, 408–410; (*c*) W. Schmidt and E. Steckhan, *J. Electroanal. Chem.*, 1978, **89**, 215–220; (*d*) W. Schmidt and E. Steckhan, *Angew. Chem., Int. Ed. Engl.*, 1978, **17**, 673–674.

46 For seminal work, see: M. F. Semmelhack, C. S. Chou and D. A. Cortes, *J. Am. Chem. Soc.*, 1983, **105**, 4492–4494.

47 For recent applications of N-oxyl radicals as redox mediators, see: (*a*) M. Rafiee, K. C. Miles and S. S. Stahl, *J. Am. Chem. Soc.*, 2015, **137**, 14751–14757; (*b*) D. P. Hickey, D. A. Schiedler, I. Matanovic, P. V. Doan, P. Atanassov, S. D. Minteer and M. S. Sigman, *J. Am. Chem. Soc.*, 2015, **137**, 16179–16186; (*c*) I. Bosque, G. Magallanes, M. Rigoulet, M. D. Kärkäs and C. R. J. Stephenson, *ACS Cent. Sci.*, 2017, **3**, 621–628; (*d*) T. Li, Y. Cao, J. He and C. P. Berlinguette, *ACS Cent. Sci.*, 2017, **3**, 778–783.

48 By definition, overpotential is the additional potential (beyond the thermodynamic requirement) needed to drive a reaction at a certain rate.

49 For seminal work, see: (*a*) J.-i. Yoshida, T. Murata and S. Isoe, *Tetrahedron Lett.*, 1986, **27**, 3373–3376; (*b*) J.-i. Yoshida and S. Isoe, *Tetrahedron Lett.*, 1987, **28**, 6621–6624.

50 For reviews on the use of electroauxiliaries, see: (*a*) J.-i. Yoshida and K. Nishiwaki, *J. Chem. Soc., Dalton Trans.*, 1998, 2589–2596; (*b*) J.-i. Yoshida, *Top. Curr. Chem.*, 1994, **170**, 39–81.

51 For representative examples, see: (*a*) F. Tang and K. D. Moeller, *Tetrahedron*, 2009, **65**, 10863–10875; (*b*) J. M. Campbell, J. A. Smith, L. Gonzalez and K. D. Moeller, *Tetrahedron Lett.*, 2015, **56**, 3595–3599.

52 For selected examples, see: (*a*) J.-i. Yoshida, K. Takada, Y. Ishichi and S. Isoe, *J. Chem. Soc., Chem. Commun.*, 1994, 2361–2362; (*b*) M. Jinno, Y. Kitano, M. Tada and K. Chiba, *Org. Lett.*, 1999, **1**, 435–438; (*c*) J.-i. Yoshida, M. Watanabe, H. Toshioka, M. Imagawa and S. Suga, *J. Electroanal. Chem.*, 2001, **507**, 55–65.

53 For representative examples, see: (*a*) S. Suga, M. Watanabe and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2002, **124**, 14824–14825; (*b*) D. Kesselring, K. Maurer and K. D. Moeller, *Org. Lett.*, 2008, **10**, 2501–2504.

54 For selected examples, see: (*a*) J.-i. Yoshida, M. Sugawara and N. Kise, *Tetrahedron Lett.*, 1996, **37**, 3157–3160; (*b*) S. Suzuki, K. Matsumoto, K. Kawamura, S. Suga and J.-i. Yoshida, *Org. Lett.*, 2004, **6**, 3755–3758.

55 A. Kirste, B. Elsler, G. Schnakenburg and S. R. Waldvogel, *J. Am. Chem. Soc.*, 2012, **134**, 3571–3576.

56 T. Morofuji, A. Shimizu and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2013, **135**, 5000–5003.

57 For reviews on the "cation pool" method, see: (*a*) J.-i. Yoshida and S. Suga, *Chem. – Eur. J.*, 2002, **8**, 2650–2658; (*b*) K. Matsumoto, S. Suga and J.-i. Yoshida, *Org. Biomol. Chem.*, 2011, **9**, 2586–2596; (*c*) J.-i. Yoshida, A. Shimizu, Y. Ashikari, T. Morofuji, R. Hayashi, T. Nokami and A. Nagaki, *Bull. Chem. Soc. Jpn.*, 2015, **88**, 763–775; (*d*) J.-i. Yoshida, A. Shimizu and R. Hayashi, *Chem. Rev.*, 2018, **118**, 4702–4730.

58 For a review on the preparation of "glycosyl triflate pools", see: T. Nokami, K. Saito and J.-i. Yoshida, *Carbohydr. Res.*, 2012, **363**, 1–6.

59 For a review discussing the sustainability of organic electrosynthesis, see: B. A. Frontana-Uribe, R. D. Little, J. G. Ibanez, A. Palma and R. Vasquez-Medrano, *Green Chem.*, 2010, 12, 2099–2119.

60 J. Utley, *Chem. Soc. Rev.*, 1997, **26**, 157–167.

61 R. Francke, *Beilstein J. Org. Chem.*, 2014, **10**, 2858–2873.

62 For representative reviews on umpolung reactions, see: (*a*) D. Seebach, *Angew. Chem., Int. Ed. Engl.*, 1979, **18**, 239–258; (*b*) A. B. Smith, III and C. M. Adams, *Acc. Chem. Res.*, 2004, **37**, 365–377; (*c*) D. Enders, O. Niemeier and A. Henseler, *Chem. Rev.*, 2007, **107**, 5606–5655; (*d*) X. Bugaut and F. Glorius, *Chem. Soc. Rev.*, 2012, **41**, 3511–3522; (*e*) J. Streuff, *Synthesis*, 2013, 281–307; (*f*) D. M. Flanigan, F. Romanov-Michailidis, N. A. White and T. Rovis, *Chem. Rev.*, 2015, **115**, 9307–9387.

63 For reviews on electrochemical umpolung reactions, see: (*a*) R. D. Little and K. D. Moeller, *Electrochem. Soc. Interface*, 2002, **11**, 36–42; (*b*) F. Tang, C. Chen and K. D. Moeller, *Synthesis*, 2007, 3411–3420; (*c*) K. D. Moeller, *Synlett*, 2009, 1208–1218.

64 For synthetic applications, see: (*a*) D. L. Wright, C. R. Whitehead, E. H. Sessions, I. Ghiviriga and D. A. Frey, *Org. Lett.*, 1999, **1**, 1535–1538; (*b*) S. Duan and K. D. Moeller, *Org. Lett.*, 2001, **3**, 2685–2688.

65 J. Mihelcic and K. D. Moeller, *J. Am. Chem. Soc.*, 2003, **125**, 36–37.

66 J. Mihelcic and K. D. Moeller, *J. Am. Chem. Soc.*, 2004, **126**, 9106–9111.

67 C. C. Hughes, A. K. Miller and D. Trauner, *Org. Lett.*, 2005, **7**, 3425–3428.

68 A. K. Miller, C. C. Hughes, J. J. Kennedy-Smith, S. N. Gradl and D. Trauner, *J. Am. Chem. Soc.*, 2006, **128**, 17057–17062.

69 For a detailed discussion on the ideal electrochemical device, see: M. Yan, Y. Kawamata and P. S. Baran, *Angew. Chem., Int. Ed.*, 2018, **57**, 4149–4155.

70 For a review on elucidating the mechanisms of transition metal-catalyzed reactions using electrochemistry, see: A. Jutand, *Chem. Rev.*, 2008, **108**, 2300–2347.

71 S. Wawzonek, *Science*, 1967, **155**, 39–44.

72 N. L. Weinberg and H. R. Weinberg, *Chem. Rev.*, 1968, **68**, 449–523.

73 R. N. Adams, *Acc. Chem. Res.*, 1969, **2**, 175–180.

74 L. Eberson and K. Nyberg, *Acc. Chem. Res.*, 1973, **6**, 106–112.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

75 T. Shono, *Tetrahedron*, 1984, **40**, 811–850.
76 K. A. Ogawa and A. J. Boydston, *Chem. Lett.*, 2015, **44**, 10–16.
77 K. Chiba and S. Kim, *Electrochemistry*, 2009, **77**, 21–29.
78 K. D. Moeller, *Top. Curr. Chem.*, 1997, **185**, 49–86.
79 K. D. Moeller, *Tetrahedron*, 2000, **56**, 9527–9554.
80 K. D. Moeller, *Electrochem. Soc. Interface*, 2016, **25**, 53–59.
81 R. Feng, J. A. Smith and K. D. Moeller, *Acc. Chem. Res.*, 2017, **50**, 2346–2352.
82 H. J. Schäfer, M. Harenbrock, E. Klocke, M. Plate and A. Weiper-Idelmann, *Pure Appl. Chem.*, 2007, **79**, 2047–2057.
83 H. J. Schäfer, *ChemCatChem*, 2014, **6**, 2792–2795.
84 H. J. Schäfer, *Angew. Chem., Int. Ed. Engl.*, 1981, **20**, 911–934.
85 H. J. Schäfer, *C. R. Chim.*, 2011, **14**, 745–765.
86 J. B. Sperry and D. L. Wright, *Chem. Soc. Rev.*, 2006, **35**, 605–621.
87 J.-i. Yoshida, K. Kataoka, R. Horcajada and A. Nagaki, *Chem. Rev.*, 2008, **108**, 2265–2299.
88 E. J. Horn, B. R. Rosen and P. S. Baran, *ACS Cent. Sci.*, 2016, **2**, 302–308.
89 During the preparation of this review the following review articles on organic electrochemistry were published: (*a*) M. Yan, Y. Kawamata and P. S. Baran, *Chem. Rev.*, 2017, **117**, 13230–13319; (*b*) Y. Jiang, K. Xu and C. Zeng, *Chem. Rev.*, 2018, **118**, 4485–4540; (*c*) K. D. Moeller, *Chem. Rev.*, 2018, **118**, 4817–4833; (*d*) A. Wiebe, T. Gieshoff, S. Möhle, E. Rodrigo, M. Zirbes and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2018, **57**, 5594–5619; (*e*) S. Möhle, M. Zirbes, E. Rodrigo, T. Gieshoff, A. Wiebe and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2018, **57**, 6018–6041; (*f*) Q.-L. Yang, P. Fang and T.-S. Mei, *Chin. J. Chem.*, 2018, **36**, 338–352.
90 For recent reviews on C–H activation/functionalization, see: (*a*) T. Gensch, M. N. Hopkinson, F. Glorius and J. Wencel-Delord, *Chem. Soc. Rev.*, 2016, **45**, 2900–2936; (*b*) M. Moselage, J. Li and L. Ackermann, *ACS Catal.*, 2016, **6**, 498–525; (*c*) X.-S. Xue, P. Ji, B. Zhou and J.-P. Cheng, *Chem. Rev.*, 2017, **117**, 8622–8648; (*d*) J. He, M. Wasa, K. S. L. Chan, Q. Shao and J.-Q. Yu, *Chem. Rev.*, 2017, **117**, 8754–8786; (*e*) J. R. Hummel, J. A. Boerth and J. A. Ellman, *Chem. Rev.*, 2017, **117**, 9163–9227; (*f*) M. Parasram and V. Gevorgyan, *Acc. Chem. Res.*, 2017, **50**, 2038–2053; (*g*) R. Das, G. S. Kumar and M. Kapur, *Eur. J. Org. Chem.*, 2017, 5439–5459; (*h*) M. T. Mihai, G. R. Genov and R. J. Phipps, *Chem. Soc. Rev.*, 2018, **47**, 149–171.
91 For a review on late stage functionalization of drug-like molecules, see: T. Cernak, K. D. Dykstra, S. Tyagarajan, P. Vachal and S. W. Krska, *Chem. Soc. Rev.*, 2016, **45**, 546–576.
92 T. Shono, H. Hamaguchi and Y. Matsumura, *J. Am. Chem. Soc.*, 1975, **97**, 4264–4268.
93 T. Shono, Y. Matsumura, K. Tsubata, Y. Sugihara, S.-i. Yamane, T. Kanazawa and T. Aoki, *J. Am. Chem. Soc.*, 1982, **104**, 6697–6703.
94 For early contributions, see: (*a*) S. D. Ross, M. Finkelstein and R. C. Petersen, *J. Org. Chem.*, 1966, **31**, 128–133; (*b*) S. D. Ross, M. Finkelstein and R. C. Petersen, *J. Am. Chem. Soc.*, 1966, **88**, 4657–4660.
95 For reviews on Shono-type anodic oxidations, see: (*a*) T. Shono, *Top. Curr. Chem.*, 1988, **148**, 131–162; (*b*) A. M. Jones and C. E. Banks, *Beilstein J. Org. Chem.*, 2014, **10**, 3056–3072.
96 P. Alfonso-Súarez, A. V. Kolliopoulos, J. P. Smith, C. E. Banks and A. M. Jones, *Tetrahedron Lett.*, 2015, **56**, 6863–6867.
97 P. D. Palasz, J. H. P. Utley and J. D. Hardstone, *J. Chem. Soc., Perkin Trans. 2*, 1984, 807–813.
98 T. Shone, Y. Matsumura, K. Uchida and H. Kobayashi, *J. Org. Chem.*, 1985, **50**, 3243–3245.
99 T. Shono, Y. Matsumura and K. Tsubata, *Org. Synth.*, 1985, **63**, 206–211.
100 T. Shono, Y. Matsumura and K. Inoue, *J. Org. Chem.*, 1983, **48**, 1388–1389.
101 M. Thaning and L.-G. Wistrand, *Acta Chem. Scand.*, 1989, **43**, 290–295.
102 K.-D. Ginzel, P. Brungs and E. Steckhan, *Tetrahedron*, 1989, **45**, 1691–1701.
103 K. D. Moeller, P. W. Wang, S. Tarazi, M. R. Marzabadi and P. L. Wong, *J. Org. Chem.*, 1991, **56**, 1058–1067.
104 M. Thaning and L.-G. Wistrand, *Acta Chem. Scand.*, 1992, **46**, 194–199.
105 G. Butora, J. W. Reed, T. Hudlicky, L. E. Brammer, Jr., P. I. Higgs, D. P. Simmons and N. E. Heard, *J. Am. Chem. Soc.*, 1997, **119**, 7694–7701.
106 M. K. S. Vink, C. A. Schortinghuis, J. Luten, J. H. van Maarseveen, H. E. Schoemaker, H. Hiemstra and F. P. J. T. Rutjes, *J. Org. Chem.*, 2002, **67**, 7869–7871.
107 M. David and H. Dhimane, *Synlett*, 2004, 1029–1033.
108 M. R. Faust, G. Höfner, J. Pabel and K. T. Wanner, *Eur. J. Med. Chem.*, 2010, **45**, 2453–2466.
109 T. Golub and J. Y. Becker, *Electrochim. Acta*, 2015, **173**, 408–415.
110 For a review on strategies for the direct α-functionalization of saturated cyclic amines, see: E. A. Mitchell, A. Peschiulli, N. Lefevre, L. Meerpoel and B. U. W. Maes, *Chem. – Eur. J.*, 2012, **18**, 10092–10142.
111 T. Shono, Y. Matsumura, O. Onomura and M. Sato, *J. Org. Chem.*, 1988, **53**, 4118–4121.
112 Y. Demizu, H. Shiigi, H. Mori, K. Matsumoto and O. Onomura, *Tetrahedron: Asymmetry*, 2008, **19**, 2659–2665.
113 X. Wang, J. Li, R. A. Saporito and N. Toyooka, *Tetrahedron*, 2013, **69**, 10311–10315.
114 S. S. Libendi, Y. Demizu, Y. Matsumura and O. Onomura, *Tetrahedron*, 2008, **64**, 3935–3942.
115 S. S. Libendi, Y. Demizu and O. Onomura, *Org. Biomol. Chem.*, 2009, **7**, 351–356.
116 A. D. Tereshchenko, J. S. Myronchuk, L. D. Leitchenko, I. V. Knysh, G. O. Tokmakova, O. O. Litsis, A. Tolmachev, K. Liubchak and P. Mykhailiuk, *Tetrahedron*, 2017, **73**, 750–757.
117 Y. Cao, K. Suzuki, T. Tajima and T. Fuchigami, *Tetrahedron*, 2005, **61**, 6854–6859.
118 T. Shono, Y. Matsumura, K. Tsubata and J. Takata, *Chem. Lett.*, 1981, 1121–1124.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

119 T. Shono, Y. Matsumura and K. Tsubata, *Tetrahedron Lett.*, 1981, **22**, 2411–2412.
120 T. Shono, Y. Matsumura and K. Tsubata, *J. Am. Chem. Soc.*, 1981, **103**, 1172–1176.
121 T. Shono, Y. Matsumura and K. Tsubata, *Tetrahedron Lett.*, 1981, **22**, 3249–3252.
122 For synthetic applications of N-acyl iminium ions, see: (*a*) B. E. Maryanoff, H.-C. Zhang, J. H. Cohen, I. J. Turchi and C. A. Maryanoff, *Chem. Rev.*, 2004, **104**, 1431–1628; (*b*) P. Wu and T. E. Nielsen, *Chem. Rev.*, 2017, **117**, 7811–7856.
123 A. Papadopoulos, J. Heyer, K.-D. Ginzel and E. Steckhan, *Chem. Ber.*, 1989, **122**, 2159–2164.
124 A. Papadopoulos, B. Lewall, E. Steckhan, K.-D. Ginzel, F. Knoch and M. Nieger, *Tetrahedron*, 1991, **47**, 563–572.
125 G. Kardassis, P. Brungs and E. Steckhan, *Tetrahedron*, 1998, **54**, 3471–3478.
126 G. Kardassis, P. Brungs, C. Nothhelfer and E. Steckhan, *Tetrahedron*, 1998, **54**, 3479–3488.
127 Y. Kanda, O. Onomura, T. Maki and Y. Matsumura, *Chirality*, 2003, **15**, 89–94.
128 K. Schierle-Arndt, D. Kolter, K. Danielmeier and E. Steckhan, *Eur. J. Org. Chem.*, 2001, 2425–2433.
129 M. Lennartz, M. Sadakane and E. Steckhan, *Tetrahedron*, 1999, **55**, 14407–14420.
130 M. Lennartz and E. Steckhan, *Synlett*, 2000, 319–322.
131 M. Lennartz and E. Steckhan, *Tetrahedron*, 2001, **57**, 675–680.
132 D.-S. Lee, *Tetrahedron: Asymmetry*, 2009, **20**, 2014–2020.
133 M. G. M. D'Oca, R. A. Pilli and I. Vencato, *Tetrahedron Lett.*, 2000, **41**, 9709–9712.
134 Y. Matsumura, Y. Kanda, K. Shirai, O. Onomura and T. Maki, *Tetrahedron*, 2000, **56**, 7411–7422.
135 M. G. M. D'Oca, R. A. Pilli, V. L. Pardini, D. Curi and F. C. M. Comninos, *J. Braz. Chem. Soc.*, 2001, **12**, 507–513.
136 M. Zelgert, M. Nieger, M. Lennartz and E. Steckhan, *Tetrahedron*, 2002, **58**, 2641–2646.
137 E. Sierecki, S. Turcaud, T. Martens and J. Royer, *Synthesis*, 2006, 3199–3208.
138 E. Sierecki, G. Errasti, T. Martens and J. Royer, *Tetrahedron*, 2010, **66**, 10002–10007.
139 S. Turcaud, T. Martens, E. Sierecki, J. Pérard-Viret and J. Royer, *Tetrahedron Lett.*, 2005, **46**, 5131–5134.
140 M. Gong and J.-M. Huang, *Chem. – Eur. J.*, 2016, **22**, 14293–14296.
141 *Organic electrochemistry, fourth edition, revised and expanded*, ed. O. Hammerich and H. Lund, Marcel Dekker, Inc., New York, 4th edn, 2001.
142 (*a*) B. Wermeckes and F. Beck, *Electrochim. Acta*, 1985, **30**, 1491–1500; (*b*) K. Danielmeier, K. Schierle and E. Steckhan, *Tetrahedron*, 1996, **52**, 9743–9754; (*c*) H. Senboku, K. Nagakura, T. Fukuhara and S. Hara, *Tetrahedron*, 2015, **71**, 3850–3856.
143 D. Baba and T. Fuchigami, *Tetrahedron Lett.*, 2003, **44**, 3133–3136.
144 S. Andreades and E. W. Zahnow, *J. Am. Chem. Soc.*, 1969, **91**, 4181–4190.
145 T. Chiba and Y. Takata, *J. Org. Chem.*, 1977, **42**, 2973–2977.
146 I. Gallardo and N. Vilà, *J. Org. Chem.*, 2008, **73**, 6647–6656.
147 An electrochemical approach to afford phosphinic amides has also been developed, see: Y. Wang, P. Qian, J.-H. Su, Y. Li, M. Bi, Z. Zha and Z. Wang, *Green Chem.*, 2017, **19**, 4769–4773.
148 For early studies on electrochemical oxidation of aliphatic amines, see: (*a*) R. F. Dapo and C. K. Mann, *Anal. Chem.*, 1963, **35**, 677–680; (*b*) C. K. Mann, *Anal. Chem.*, 1964, **36**, 2424–2426; (*c*) K. K. Barnes and C. K. Mann, *J. Org. Chem.*, 1967, **32**, 1474–1479; (*d*) P. J. Smith and C. K. Mann, *J. Org. Chem.*, 1968, **33**, 316–317.
149 I. Gallardo and N. Vilà, *J. Org. Chem.*, 2010, **75**, 680–689.
150 O. Onomura, Y. Ishida, T. Maki, D. Minato, Y. Demizu and Y. Matsumura, *Electrochemistry*, 2006, **74**, 645–648.
151 V. H. Vu, F. Louafi, N. Girard, R. Marion, T. Roisnel, V. Dorcet and J.-P. Hurvois, *J. Org. Chem.*, 2014, **79**, 3358–3373.
152 L. Zhang, J.-H. Su, S. Wang, C. Wan, Z. Zha, J. Du and Z. Wang, *Chem. Commun.*, 2011, **47**, 5488–5490.
153 K. D. Moeller, S. Tarazi and M. R. Marzabadi, *Tetrahedron Lett.*, 1989, **30**, 1213–1216.
154 K. D. Moeller, S. L. Rothfus and P. L. Wang, *Tetrahedron*, 1991, **47**, 583–592.
155 M. A. Endoma, G. Butora, C. D. Claeboe, T. Hudlicky and K. A. Abboud, *Tetrahedron Lett.*, 1997, **38**, 8833–8836.
156 M. Okimoto, T. Yoshida, M. Hoshi, K. Hattori, M. Komata, K. Numata and K. Tomozawa, *Heterocycles*, 2006, **68**, 2563–2570.
157 M. Okimoto, T. Yoshida, M. Hoshi, K. Hattori, M. Komata, K. Numata and K. Tomozawa, *Aust. J. Chem.*, 2007, **60**, 236–242.
158 M. Okimoto, K. Ohashi, H. Yamamori, S. Nishikawa, M. Hoshi and T. Yoshida, *Synthesis*, 2012, 1315–1322.
159 For electrochemical oxidation of benzylic amines to imines, see: M. Okimoto, Y. Takahashi, K. Numata, Y. Nagata and G. Sasaki, *Synth. Commun.*, 2005, **35**, 1989–1995.
160 For anodic cyanation of tetrahydroisoquinolines, see: F. Louafi, J.-P. Hurvois, A. Chibani and T. Roisnel, *J. Org. Chem.*, 2010, **75**, 5721–5724.
161 Q. Yang, L. Zhang, C. Ye, S. Luo, L.-Z. Wu and C.-H. Tung, *Angew. Chem., Int. Ed.*, 2017, **56**, 3694–3698.
162 N. Fu, L. Li, Q. Yang and S. Luo, *Org. Lett.*, 2017, **19**, 2122–2125.
163 O. Baslé, N. Borduas, P. Dubois, J. M. Chapuzet, T.-H. Chan, J. Lessard and C.-J. Li, *Chem. – Eur. J.*, 2010, **16**, 8162–8166.
164 C. Li, C.-C. Zeng, L.-M. Hu, F.-L. Yang, S. J. Yoo and R. D. Little, *Electrochim. Acta*, 2013, **114**, 560–566.
165 For acceptorless dehydrogenation of N-heterocycles using TEMPO as mediator, see: Y. Wu, H. Yi and A. Lei, *ACS Catal.*, 2018, **8**, 1192–1196.
166 Y. Kashiwagi and J.-i. Anzai, *Chem. Pharm. Bull.*, 2001, **49**, 324–326.
167 J.-i. Yoshida, S. Suga, S. Suzuki, N. Kinomura, A. Yamamoto and K. Fujiwara, *J. Am. Chem. Soc.*, 1999, **121**, 9546–9549.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

168 S. Suga, M. Okajima and J.-i. Yoshida, *Tetrahedron Lett.*, 2001, **42**, 2173–2176.
169 S. Suga, T. Nishida, D. Yamada, A. Nagaki and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2004, **126**, 14338–14339.
170 T. Maruyama, Y. Mizuno, I. Shimizu, S. Suga and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2007, **129**, 1902–1903.
171 S. Suga, M. Okajima, K. Fujiwara and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2001, **123**, 7941–7942.
172 S. Suga, M. Okajima, K. Fujiwara and J.-i. Yoshida, *QSAR Comb. Sci.*, 2005, **24**, 728–741.
173 S. Suga, D. Yamada and J.-i. Yoshida, *Chem. Lett.*, 2010, **39**, 404–406.
174 J. Kuleshova, J. T. Hill-Cousins, P. R. Birkin, R. C. D. Brown, D. Pletcher and T. J. Underwood, *Electrochim. Acta*, 2011, **56**, 4322–4326.
175 K. Watts, W. Gattrell and T. Wirth, *Beilstein J. Org. Chem.*, 2011, **7**, 1108–1114.
176 B. Giese and J. Dupuis, *Angew. Chem., Int. Ed. Engl.*, 1983, **22**, 622–623.
177 For reviews on Giese-type reactions, see: (*a*) B. Giese, *Angew. Chem., Int. Ed. Engl.*, 1983, **22**, 753–764; (*b*) W. Zhang, *Tetrahedron*, 2001, **57**, 7237–7262; (*c*) G. S. C. Srikanth and S. L. Castle, *Tetrahedron*, 2005, **61**, 10377–10441.
178 S. Suga, S. Suzuki and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2002, **124**, 30–31.
179 S. Suga, S. Suzuki, T. Maruyama and J.-i. Yoshida, *Bull. Chem. Soc. Jpn.*, 2004, **77**, 1545–1554.
180 D. Horii, T. Fuchigami and M. Atobe, *J. Am. Chem. Soc.*, 2007, **129**, 11692–11693.
181 D. Horii, F. Amemiya, T. Fuchigami and M. Atobe, *Chem. – Eur. J.*, 2008, **14**, 10382–10387.
182 T. Tajima and A. Nakajima, *J. Am. Chem. Soc.*, 2008, **130**, 10496–10497.
183 For seminal studies on the concept of "site isolation", see: (*a*) B. J. Cohen, M. A. Kraus and A. Patchornik, *J. Am. Chem. Soc.*, 1977, **99**, 4165–4167; (*b*) B. J. Cohen, M. A. Kraus and A. Patchornik, *J. Am. Chem. Soc.*, 1981, **103**, 7620–7629.
184 R. Asami, M. Atobe and T. Fuchigami, *J. Am. Chem. Soc.*, 2005, **127**, 13160–13161.
185 R. Asami, T. Fuchigami and M. Atobe, *Langmuir*, 2006, **22**, 10258–10263.
186 R. Asami, T. Fuchigami and M. Atobe, *Chem. Commun.*, 2008, 244–246.
187 R. Asami, T. Fuchigami and M. Atobe, *Org. Biomol. Chem.*, 2008, **6**, 1938–1943.
188 M. Atobe, S. Ikari, K. Nakabayashi, F. Amemiya and T. Fuchigami, *Langmuir*, 2010, **26**, 9111–9115.
189 T. Tajima and T. Fuchigami, *Angew. Chem., Int. Ed.*, 2005, **44**, 4760–4763.
190 H. Kurihara, T. Tajima and T. Fuchigami, *Electrochemistry*, 2006, **74**, 615–617.
191 T. Tajima, H. Kurihara and T. Fuchigami, *J. Am. Chem. Soc.*, 2007, **129**, 6680–6681.
192 H. Kurihara, T. Fuchigami and T. Tajima, *J. Org. Chem.*, 2008, **73**, 6888–6890.
193 T. Tajima and H. Kurihara, *Chem. Commun.*, 2008, 5167–5169.
194 T. Tajima, S. Ishino and H. Kurihara, *Chem. Lett.*, 2008, **37**, 1036–1037.
195 T. Tajima and A. Nakajima, *Chem. Lett.*, 2009, **38**, 160–161.
196 T. Tajima, H. Kurihara, S. Shimizu and H. Tateno, *Electrochemistry*, 2013, **81**, 353–355.
197 T. Tajima and T. Fuchigami, *J. Am. Chem. Soc.*, 2005, **127**, 2848–2849.
198 T. Tajima and T. Fuchigami, *Chem. – Eur. J.*, 2005, **11**, 6192–6196.
199 K. Suds, K. Hotoda, J.-i. Watanabe, K. Shiozawa and T. Takanami, *J. Chem. Soc., Perkin Trans. 1*, 1992, 1283–1284.
200 K. Suda, K. Hotoda, F. Iemuroe and T. Takanami, *J. Chem. Soc., Perkin Trans. 1*, 1993, 1553–1555.
201 E. Le Gall, J.-P. Hurvois and S. Sinbandhit, *Eur. J. Org. Chem.*, 1999, 2645–2653.
202 S. Suga, M. Watanabe and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2002, **124**, 14824–14825.
203 T. Kamada and A. Oku, *J. Chem. Soc., Perkin Trans. 1*, 2002, 1105–1110.
204 S. Suga, A. Nagaki, Y. Tsutsui and J.-i. Yoshida, *Org. Lett.*, 2003, **5**, 945–947.
205 A. Nagaki, K. Kawamura, S. Suga, T. Ando, M. Sawamoto and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2004, **126**, 14702–14703.
206 S. Suga, Y. Kageyama, G. Babu, K. Itami and J.-i. Yoshida, *Org. Lett.*, 2004, **6**, 2709–2711.
207 T. Maruyama, S. Suga and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2005, **127**, 7324–7325.
208 A. Nagaki, M. Togai, S. Suga, N. Aoki, K. Mae and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2005, **127**, 11666–11675.
209 S. Suga, Y. Tsutsui, A. Nagaki and J.-i. Yoshida, *Bull. Chem. Soc. Jpn.*, 2005, **78**, 1206–1217.
210 T. Maruyama, S. Suga and J.-i. Yoshida, *Tetrahedron*, 2006, **62**, 6519–6525.
211 S. Suga, M. Watanabe, C.-H. Song and J.-i. Yoshida, *Electrochemistry*, 2006, **74**, 672–679.
212 A. Nagaki, T. Iwasaki, K. Kawamura, D. Yamada, S. Suga, T. Ando, M. Sawamoto and J.-i. Yoshida, *Chem. – Asian J.*, 2008, **3**, 1558–1567.
213 Y. Ashikari, Y. Kiuchi, T. Takeuchi, K. Ueoka, S. Suga and J.-i. Yoshida, *Chem. Lett.*, 2014, **43**, 210–212.
214 T. Shoji, S. Kim, K. Yamamoto, T. Kawai, Y. Okada and K. Chiba, *Org. Lett.*, 2014, **16**, 6404–6407.
215 M. Sugawara, K. Mori and J.-i. Yoshida, *Electrochim. Acta*, 1997, **42**, 1995–2003.
216 T. Fuchigami, M. Tetsu, T. Tajima and H. Ishii, *Synlett*, 2001, 1269–1271.
217 S. Kim, K. Hayashi, Y. Kitano, M. Tada and K. Chiba, *Org. Lett.*, 2002, **4**, 3735–3737.
218 K. J. Frankowski, R. Liu, G. L. Milligan, K. D. Moeller and J. Aubé, *Angew. Chem., Int. Ed.*, 2015, **54**, 10555–10558.
219 For leading reviews on azanucleosides and azanucleotides, see: (*a*) C. Nájera and J. M. Sansano, *Org. Biomol. Chem.*, 2009, **7**, 4567–4581; (*b*) G. Romeo, U. Chiacchio, A. Corsaro and P. Merino, *Chem. Rev.*, 2010, **110**, 3337–3370; (*c*) B. L. Stocker, E. M. Dangerfield, A. L. Win-Mason, G. W. Haslett

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

and M. S. M. Timmer, *Eur. J. Org. Chem.*, 2010, 1615–1637; (*d*) D. Hernández and A. Boto, *Eur. J. Org. Chem.*, 2014, 2201–2220.
220 S. Furukubo, N. Moriyama, O. Onomura and Y. Matsumura, *Tetrahedron Lett.*, 2004, **45**, 8177–8181.
221 S. Kim, T. Shoji, Y. Kitano and K. Chiba, *Chem. Commun.*, 2013, **49**, 6525–6527.
222 T. Shoji, S. Haraya, S. Kim and K. Chiba, *Electrochim. Acta*, 2016, **200**, 290–295.
223 T. Shoji, S. Kim and K. Chiba, *Angew. Chem., Int. Ed.*, 2017, **56**, 4011–4014.
224 K. Bodmann, T. Bug, S. Steinbeisser, R. Kreuder and O. Reiser, *Tetrahedron Lett.*, 2006, **47**, 2061–2064.
225 F. Cornille, U. Slomczynska, M. L. Smythe, D. D. Beusen, K. D. Moeller and G. R. Marshall, *J. Am. Chem. Soc.*, 1995, **117**, 909–917.
226 U. Slomczynska, D. K. Chalmers, F. Cornille, M. L. Smythe, D. D. Beusen, K. D. Moeller and G. R. Marshall, *J. Org. Chem.*, 1996, **61**, 1198–1204.
227 W. Li and K. D. Moeller, *J. Am. Chem. Soc.*, 1996, **118**, 10106–10112.
228 W. Chu and K. D. Moeller, *Tetrahedron Lett.*, 1999, **40**, 7939–7943.
229 L. M. Beal, B. Liu, W. Chu and K. D. Moeller, *Tetrahedron*, 2000, **56**, 10113–10125.
230 H. Sun and K. D. Moeller, *Org. Lett.*, 2002, **4**, 1547–1550.
231 H. Sun and K. D. Moeller, *Org. Lett.*, 2003, **5**, 3189–3192.
232 H. Sun, C. Martin, D. Kesselring, R. Keller and K. D. Moeller, *J. Am. Chem. Soc.*, 2006, **128**, 13761–13771.
233 For selected reviews on synthetic approaches towards peptidomimetic design and applications, see: (*a*) A. Giannis and T. Kolter, *Angew. Chem., Int. Ed. Engl.*, 1993, **32**, 1244–1267; (*b*) I. Avan, C. D. Hall and A. R. Katritzky, *Chem. Soc. Rev.*, 2014, **43**, 3575–3594; (*c*) M. Pelay-Gimeno, A. Glas, O. Koch and T. N. Grossmann, *Angew. Chem., Int. Ed.*, 2015, **54**, 8896–8927; (*d*) A. K. Yudin, *Chem. Sci.*, 2015, **6**, 30–49; (*e*) D. E. Scott, A. R. Bayly, C. Abell and J. Skidmore, *Nat. Rev. Drug Discovery*, 2016, **15**, 533–550; (*f*) O. Van der Poorten, A. Knuhtsen, D. Sejer Pedersen, S. Ballet and D. Tourwé, *J. Med. Chem.*, 2016, **59**, 10865–10890; (*g*) A. Mizuno, K. Matsui and S. Shuto, *Chem. – Eur. J.*, 2017, **23**, 14394–14409; (*h*) N. Qvit, S. J. S. Rubin, T. J. Urban, D. Mochly-Rosen and E. R. Gross, *Drug Discovery Today*, 2017, **22**, 454–462.
234 For a review on radical reactions in the synthesis of α-amino acids and related derivatives, see: C. J. Easton, *Chem. Rev.*, 1997, **97**, 53–82.
235 For synthesis and applications of silicon-containing α-amino acids, see: M. Mortensen, R. Husmann, E. Veri and C. Bolm, *Chem. Soc. Rev.*, 2009, **38**, 1002–1010.
236 C. Reuter, P. Huy, J.-M. Neudörfl, R. Kühne and H.-G. Schmalz, *Chem. – Eur. J.*, 2011, **17**, 12037–12044.
237 T. Shono, Y. Matsumura and T. Kanazawa, *Tetrahedron Lett.*, 1983, **24**, 1259–1262.
238 M. A. Kabeshov, B. Musio, P. R. D. Murray, D. L. Browne and S. V. Ley, *Org. Lett.*, 2014, **16**, 4618–4621.
239 F. Louafi, J. Moreau, S. Shahane, S. Golhen, T. Roisnel, S. Sinbandhit and J.-P. Hurvois, *J. Org. Chem.*, 2011, **76**, 9720–9732.
240 N. Girard, J.-P. Hurvois, C. Moinet and L. Toupet, *Eur. J. Org. Chem.*, 2005, 2269–2280.
241 N. Shankaraiah, R. A. Pilli and L. S. Santos, *Tetrahedron Lett.*, 2008, **49**, 5098–5100.
242 N. Shankaraiah, W. A. da Silva, C. K. Z. Andrade and L. S. Santos, *Tetrahedron Lett.*, 2008, **49**, 4289–4291.
243 Y. Mirabal-Gallardo, J. Piérola, N. Shankaraiah and L. S. Santos, *Tetrahedron Lett.*, 2012, **53**, 3672–3675.
244 T.-S. Kam, T.-M. Lim and G.-H. Tan, *J. Chem. Soc., Perkin Trans. 1*, 2001, 1594–1604.
245 K. D. Moeller and P. L. Wong, *Bioorg. Med. Chem. Lett.*, 1992, **2**, 739–742.
246 A. Paci, T. Martens and J. Royer, *Bioorg. Med. Chem. Lett.*, 2001, **11**, 1347–1349.
247 P. J. Smith and C. K. Mann, *J. Org. Chem.*, 1969, **34**, 1821–1826.
248 L. C. Portis, V. V. Bhat and C. K. Mann, *J. Org. Chem.*, 1970, **35**, 2175–2178.
249 S. D. Ross, *Tetrahedron Lett.*, 1973, **14**, 1237–1240.
250 L. R. Hall, R. T. Iwamoto and R. P. Hanzlik, *J. Org. Chem.*, 1989, **54**, 2446–2451.
251 P. G. Kirira, M. Kuriyama and O. Onomura, *Chem. – Eur. J.*, 2010, **16**, 3970–3982.
252 E. P. Gillis, K. J. Eastman, M. D. Hill, D. J. Donnelly and N. A. Meanwell, *J. Med. Chem.*, 2015, **58**, 8315–8359.
253 Y. Zhou, J. Wang, Z. Gu, S. Wang, W. Zhu, J. L. Aceña, V. A. Soloshonok, K. Izawa and H. Liu, *Chem. Rev.*, 2016, **116**, 422–518.
254 D. E. Yerien, S. Bonesi and A. Postigo, *Org. Biomol. Chem.*, 2016, **14**, 8398–8427.
255 P. Jeschke, *ChemBioChem*, 2004, **5**, 570–589.
256 T. Fujiwara and D. O'Hagan, *J. Fluorine Chem.*, 2014, **167**, 16–29.
257 R. Berger, G. Resnati, P. Metrangolo, E. Weber and J. Hulliger, *Chem. Soc. Rev.*, 2011, **40**, 3496–3508.
258 For selected reviews on electrophilic fluorination, see: (*a*) G. S. Lal, G. P. Pez and R. G. Syvret, *Chem. Rev.*, 1996, **96**, 1737–1756; (*b*) S. D. Taylor, C. C. Kotoris and G. Hum, *Tetrahedron*, 1999, **55**, 12431–12477; (*c*) S. Barata-Vallejo, B. Lantaño and A. Postigo, *Chem. – Eur. J.*, 2014, **20**, 16806–16829; (*d*) T. Koike and M. Akita, *J. Fluorine Chem.*, 2014, **167**, 30–36; (*e*) J. Charpentier, N. Früh and A. Togni, *Chem. Rev.*, 2015, **115**, 650–682.
259 For selected reviews on nucleophilic fluorination, see: (*a*) R. P. Singh and J. M. Shreeve, *Synthesis*, 2002, 2561–2578; (*b*) V. V. Grushin, *Acc. Chem. Res.*, 2010, **43**, 160–171; (*c*) C. Hollingworth and V. Gouverneur, *Chem. Commun.*, 2012, **48**, 2929–2942; (*d*) J.-W. Lee, M. T. Oliveira, H. B. Jang, S. Lee, D. Y. Chi, D. W. Kim and C. E. Song, *Chem. Soc. Rev.*, 2016, **45**, 4638–4650; (*e*) S. V. Kohlhepp and T. Gulder, *Chem. Soc. Rev.*, 2016, **45**, 6270–6288; (*f*) S. Liang, G. B. Hammond and B. Xu, *Chem. – Eur. J.*, 2017, **23**, 17850–17861; (*g*) C. N. Neumann and T. Ritter, *Acc. Chem. Res.*, 2017, **50**, 2822–2833.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

260 For selected reviews on radical fluorination, see: (*a*) C. Chatalova-Sazepin, R. Hemelaere, J.-F. Paquin and G. M. Sammis, *Synthesis*, 2015, 2554–2569; (*b*) M. P. Sibi and Y. Landais, *Angew. Chem., Int. Ed.*, 2013, **52**, 3570–3572; (*c*) X. Pan, H. Xia and J. Wu, *Org. Chem. Front.*, 2016, **3**, 1163–1185; (*d*) B. Lantaño and A. Postigo, *Org. Biomol. Chem.*, 2017, **15**, 9954–9973; (*e*) T. Koike and M. Akita, *Chem*, 2018, **4**, 409–437.
261 For general reviews on synthetic approaches for the synthesis of organofluorine compounds, see: (*a*) T. Liang, C. N. Neumann and T. Ritter, *Angew. Chem., Int. Ed.*, 2013, **52**, 8214–8264; (*b*) P. A. Champagne, J. Desroches, J.-D. Hamel, M. Vandamme and J.-F. Paquin, *Chem. Rev.*, 2015, **115**, 9073–9174; (*c*) C. N. Neumann and T. Ritter, *Angew. Chem., Int. Ed.*, 2015, **54**, 3216–3221; (*d*) D. L. Orsi and R. A. Altman, *Chem. Commun.*, 2017, **53**, 7168–7181; (*e*) D. E. Yerien, S. Barata-Vallejo and A. Postigo, *Chem. – Eur. J.*, 2017, **23**, 14676–14701.
262 For a review on electrochemical fluoroalkylation, see: Y. B. Dudkina, M. N. Khrizanforov, T. V. Gryaznova and Y. H. Budnikova, *J. Organomet. Chem.*, 2014, **751**, 301–305.
263 I. L. Knunyants, I. N. Rozhkov, A. V. Bukhtiarov, M. M. Gol'din and R. V. Kudryavtsev, *Izv. Akad. Nauk SSSR, Ser. Khim.*, 1970, 1207–1208.
264 I. N. Rozhkov, A. B. Bukhtiarov, N. D. Kuleshova and I. L. Knunyants, *Dokl. Akad. Nauk SSSR*, 1970, **193**, 1322–1325.
265 I. N. Rozhkov, *Russ. Chem. Rev.*, 1976, **45**, 615–629.
266 K. Momota, M. Morita and Y. Matsuda, *Electrochim. Acta*, 1993, **38**, 619–624.
267 K. Momota, M. Morita and Y. Matsuda, *Electrochim. Acta*, 1993, **38**, 1123–1130.
268 K. Momota, K. Kato, M. Morita and Y. Matsuda, *Electrochim. Acta*, 1994, **39**, 41–49.
269 K. Momota, H. Horio, K. Kato, M. Morita and Y. Matsuda, *Electrochim. Acta*, 1995, **40**, 233–240.
270 K. Momota, K. Kato, M. Morita and Y. Matsuda, *Electrochim. Acta*, 1994, **39**, 681–690.
271 K. Momota, T. Yonezawa, Y. Hayakawa, K. Kato, M. Morita and Y. Matsuda, *J. Appl. Electrochem.*, 1995, **25**, 651–658.
272 K. Momota, K. Mukai, K. Kato and M. Morita, *Electrochim. Acta*, 1998, **43**, 2503–2514.
273 K. Momota, T. Yonezawa, K. Mukai and M. Morita, *J. Fluorine Chem.*, 1998, **87**, 173–178.
274 T. Tajima, H. Ishii and T. Fuchigami, *Electrochem. Commun.*, 2002, **4**, 589–592.
275 N. Ilayaraja and M. Noel, *J. Electroanal. Chem.*, 2010, **638**, 39–45.
276 T. Tajima, H. Ishii and T. Fuchigami, *Electrochem. Commun.*, 2001, **3**, 467–471.
277 T. Tajima, H. Ishii and T. Fuchigami, *Tetrahedron Lett.*, 2001, **42**, 4857–4860.
278 M. R. Shaaban and T. Fuchigami, *Synlett*, 2001, 1644–1646.
279 S. M. Riyadh, H. Ishii and T. Fuchigami, *Tetrahedron*, 2002, **58**, 9273–9278.
280 K. M. Dawood and T. Fuchigami, *J. Org. Chem.*, 2004, **69**, 5302–5306.
281 T. Tajima, A. Nakajima and T. Fuchigami, *J. Org. Chem.*, 2006, **71**, 1436–1441.
282 B. Yin, L. Wang, S. Inagi and T. Fuchigami, *Tetrahedron*, 2010, **66**, 6820–6825.
283 B. Yin, S. Inagi and T. Fuchigami, *Synlett*, 2011, 1313–1317.
284 S. M. Riyadh and T. Fuchigami, *J. Org. Chem.*, 2002, **67**, 9379–9383.
285 M. R. Shaaban and T. Fuchigami, *Tetrahedron Lett.*, 2002, **43**, 273–276.
286 M. R. Shaaban, S. Inagi and T. Fuchigami, *Electrochim. Acta*, 2009, **54**, 2635–2639.
287 T. Fukuhara, M. Sawaguchi and N. Yoneda, *Electrochem. Commun.*, 2000, **2**, 259–261.
288 M. Sawaguchi, T. Fukuhara and N. Yoneda, *J. Electroanal. Chem.*, 2001, **507**, 66–70.
289 T. Fukuhara, Y. Akiyama, N. Yoneda, T. Tada and S. Hara, *Tetrahedron Lett.*, 2002, **43**, 6583–6585.
290 For electrochemical reduction of aromatic systems, see: M. Ishifune, H. Yamashita, Y. Kera, N. Yamashita, K. Hirata, H. Murase and S. Kashimura, *Electrochim. Acta*, 2003, **48**, 2405–2409.
291 For electrochemical reduction of pyridine derivatives, see: Y. Kita, H. Maekawa, Y. Yamasaki and I. Nishiguchi, *Tetrahedron*, 2001, **57**, 2095–2102.
292 J.-B. Tommasino, A. Brondex, M. Médebielle, M. Thomalla, B. R. Langlois and T. Billard, *Synlett*, 2002, 1697–1699.
293 A. G. O'Brien, A. Maruyama, Y. Inokuma, M. Fujita, P. S. Baran and D. G. Blackmond, *Angew. Chem., Int. Ed.*, 2014, **53**, 11868–11871.
294 T. Fuchigami, M. Shimojo, A. Konno and K. Nakagawa, *J. Org. Chem.*, 1990, **55**, 6074–6075.
295 A. Konno, K. Nakagawa and T. Fuchigami, *J. Chem. Soc., Chem. Commun.*, 1991, 1027–1029.
296 T. Fuchigami, H. Yano and A. Konno, *J. Org. Chem.*, 1991, **56**, 6731–6733.
297 T. Fuchigami, A. Konno, K. Nakagawa and M. Shimojo, *J. Org. Chem.*, 1994, **59**, 5937–5941.
298 A. Konno and T. Fuchigami, *J. Appl. Electrochem.*, 1995, **25**, 173–175.
299 T. Fuchigami, M. Shimojo and A. Konno, *J. Org. Chem.*, 1995, **60**, 3459–3464.
300 A. Konno and T. Fuchigami, *J. Org. Chem.*, 1997, **62**, 8579–8581.
301 S. Higashiya, T. Sato and T. Fuchigami, *J. Fluorine Chem.*, 1998, **87**, 203–208.
302 See also: T. Fuchigami, K. Yamamoto and A. Konno, *Tetrahedron*, 1991, **47**, 625–634.
303 T. Brigaud and E. Laurent, *Tetrahedron Lett.*, 1990, **31**, 2287–2290.
304 T. Brigaud, A. Laurent and E. Laurent, *Phosphorus, Sulfur Silicon Relat. Elem.*, 1991, **59**, 153–156.
305 For discussions on the parameters affecting fluorination, see: (*a*) V. Suryanarayanan and M. Noel, *J. Fluorine Chem.*, 1998, **91**, 153–157; (*b*) K. M. Dawood and T. Fuchigami, *J. Org. Chem.*, 1999, **64**, 138–143; (*c*) K. Suzuki, S. Inagi and T. Fuchigami, *Electrochim. Acta*, 2009, **54**, 961–965.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

306 P. Baroux, R. Tardivel and J. Simonet, *Tetrahedron Lett.*, 1995, **36**, 3851–3852.
307 P. Baroux, R. Tardivel and J. Simonet, *J. Electrochem. Soc.*, 1997, **144**, 841–846.
308 A. Konno, W. Naito and T. Fuchigami, *Tetrahedron Lett.*, 1992, **33**, 7017–7020.
309 Y. Hou, S. Higashiya and T. Fuchigami, *J. Org. Chem.*, 1997, **62**, 9173–9176.
310 S. M. Riyadh, H. Ishii and T. Fuchigami, *Tetrahedron*, 2001, **57**, 8817–8821.
311 S. Kuribayashi, N. Shida, S. Inagi and T. Fuchigami, *Tetrahedron*, 2016, **72**, 5343–5349.
312 K. M. Dawood, H. Ishii and T. Fuchigami, *J. Org. Chem.*, 2001, **66**, 7030–7034.
313 T. Fuchigami, S. Narizuka, A. Konno and K. Momota, *Electrochim. Acta*, 1998, **43**, 1985–1989.
314 M. R. Shaaban, H. Ishii and T. Fuchigami, *J. Org. Chem.*, 2001, **66**, 5633–5636.
315 S. Higashiya, S. Narizuka, A. Konno, T. Maeda, K. Momota and T. Fuchigami, *J. Org. Chem.*, 1999, **64**, 133–137.
316 M. R. Shaaban, H. Ishii and T. Fuchigami, *J. Org. Chem.*, 2000, **65**, 8685–8689.
317 Y. Cao, A. Hidaka, T. Tajima and T. Fuchigami, *J. Org. Chem.*, 2005, **70**, 9614–9617.
318 B. Zagipa, A. Hidaka, Y. Cao and T. Fuchigami, *J. Fluorine Chem.*, 2006, **127**, 552–557.
319 K. M. Dawood, S. Higashiya, Y. Hou and T. Fuchigami, *J. Org. Chem.*, 1999, **64**, 7935–7939.
320 S. Inagi, T. Sawamura and T. Fuchigami, *Electrochem. Commun.*, 2008, **10**, 1158–1160.
321 M. Noel and V. Suryanarayanan, *J. Appl. Electrochem.*, 2004, **34**, 357–369.
322 K. M. Dawood, *Tetrahedron*, 2004, **60**, 1435–1451.
323 T. Fuchigami and T. Tajima, *J. Fluorine Chem.*, 2005, **126**, 181–187.
324 T. Fuchigami, *J. Fluorine Chem.*, 2007, **128**, 311–316.
325 T. Fuchigami and S. Inagi, *Chem. Commun.*, 2011, **47**, 10211–10223.
326 Z. U. H. Khan, D. Kong, Y. Chen, N. Muhammad, A. U. Khan, F. U. Khan, K. Tahir, A. Ahmad, L. Wang and P. Wan, *J. Ind. Eng. Chem.*, 2015, **31**, 26–38.
327 T. Sunaga, M. Atobe, S. Inagi and T. Fuchigami, *Chem. Commun.*, 2009, 956–958.
328 T. Tajima, A. Nakajima, Y. Doi and T. Fuchigami, *Angew. Chem., Int. Ed.*, 2007, **46**, 3550–3552.
329 T. Sawamura, S. Kuribayashi, S. Inagi and T. Fuchigami, *Org. Lett.*, 2010, **12**, 644–646.
330 D. Baba, Y.-J. Yang, B.-J. Uang and T. Fuchigami, *J. Fluorine Chem.*, 2003, **121**, 93–96.
331 K. Suzuki and T. Fuchigami, *J. Org. Chem.*, 2004, **69**, 1276–1282.
332 D. Baba, H. Ishii, S. Higashiya, K. Fujisawa and T. Fuchigami, *J. Org. Chem.*, 2001, **66**, 7020–7024.
333 S. Narizuka, H. Koshiyama, A. Konno and T. Fuchigami, *J. Fluorine Chem.*, 1995, **73**, 121–127.
334 M. Aoyama, T. Fukuhara and S. Hara, *J. Org. Chem.*, 2008, **73**, 4186–4189.
335 M. Monoi and S. Hara, *J. Fluorine Chem.*, 2012, **140**, 28–30.
336 M. Hasegawa, H. Ishii and T. Fuchigami, *Tetrahedron Lett.*, 2002, **43**, 1503–1505.
337 M. Hasegawa, H. Ishii, Y. Cao and T. Fuchigami, *J. Electrochem. Soc.*, 2006, **153**, D162–D166.
338 See also: K. M. Dawood and T. Fuchigami, *J. Org. Chem.*, 2001, **66**, 7691–7695.
339 K. M. Dawood and T. Fuchigami, *Tetrahedron Lett.*, 2001, **42**, 2513–2515.
340 K. M. Dawood and T. Fuchigami, *J. Org. Chem.*, 2005, **70**, 7537–7541.
341 H. Ishii, N. Yamada and T. Fuchigami, *Chem. Commun.*, 2000, 1617–1618.
342 H. Ishii, N. Yamada and T. Fuchigami, *Tetrahedron*, 2001, **57**, 9067–9072.
343 M. Hasegawa, H. Ishii and T. Fuchigami, *Green Chem.*, 2003, **5**, 512–515.
344 M. Hasegawa and T. Fuchigami, *Electrochim. Acta*, 2004, **49**, 3367–3372.
345 T. Sawamura, S. Kuribayashi, S. Inagi and T. Fuchigami, *Adv. Synth. Catal.*, 2010, **352**, 2757–2760.
346 T. Sawamura, K. Takahashi, S. Inagi and T. Fuchigami, *Angew. Chem., Int. Ed.*, 2012, **51**, 4413–4416.
347 B. Yin, S. Inagi and T. Fuchigami, *Beilstein J. Org. Chem.*, 2015, **11**, 85–91.
348 For fluorodesulfurization of polymers, see: (*a*) S. Inagi, S. Hayashi and T. Fuchigami, *Chem. Commun.*, 2009, 1718–1720; (*b*) S. Inagi and T. Fuchigami, *Macromol. Rapid Commun.*, 2014, **35**, 854–867.
349 For the use of boron derivatives as electroauxiliaries, see: (*a*) M. Tanigawa, Y. Kuriyama, S. Inagi and T. Fuchigami, *Electrochim. Acta*, 2016, **199**, 314–318; (*b*) J. Suzuki, N. Shida, S. Inagi and T. Fuchigami, *Electroanalysis*, 2016, **28**, 2797–2801.
350 Y. Hou, S. Higashiya and T. Fuchigami, *J. Org. Chem.*, 1999, **64**, 3346–3349.
351 V. Dinoiu, K. Kanno, T. Fukuhara and N. Yoneda, *J. Fluorine Chem.*, 2005, **126**, 753–758.
352 H. Nagura, S. Inagi and T. Fuchigami, *Tetrahedron*, 2009, **65**, 1559–1566.
353 H. Nagura, S. Kuribayashi, Y. Ishiguro, S. Inagi and T. Fuchigami, *Tetrahedron*, 2010, **66**, 183–186.
354 S. Fujie, K. Matsumoto, S. Suga and J.-i. Yoshida, *Chem. Lett.*, 2009, **38**, 1186–1187.
355 V. Dinoiu, T. Fukuhara, S. Hara and N. Yoneda, *J. Fluorine Chem.*, 2000, **103**, 75–80.
356 For an electrochemical approach to hydrodefluorination of fluoroaromatic compounds, see: W.-B. Wu, M.-L. Li and J.-M. Huang, *Tetrahedron Lett.*, 2015, **56**, 1520–1523.
357 G. Faita, M. Fleischmann and D. Pletcher, *J. Electroanal. Chem.*, 1970, **25**, 455–459.
358 L. Appelbaum, D. Danovich, G. Lazanes, M. Michman and M. Oron, *J. Electroanal. Chem.*, 2001, **499**, 39–47.
359 For anodic chlorination of aniline, see: Y. Matsuda and T. Nishiki, *Electrochim. Acta*, 1984, **29**, 35–39.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

360 G. Casalbore, M. Mastragostino and S. Valcher, *J. Electroanal. Chem.*, 1978, **87**, 411–418.
361 I. Taniguchi, M. Yano, H. Yamaguchi and K. Yasukouchi, *J. Electroanal. Chem.*, 1982, **132**, 233–245.
362 M. Mastragostino, S. Valcher and M. Biserni, *J. Electroanal. Chem.*, 1983, **158**, 369–373.
363 C. Visy and M. Novák, *Electrochim. Acta*, 1987, **32**, 1757–1759.
364 G. D. Allen, M. C. Buzzeo, I. G. Davies, C. Villagrán, C. Hardacre and R. G. Compton, *J. Phys. Chem. B*, 2004, **108**, 16322–16327.
365 E. I. Rogers, D. S. Silvester, L. Aldous, C. Hardacre and R. G. Compton, *J. Phys. Chem. C*, 2008, **112**, 6551–6557.
366 For a review on electrochemical halogenation of organic compounds, see: B. V. Lyalin and V. A. Petrosyan, *Russ. J. Electrochem.*, 2013, **49**, 497–529.
367 For a review on halogenation of organic compounds using continuous flow and microreactor technology, see: D. Cantillo and C. O. Kappe, *React. Chem. Eng.*, 2017, **2**, 7–19.
368 S. R. Ellis, D. Pletcher, W. N. Brooks and K. P. Healy, *J. Appl. Electrochem.*, 1983, **13**, 735–741.
369 S. R. Forsyth, D. Pletcher and K. P. Healy, *J. Appl. Electrochem.*, 1987, **17**, 905–913.
370 B. V. Lyalin, V. A. Petrosyan and B. I. Ugrak, *Russ. J. Electrochem.*, 2008, **44**, 1320–1326.
371 T. Raju, G. Kalpana Devi and K. Kulangiappar, *Electrochim. Acta*, 2006, **51**, 4596–4600.
372 Takahiro Sawamura, Kohta Takahashi, Shinsuke Inagi and Toshio Fuchigami, *Electrochemistry*, 2013, **81**, 365–367.
373 T. Raju, K. Kulangiappar, M. Anbu Kulandainathan, U. Uma, R. Malini and A. Muthukumaran, *Tetrahedron Lett.*, 2006, **47**, 4581–4584.
374 T. Raju, K. Kulangiappar, M. Anbu Kulandainathan and A. Muthukumaran, *Tetrahedron Lett.*, 2005, **46**, 7047–7050.
375 K. Kulangiappar, G. Karthik and M. Anbu Kulandainathan, *Synth. Commun.*, 2009, **39**, 2304–2309.
376 K. Tsuchida, T. Kochi and F. Kakiuchi, *Asian J. Org. Chem.*, 2013, **2**, 935–937.
377 E. M. Ungureanu, A. C. Razus, L. Birzan, G. Buica, M. Cretu and E. Cristian, *Electrochim. Acta*, 2006, **52**, 794–803.
378 For an electrochemical study of azulene derivatives, see: E.-M. Ungureanu, A. C. Razus, L. Birzan, M.-S. Cretu and G.-O. Buica, *Electrochim. Acta*, 2008, **53**, 7089–7099.
379 M. Verniette, C. Daremon and J. Simonet, *Electrochim. Acta*, 1978, **23**, 929–934.
380 M. Novák, C. Visy and K. Bodor, *Electrochim. Acta*, 1982, **27**, 1293–1295.
381 M. Novák and C. Visy, *Electrochim. Acta*, 1983, **28**, 507–510.
382 M. Novák and C. Visy, *Electrochim. Acta*, 1983, **28**, 511–513.
383 C. Visy and M. Novák, *J. Electroanal. Chem.*, 1988, **252**, 91–97.
384 C. Visy and M. Novák, *J. Electroanal. Chem.*, 1990, **296**, 571–581.
385 D. Stevanović, I. Damljanović, M. Vukićević, N. Manojlović, N. S. Radulović and R. D. Vukićević, *Helv. Chim. Acta*, 2011, **94**, 1406–1415.
386 B. V. Lyalin, V. A. Petrosyan and B. I. Ugrak, *Russ. Chem. Bull.*, 2009, **58**, 291–296.
387 Z. Tan, Y. Liu, R. Helmy, N. R. Rivera, D. Hesk, S. Tyagarajan, L. Yang and J. Su, *Tetrahedron Lett.*, 2017, **58**, 3014–3018.
388 P. Pouillen, R. Minko, M. Verniette, P. Martinet and A. M. Martre, *Electrochim. Acta*, 1979, **24**, 1189–1194.
389 Y. Ashikari, A. Shimizu, T. Nokami and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2013, **135**, 16070–16073.
390 A. Shimizu, R. Hayashi, Y. Ashikari, T. Nokami and J.-i. Yoshida, *Beilstein J. Org. Chem.*, 2015, **11**, 242–248.
391 K. Kulangiappar, M. Ramaprakash, D. Vasudevan and T. Raju, *Synth. Commun.*, 2016, **46**, 145–153.
392 B. V. Lyalin, V. A. Petrosyan and B. I. Ugrak, *Russ. Chem. Bull.*, 2010, **59**, 1549–1555.
393 For electrochemical synthesis of pyrazoles, see: X. Wei and B. Speiser, *Electrochim. Acta*, 1997, **42**, 73–79.
394 K. Kataoka, Y. Hagiwara, K. Midorikawa, S. Suga and J.-i. Yoshida, *Org. Process Res. Dev.*, 2008, **12**, 1130–1136.
395 K. Midorikawa, S. Suga and J.-i. Yoshida, *Chem. Commun.*, 2006, 3794–3796.
396 R. Möckel, J. Hille, E. Winterling, S. Weidemüller, T. M. Faber and G. Hilt, *Angew. Chem., Int. Ed.*, 2018, **57**, 442–445.
397 Y. Zhu, Y. Zhu, H. Zeng, Z. Chen, R. D. Little and C. Ma, *J. Electroanal. Chem.*, 2015, **751**, 105–110.
398 M. Rafiee, F. Wang, D. P. Hruszkewycz and S. S. Stahl, *J. Am. Chem. Soc.*, 2018, **140**, 22–25.
399 For selective electrochemical generation of benzylic radicals using ferrocene-based mediators, see: A. J. J. Lennox, J. E. Nutting and S. S. Stahl, *Chem. Sci.*, 2018, **9**, 356–361.
400 M. Masui, S. Hara, T. Ueshima, T. Kawaguchi and S. Ozaki, *Chem. Pharm. Bull.*, 1983, **31**, 4209–4211.
401 M. Masui, T. Kawaguchi and S. Ozaki, *J. Chem. Soc., Chem. Commun.*, 1985, 1484–1485.
402 For reviews on NHPI-mediated reactions, see: (*a*) Y. Ishii, S. Sakaguchi and T. Iwahama, *Adv. Synth. Catal.*, 2001, **343**, 393–427; (*b*) R. A. Sheldon and I. W. C. E. Arends, *Adv. Synth. Catal.*, 2004, **346**, 1051–1071; (*c*) F. Recupero and C. Punta, *Chem. Rev.*, 2007, **107**, 3800–3842; (*d*) S. Coseri, *Catal. Rev.*, 2009, **51**, 218–292; (*e*) S. Wertz and A. Studer, *Green Chem.*, 2013, **15**, 3116–3134; (*f*) L. Melone and C. Punta, *Beilstein J. Org. Chem.*, 2013, **9**, 1296–1310.
403 The bond dissociation energy (BDE) of NHPI has been determined to 88 kcal mol^{-1} and the BDE of toluene derivatives ranges from 88–90 kcal mol^{-1}. See: (*a*) T. Fox and P. A. Kollman, *J. Phys. Chem.*, 1996, **100**, 2950–2956; (*b*) G. P. Bean, *Tetrahedron*, 2002, **58**, 9941–9948; (*c*) R. Amorati, M. Lucarini, V. Mugnaini, G. F. Pedulli, F. Minisci, F. Recupero, F. Fontana, P. Astolfi and L. Greci, *J. Org. Chem.*, 2003, **68**, 1747–1754; (*d*) N. Hirai, N. Sawatari, N. Nakamura, S. Sakaguchi and Y. Ishii, *J. Org. Chem.*, 2003, **68**, 6587–6590; (*e*) D. A. Pratt, G. A. DiLabio, P. Mulder and K. U. Ingold, *Acc. Chem. Res.*, 2004, **37**, 334–340.
404 I. Nishiguchi, O. Kanbe, K. Itoh and H. Maekawa, *Synlett*, 2000, 89–91.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

405 T. Shono, Y. Matsumura, K. Inoue and F. Lwasaki, *J. Chem. Soc., Perkin Trans. 1*, 1986, 73–77.
406 F. Barba, M. N. Elinson, J. Escudero and S. K. Feducovich, *Tetrahedron*, 1997, **53**, 4427–4436.
407 F. Barba, M. N. Elinson, J. Escudero, M. Guirado and S. K. Feducovich, *Electrochim. Acta*, 1998, **43**, 973–976.
408 M. N. Elinson, S. K. Feducovich, A. S. Dorofeev, A. N. Vereshchagin and G. I. Nikishin, *Tetrahedron*, 2000, **56**, 9999–10003.
409 M. N. Elinson, S. K. Feducovich, D. E. Dmitriev, A. S. Dorofeev, A. N. Vereshchagin and G. I. Nikishin, *Tetrahedron Lett.*, 2001, **42**, 5557–5559.
410 M. N. Elinson, A. S. Dorofeev, S. K. Feducovich, R. F. Nasybullin, E. F. Litvin, M. V. Kopyshev and G. I. Nikishin, *Tetrahedron*, 2006, **62**, 8021–8028.
411 C. Yao, Y. Wang, T. Li, C. Yu, L. Li and C. Wang, *Tetrahedron*, 2013, **69**, 10593–10597.
412 H. Gao, Z. Zha, Z. Zhang, H. Ma and Z. Wang, *Chem. Commun.*, 2014, **50**, 5034–5036.
413 M. N. Elinson, S. K. Feducovich, S. G. Bushuev, A. A. Zakharenkov, D. V. Pashchenko and G. I. Nikishin, *Mendeleev Commun.*, 1998, **8**, 15–16.
414 M. N. Elinson, S. K. Feducovich, T. L. Lizunova and G. I. Nikishin, *Tetrahedron*, 2000, **56**, 3063–3069.
415 M. N. Elinson, S. K. Feducovich, Z. A. Starikova, O. S. Olessova, A. N. Vereshchagin and G. I. Nikishin, *Tetrahedron Lett.*, 2000, **41**, 4937–4941.
416 M. N. Elinson, S. K. Feducovich and G. I. Nikishin, *ARKIVOC*, 2001, 31–36.
417 O. Mitsuhiro, T. Yukio and K. Toyoji, *Bull. Chem. Soc. Jpn.*, 2003, **76**, 207–208.
418 M. N. Elinson, S. K. Fedukovich, T. A. Zaimovskaya, A. N. Vereshchagin and G. I. Nikishin, *Russ. Chem. Bull.*, 2003, **52**, 2241–2246.
419 M. N. Elinson, S. K. Feducovich, Z. A. Starikova, A. N. Vereshchagin, S. V. Gorbunov and G. I. Nikishin, *Tetrahedron Lett.*, 2005, **46**, 6389–6393.
420 M. N. Elinson, S. K. Feducovich, Z. A. Starikova, A. N. Vereshchagin, P. A. Belyakov and G. I. Nikishin, *Tetrahedron*, 2006, **62**, 3989–3996.
421 A. N. Vereshchagin, M. N. Elinson, T. A. Zaimovskaya and G. I. Nikishin, *Tetrahedron*, 2008, **64**, 9766–9770.
422 D. Minato, S. Mizuta, M. Kuriyama, Y. Matsumura and O. Onomura, *Tetrahedron*, 2009, **65**, 9742–9748.
423 M. Okimoto, H. Yamamori, K. Ohashi, S. Nishikawa, M. Hoshi and T. Yoshida, *Synlett*, 2012, 2544–2548.
424 M. Okimoto, H. Yamamori, K. Ohashi, M. Hoshi and T. Yoshida, *Synlett*, 2013, 1568–1572.
425 A. N. Vereshchagin, M. N. Elinson, E. O. Dorofeeva, D. V. Demchuk, I. S. Bushmarinov, A. S. Goloveshkin and G. I. Nikishin, *Tetrahedron*, 2013, **69**, 5234–5241.
426 M. Okimoto, H. Yamamori, M. Hoshi and T. Yoshida, *Tetrahedron Lett.*, 2014, **55**, 1299–1302.
427 A. N. Vereshchagin, M. N. Elinson, E. O. Dorofeeva, R. F. Nasybullin, I. S. Bushmarinov, A. S. Goloveshkin and M. P. Egorov, *Electrochim. Acta*, 2015, **165**, 116–121.
428 M. N. Elinson, E. O. Dorofeeva, A. N. Vereshchagin, R. F. Nasybullin and M. P. Egorov, *Catal. Sci. Technol.*, 2015, **5**, 2384–2387.
429 Z. Zhang, J. Su, Z. Zha and Z. Wang, *Chem. – Eur. J.*, 2013, **19**, 17711–17714.
430 For electrochemical C—H acylation of electron-deficient N-heteroarenes with α-keto acids, see: Q.-Q. Wang, K. Xu, Y.-Y. Jiang, Y.-G. Liu, B.-G. Sun and C.-C. Zeng, *Org. Lett.*, 2017, **19**, 5517–5520.
431 For preparation of benzimidazoles through decarboxylation of α-keto acids, see: H.-B. Wang and J.-M. Huang, *Adv. Synth. Catal.*, 2016, **358**, 1975–1981.
432 Z. Zhang, J. Su, Z. Zha and Z. Wang, *Chem. Commun.*, 2013, **49**, 8982–8984.
433 P. Qian, J.-H. Su, Y. Wang, M. Bi, Z. Zha and Z. Wang, *J. Org. Chem.*, 2017, **82**, 6434–6440.
434 N.-N. Bui, X.-H. Ho, S.-i. Mho and H.-Y. Jang, *Eur. J. Org. Chem.*, 2009, 5309–5312.
435 (*a*) M. Marigo, T. C. Wabnitz, D. Fielenbach and K. A. Jørgensen, *Angew. Chem., Int. Ed.*, 2005, **44**, 794–797; (*b*) Y. Hayashi, H. Gotoh, T. Hayashi and M. Shoji, *Angew. Chem., Int. Ed.*, 2005, **44**, 4212–4215.
436 For reviews on diarylprolinol silyl ethers as organocatalysts, see: (*a*) K. L. Jensen, G. Dickmeiss, H. Jiang, Ł. Albrecht and K. A. Jørgensen, *Acc. Chem. Res.*, 2012, **45**, 248–264; (*b*) L. Klier, F. Tur, P. H. Poulsen and K. A. Jørgensen, *Chem. Soc. Rev.*, 2017, **46**, 1080–1102.
437 E. E. Finney, K. A. Ogawa and A. J. Boydston, *J. Am. Chem. Soc.*, 2012, **134**, 12374–12377.
438 For electrochemical characterization of azolium salts, see: K. A. Ogawa and A. J. Boydston, *Chem. Lett.*, 2014, **43**, 907–909.
439 For recent reviews on the oxidative esterification and amidation of aldehydes, see: (*a*) K. Ekoue-Kovi and C. Wolf, *Chem. – Eur. J.*, 2008, **14**, 6302–6315; (*b*) S. Gaspa, A. Porcheddu and L. De Luca, *Tetrahedron Lett.*, 2016, **57**, 3433–3440.
440 K. A. Ogawa and A. J. Boydston, *Org. Lett.*, 2014, **16**, 1928–1931.
441 R. A. Green, D. Pletcher, S. G. Leach and R. C. D. Brown, *Org. Lett.*, 2015, **17**, 3290–3293.
442 R. A. Green, D. Pletcher, S. G. Leach and R. C. D. Brown, *Org. Lett.*, 2016, **18**, 1198–1201.
443 N. Sauermann, T. H. Meyer, C. Tian and L. Ackermann, *J. Am. Chem. Soc.*, 2017, **139**, 18452–18455.
444 Y. Qiu, W.-J. Kong, J. Struwe, N. Sauermann, T. Rogge, A. Scheremetjew and L. Ackermann, *Angew. Chem., Int. Ed.*, 2018, **57**, 5828–5832.
445 Y. Qiu, C. Tian, L. Massignan, T. Rogge and L. Ackermann, *Angew. Chem., Int. Ed.*, 2018, **57**, 5818–5822.
446 F. Xu, X.-Y. Qian, Y.-J. Li and H.-C. Xu, *Org. Lett.*, 2017, **19**, 6332–6335.
447 S. Zhang, L. Li, H. Wang, Q. Li, W. Liu, K. Xu and C. Zeng, *Org. Lett.*, 2018, **20**, 252–255.
448 For a related approach, see: L. Zhang, Z. Zhang, J. Hong, J. Yu, J. Zhang and F. Mo, *J. Org. Chem.*, 2018, **83**, 3200–3207.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

449 For a related electrocatalytic dehydrogenative esterification strategy, see: S. Zhang, F. Lian, M. Xue, T. Qin, L. Li, X. Zhang and K. Xu, *Org. Lett.*, 2017, **19**, 6622–6625.
450 For electrochemical synthesis of phthalides, see: D. Hayrapetyan, V. Shkepu, O. T. Seilkhanov, Z. Zhanabil and K. Lam, *Chem. Commun.*, 2017, **53**, 8451–8454.
451 S. Nad and R. Breinbauer, *Angew. Chem., Int. Ed.*, 2004, **43**, 2297–2299.
452 S. Nad, S. Roller, R. Haag and R. Breinbauer, *Org. Lett.*, 2006, **8**, 403–406.
453 C. Laurencé, M. Rivard, I. Lachaise, J. Bensemhoun and T. Martens, *Tetrahedron*, 2011, **67**, 9518–9521.
454 E. Nieto-Mendoza, J. A. Guevara-Salazar, M. T. Ramírez-Apan, B. A. Frontana-Uribe, J. A. Cogordan and J. Cárdenas, *J. Org. Chem.*, 2005, **70**, 4538–4541.
455 M. Albert, D. De Souza, P. Feiertag and H. Hönig, *Org. Lett.*, 2002, **4**, 3251–3254.
456 For issues associated with the use of oxidants in the pharmaceutical industry, see: S. Caron, R. W. Dugger, S. G. Ruggeri, J. A. Ragan and D. H. B. Ripin, *Chem. Rev.*, 2006, **106**, 2943–2989.
457 L. Meng, J. Su, Z. Zha, L. Zhang, Z. Zhang and Z. Wang, *Chem. – Eur. J.*, 2013, **19**, 5542–5545.
458 M. Okajima, K. Soga, T. Nokami, S. Suga and J.-i. Yoshida, *Org. Lett.*, 2006, **8**, 5005–5007.
459 T. Nokami, K. Ohata, M. Inoue, H. Tsuyama, A. Shibuya, K. Soga, M. Okajima, S. Suga and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2008, **130**, 10864–10865.
460 M. Okajima, K. Soga, T. Watanabe, K. Terao, T. Nokami, S. Suga and J.-i. Yoshida, *Bull. Chem. Soc. Jpn.*, 2009, **82**, 594–599.
461 K. Terao, T. Watanabe, T. Suehiro, T. Nokami and J.-i. Yoshida, *Tetrahedron Lett.*, 2010, **51**, 4107–4109.
462 T. Nokami, T. Watanabe, N. Musya, T. Morofuji, K. Tahara, Y. Tobe and J.-i. Yoshida, *Chem. Commun.*, 2011, **47**, 5575–5577.
463 T. Nokami, T. Watanabe, N. Musya, T. Suehiro, T. Morofuji and J.-i. Yoshida, *Tetrahedron*, 2011, **67**, 4664–4671.
464 T. Nokami, T. Watanabe, K. Terao, K. Soga, K. Ohata and J.-i. Yoshida, *Electrochemistry*, 2013, **81**, 399–401.
465 T. Nokami, N. Musya, T. Morofuji, K. Takeda, M. Takumi, A. Shimizu and J.-i. Yoshida, *Beilstein J. Org. Chem.*, 2014, **10**, 3097–3103.
466 For generation of benzylic cations through carbon–carbon bond cleavage, see: M. Okajima, S. Suga, K. Itami and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2005, **127**, 6930–6931.
467 Y. Ashikari, T. Nokami and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2011, **133**, 11840–11843.
468 Y. Ashikari, T. Nokami and J.-i. Yoshida, *Org. Lett.*, 2012, **14**, 938–941.
469 T. Tajima, Y. Kishi and A. Nakajima, *Electrochim. Acta*, 2009, **54**, 5959–5963.
470 For early work on anodic acetoxylation, see: (*a*) L. Eberson and K. Nyberg, *J. Am. Chem. Soc.*, 1966, **88**, 1686–1691; (*b*) L. Eberson, *J. Am. Chem. Soc.*, 1967, **89**, 4669–4677.
471 J. H. P. Utley and G. G. Rozenberg, *J. Appl. Electrochem.*, 2003, **33**, 525–532.
472 For electrochemical synthesis of benzoxazoles using DDQ as a redox catalyst, see: L.-S. Kang, H.-l. Xiao, C.-C. Zeng, L.-M. Hu and R. D. Little, *J. Electroanal. Chem.*, 2016, **767**, 13–17.
473 D. P. Hruszkewycz, K. C. Miles, O. R. Thiel and S. S. Stahl, *Chem. Sci.*, 2017, **8**, 1282–1287.
474 N.-t. Zhang, C.-c. Zeng, C. M. Lam, R. K. Gbur and R. D. Little, *J. Org. Chem.*, 2013, **78**, 2104–2110.
475 R. Francke and R. D. Little, *J. Am. Chem. Soc.*, 2014, **136**, 427–435.
476 N.-n. Lu, S. J. Yoo, L.-J. Li, C.-C. Zeng and R. D. Little, *Electrochim. Acta*, 2014, **142**, 254–260.
477 K.-Y. Zhang, N.-n. Lu, S. J. Yoo, L.-M. Hu, R. D. Little and C.-C. Zeng, *Electrochim. Acta*, 2016, **199**, 357–365.
478 C.-c. Zeng, N.-t. Zhang, C. M. Lam and R. D. Little, *Org. Lett.*, 2012, **14**, 1314–1317.
479 See also: S. J. Yoo, L.-J. Li, C.-C. Zeng and R. D. Little, *Angew. Chem., Int. Ed.*, 2015, **54**, 3744–3747.
480 For arylative ring-opening of epoxides using triarylimidazole redox mediators, see: N.-n. Lu, N.-t. Zhang, C.-C. Zeng, L.-M. Hu, S. J. Yoo and R. D. Little, *J. Org. Chem.*, 2015, **80**, 781–789.
481 E. J. Horn, B. R. Rosen, Y. Chen, J. Tang, K. Chen, M. D. Eastgate and P. S. Baran, *Nature*, 2016, **533**, 77–81.
482 S. R. Waldvogel and M. Selt, *Angew. Chem., Int. Ed.*, 2016, **55**, 12578–12580.
483 For seminal work on electrochemical allylic C–H oxidation using NHPI as mediator, see: M. Masui, K. Hosomi, K. Tsuchida and S. Ozaki, *Chem. Pharm. Bull.*, 1985, **33**, 4798–4802.
484 J. Kowalski, J. Płoszyńska, A. Sobkowiak, J. W. Morzycki and A. Z. Wilczewska, *J. Electroanal. Chem.*, 2005, **585**, 275–280.
485 J. Kowalski, Z. Łotowski, J. W. Morzycki, J. Płoszyńska, A. Sobkowiak and A. Z. Wilczewska, *Steroids*, 2008, **73**, 543–548.
486 Y.-Y. Hosokawa, H. Hakamata, T. Murakami, S. Aoyagi, M. Kuroda, Y. Mimaki, A. Ito, S. Morosawa and F. Kusu, *Electrochim. Acta*, 2009, **54**, 6412–6416.
487 Y.-Y. Hosokawa, H. Hakamata, T. Murakami and F. Kusu, *Tetrahedron Lett.*, 2010, **51**, 129–132.
488 For electrochemical synthesis of glycoconjugates, see: A. M. Tomkiel, J. Kowalski, J. Płoszyńska, L. Siergiejczyk, Z. Łotowski, A. Sobkowiak and J. W. Morzycki, *Steroids*, 2014, **82**, 60–67.
489 For electrochemical synthesis of glycoconjugates using nonactivated sugars, see: A. M. Tomkiel, K. Brzezinski, Z. Łotowski, L. Siergiejczyk, P. Wałejko, S. Witkowski, J. Kowalski, J. Płoszyńska, A. Sobkowiak and J. W. Morzycki, *Tetrahedron*, 2013, **69**, 8904–8913.
490 Y. Kawamata, M. Yan, Z. Liu, D.-H. Bao, J. Chen, J. T. Starr and P. S. Baran, *J. Am. Chem. Soc.*, 2017, **139**, 7448–7451.
491 C. Sambiagio, H. Sterckx and B. U. W. Maes, *ACS Cent. Sci.*, 2017, **3**, 686–688.
492 For general reviews on palladium-catalyzed reactions, see: (*a*) K. C. Nicolaou, P. G. Bulger and D. Sarlah, *Angew. Chem., Int. Ed.*, 2005, **44**, 4442–4489; (*b*) R. Martin and

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

S. L. Buchwald, *Acc. Chem. Res.*, 2008, **41**, 1461–1473; (*c*) G. C. Fu, *Acc. Chem. Res.*, 2008, **41**, 1555–1564; (*d*) X. Chen, K. M. Engle, D.-H. Wang and J.-Q. Yu, *Angew. Chem., Int. Ed.*, 2009, **48**, 5094–5115; (*e*) X.-F. Wu, H. Neumann and M. Beller, *Chem. Soc. Rev.*, 2011, **40**, 4986–5009; (*f*) C. C. C. Johansson Seechurn, M. O. Kitching, T. J. Colacot and V. Snieckus, *Angew. Chem., Int. Ed.*, 2012, **51**, 5062–5085; (*g*) H. Li, C. C. C. Johansson Seechurn and T. J. Colacot, *ACS Catal.*, 2012, **2**, 1147–1164; (*h*) P. G. Gildner and T. J. Colacot, *Organometallics*, 2015, **34**, 5497–5508; (*i*) R. Takise, K. Muto and J. Yamaguchi, *Chem. Soc. Rev.*, 2017, **46**, 5864–5888; (*j*) M. M. Lorion, K. Maindan, A. R. Kapdi and L. Ackermann, *Chem. Soc. Rev.*, 2017, **46**, 7399–7420; (*k*) N. Hazari, P. R. Melvin and M. M. Beromi, *Nat. Rev. Chem.*, 2017, **1**, 0025.

493 K. D. Moeller, *Organometallics*, 2014, **33**, 4607–4616.

494 Y. B. Dudkina, T. V. Gryaznova, O. G. Sinyashin and Y. H. Budnikova, *Russ. Chem. Bull.*, 2015, **64**, 1713–1725.

495 K.-J. Jiao, C.-Q. Zhao, P. Fang and T.-S. Mei, *Tetrahedron Lett.*, 2017, **58**, 797–802.

496 F. Kakiuchi and T. Kochi, *Isr. J. Chem.*, 2017, **57**, 953–963.

497 For an example on the use of electrosynthesis for generation of catalytic Pd intermediates, see: V. A. Zinovyeva, C. Luo, S. Fournier, C. H. Devillers, H. Cattey, H. Doucet, J.-C. Hierso and D. Lucas, *Chem. – Eur. J.*, 2011, **17**, 9901–9906.

498 J. Tsuji and M. Minato, *Tetrahedron Lett.*, 1987, **28**, 3683–3686.

499 See also: K. Mitsudo, S. Fukunaga, T. Fujita, H. Mandai, S. Suga and H. Tanaka, *Electrochemistry*, 2013, **81**, 347–349.

500 C. Amatore, C. Cammoun and A. Jutand, *Adv. Synth. Catal.*, 2007, **349**, 292–296.

501 For electrochemically-mediated Heck reactions that rely on cathodic reduction of PdI^{I} to Pd0, see: (*a*) J. Tian, K. Maurer, E. Tesfu and K. D. Moeller, *J. Am. Chem. Soc.*, 2005, **127**, 1392–1393; (*b*) J. Tian and K. D. Moeller, *Org. Lett.*, 2005, **7**, 5381–5383; (*c*) L. Hu, M. Stuart, J. Tian, K. Maurer and K. D. Moeller, *J. Am. Chem. Soc.*, 2010, **132**, 16610–16616.

502 K. Mitsudo, N. Kamimoto, H. Murakami, H. Mandai, A. Wakamiya, Y. Murata and S. Suga, *Org. Biomol. Chem.*, 2012, **10**, 9562–9569.

503 C. Amatore, C. Cammoun and A. Jutand, *Eur. J. Org. Chem.*, 2008, 4567–4570.

504 C. Amatore, C. Cammoun and A. Jutand, *Synlett*, 2007, 2173–2178.

505 K. Mitsudo, T. Shiraga and H. Tanaka, *Tetrahedron Lett.*, 2008, **49**, 6593–6595.

506 K. Mitsudo, T. Shiraga, D. Kagen, D. Shi, J. Y. Becker and H. Tanaka, *Tetrahedron*, 2009, **65**, 8384–8388.

507 or electroreductive coupling of aryl bromides using a Pd/viologen double mediatory system, see: M. Kuroboshi, T. Shiba and H. Tanaka, *Tetrahedron Lett.*, 2013, **54**, 3666–3668.

508 K. Mitsudo, T. Shiraga, J.-i. Mizukawa, S. Suga and H. Tanaka, *Chem. Commun.*, 2010, **46**, 9256–9258.

509 K. Mitsudo, T. Kaide, E. Nakamoto, K. Yoshida and H. Tanaka, *J. Am. Chem. Soc.*, 2007, **129**, 2246–2247.

510 K. Mitsudo, T. Ishii and H. Tanaka, *Electrochemistry*, 2008, **76**, 859–861.

511 E. Tesfu, K. Maurer, S. R. Ragsdale and K. D. Moeller, *J. Am. Chem. Soc.*, 2004, **126**, 6212–6213.

512 E. Tesfu, K. Maurer and K. D. Moeller, *J. Am. Chem. Soc.*, 2006, **128**, 70–71.

513 E. Tesfu, K. Roth, K. Maurer and K. D. Moeller, *Org. Lett.*, 2006, **8**, 709–712.

514 B. H. Nguyen, D. Kesselring, E. Tesfu and K. D. Moeller, *Langmuir*, 2014, **30**, 2280–2286.

515 For a review on he site-selective functionalization of electrode surfaces, see: M. D. Graaf and K. D. Moeller, *Langmuir*, 2015, **31**, 7697–7706.

516 For seminal work on electrochemical regeneration of PdI^{I} using triarylamines as recyclable mediators, see: T. Inokuchi, L. Ping, F. Hamaue, M. Izawa and S. Torii, *Chem. Lett.*, 1994, 121–124.

517 F. Kakiuchi, T. Kochi, H. Mutsutani, N. Kobayashi, S. Urano, M. Sato, S. Nishiyama and T. Tanabe, *J. Am. Chem. Soc.*, 2009, **131**, 11310–11311.

518 M. Konishi, K. Tsuchida, K. Sano, T. Kochi and F. Kakiuchi, *J. Org. Chem.*, 2017, **82**, 8716–8724.

519 K. Sano, N. Kimura, T. Kochi and F. Kakiuchi, *Asian J. Org. Chem.*, DOI: 10.1002/ajoc.201800202.

520 For Pd-catalyzed electroreductive cross-coupling of alkyl and allylic halides, see: Y.-L. Lai and J.-M. Huang, *Org. Lett.*, 2017, **19**, 2022–2025.

521 F. Saito, H. Aiso, T. Kochi and F. Kakiuchi, *Organometallics*, 2014, **33**, 6704–6707.

522 Y. B. Dudkina, D. Y. Mikhaylov, T. V. Gryaznova, A. I. Tufatullin, O. N. Kataeva, D. A. Vicic and Y. H. Budnikova, *Organometallics*, 2013, **32**, 4785–4792.

523 Y. B. Dudkina, D. Y. Mikhaylov, T. V. Gryaznova, O. G. Sinyashin, D. A. Vicic and Y. H. Budnikova, *Eur. J. Org. Chem.*, 2012, 2114–2117.

524 See also: (*a*) D. Y. Mikhaylov, Y. H. Budnikova, T. V. Gryaznova, D. V. Krivolapov, I. A. Litvinov, D. A. Vicic and O. G. Sinyashin, *J. Organomet. Chem.*, 2009, **694**, 3840–3843; (*b*) D. Mikhaylov, T. Gryaznova, Y. Dudkina, M. Khrizanphorov, S. Latypov, O. Kataeva, D. A. Vicic, O. G. Sinyashin and Y. Budnikova, *Dalton Trans.*, 2012, **41**, 165–172.

525 T. V. Grayaznova, Y. B. Dudkina, D. R. Islamov, O. N. Kataeva, O. G. Sinyashin, D. A. Vicic and Y. H. Budnikova, *J. Organomet. Chem.*, 2015, **785**, 68–71.

526 See also: S. O. Strekalova, M. N. Khrizanforov, T. V. Gryaznova, V. V. Khrizanforova and Y. H. Budnikova, *Russ. Chem. Bull.*, 2016, **65**, 1295–1298.

527 H. Aiso, T. Kochi, H. Mutsutani, T. Tanabe, S. Nishiyama and F. Kakiuchi, *J. Org. Chem.*, 2012, **77**, 7718–7724.

528 N. Kamimoto, N. Nakamura, A. Tsutsumi, H. Mandai, K. Mitsudo, A. Wakamiya, Y. Murata, J.-y. Hasegawa and S. Suga, *Asian J. Org. Chem.*, 2016, **5**, 373–379.

529 A. Shrestha, M. Lee, A. L. Dunn and M. S. Sanford, *Org. Lett.*, 2018, **20**, 204–207.

530 Y.-Q. Li, Q.-L. Yang, P. Fang, T.-S. Mei and D. Zhang, *Org. Lett.*, 2017, **19**, 2905–2908.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

531 C. Ma, C.-Q. Zhao, Y.-Q. Li, L.-P. Zhang, X.-T. Xu, K. Zhang and T.-S. Mei, *Chem. Commun.*, 2017, **53**, 12189–12192.
532 Q.-L. Yang, Y.-Q. Li, C. Ma, P. Fang, X.-J. Zhang and T.-S. Mei, *J. Am. Chem. Soc.*, 2017, **139**, 3293–3298.
533 For reviews on metal-catalyzed oxidative cross-couplings, see: (*a*) C.-J. Li, *Acc. Chem. Res.*, 2009, **42**, 335–344; (*b*) C. J. Scheuermann, *Chem. – Asian J.*, 2010, **5**, 436–451; (*c*) J. A. Ashenhurst, *Chem. Soc. Rev.*, 2010, **39**, 540–548; (*d*) C. S. Yeung and V. M. Dong, *Chem. Rev.*, 2011, **111**, 1215–1292; (*e*) S. H. Cho, J. Y. Kim, J. Kwak and S. Chang, *Chem. Soc. Rev.*, 2011, **40**, 5068–5083; (*f*) K. Hirano and M. Miura, *Chem. Commun.*, 2012, **48**, 10704–10714; (*g*) G. Song, F. Wang and X. Li, *Chem. Soc. Rev.*, 2012, **41**, 3651–3678; (*h*) X. Shang and Z.-Q. Liu, *Chem. Soc. Rev.*, 2013, **42**, 3253–3260; (*i*) S. I. Kozhushkov and L. Ackermann, *Chem. Sci.*, 2013, **4**, 886–896; (*j*) S. A. Girard, T. Knauber and C.-J. Li, *Angew. Chem., Int. Ed.*, 2014, **53**, 74–100; (*k*) Y. Wu, J. Wang, F. Mao and F. Y. Kwong, *Chem. – Asian J.*, 2014, **9**, 26–47; (*l*) C. Liu, J. Yuan, M. Gao, S. Tang, W. Li, R. Shi and A. Lei, *Chem. Rev.*, 2015, **115**, 12138–12204; (*m*) X.-H. Yang, R.-J. Song, Y.-X. Xie and J.-H. Li, *ChemCatChem*, 2016, **8**, 2429–2445; (*n*) Y. Yang, J. Lan and J. You, *Chem. Rev.*, 2017, **117**, 8787–8863.
534 For a review on oxidative cross-coupling for C–O bond formation, see: I. B. Krylov, V. A. Vil' and A. O. Terent'ev, *Beilstein J. Org. Chem.*, 2015, **11**, 92–146.
535 For recent reviews on metal-free oxidative cross-coupling reactions, see: (*a*) R. Samanta, K. Matcha and A. P. Antonchick, *Eur. J. Org. Chem.*, 2013, 5769–5804; (*b*) K. Morimoto, T. Dohi and Y. Kita, *Synlett*, 2017, 1680–1694.
536 For a recent review on electrochemical oxidative cross-couplings, see: S. Tang, Y. Liu and A. Lei, *Chem*, 2018, **4**, 27–45.
537 For recent reviews on radical cross-coupling reactions, see: (*a*) C. Zhang, C. Tang and N. Jiao, *Chem. Soc. Rev.*, 2012, **41**, 3464–3484; (*b*) C. Liu, D. Liu and A. Lei, *Acc. Chem. Res.*, 2014, **47**, 3459–3470; (*c*) X. Zhu and S. Chiba, *Chem. Soc. Rev.*, 2016, **45**, 4504–4523; (*d*) S.-r. Guo, P. S. Kumar and M. Yang, *Adv. Synth. Catal.*, 2017, **359**, 2–25; (*e*) K. Liang and C. Xia, *Chin. J. Chem.*, 2017, **35**, 255–270; (*f*) J.-R. Zhang, L. Xu, Y.-Y. Liao, J.-C. Deng and R.-Y. Tang, *Chin. J. Chem.*, 2017, **35**, 271–279.
538 K. M. Johnston, *Tetrahedron Lett.*, 1967, **8**, 837–839.
539 P. Kočovský, Š. Vyskočil and M. Smrčina, *Chem. Rev.*, 2003, **103**, 3213–3246.
540 M. C. Kozlowski, B. J. Morgan and E. C. Linton, *Chem. Soc. Rev.*, 2009, **38**, 3193–3207.
541 H. Wang, *Chirality*, 2010, **22**, 827–837.
542 A. E. Wendlandt, A. M. Suess and S. S. Stahl, *Angew. Chem., Int. Ed.*, 2011, **50**, 11062–11087.
543 M. C. Kozlowski, *Acc. Chem. Res.*, 2017, **50**, 638–643.
544 J. M. Bobbitt, J. T. Stock, A. Marchand and K. H. Weisgraber, *Chem. Ind.*, 1966, 2127–2128.
545 G. F. Kirkbright, J. T. Stock, R. D. Pugliese and J. M. Bobbitt, *J. Electrochem. Soc.*, 1969, **116**, 219–221.
546 J. M. Bobbitt, K. H. Weisgraber, A. S. Steinfeld and S. G. Weiss, *J. Org. Chem.*, 1970, **35**, 2884–2888.
547 J. M. Bobbitt, H. Yagi, S. Shibuya and J. T. Stock, *J. Org. Chem.*, 1971, **36**, 3006–3010.
548 For anodic oxidation of *p*-cresol, see: T. Kametani, K. Ohkubo and S. Takano, *Chem. Pharm. Bull.*, 1968, **16**, 1095–1097.
549 L. L. Miller, F. R. Stermitz and J. R. Falck, *J. Am. Chem. Soc.*, 1971, **93**, 5941–5942.
550 L. L. Miller, F. R. Stermitz and J. R. Falck, *J. Am. Chem. Soc.*, 1973, **95**, 2651–2656.
551 T. Kametani, K. Shishido and S. Takano, *J. Heterocycl. Chem.*, 1975, **12**, 305–307.
552 J. B. Kerr, T. C. Jempty and L. Miller, *J. Am. Chem. Soc.*, 1979, **101**, 7338–7346.
553 Anodic oxidation of phenolic compounds has also been reported to produce C–O coupled products, see: J. M. Bobbitt and R. C. Hallcher, *J. Chem. Soc. D*, 1971, 543–544.
554 E. Kotani, N. Takeuchi and S. Tobinaga, *J. Chem. Soc., Chem. Commun.*, 1973, 550–551.
555 E. Kotani and S. Tobinaga, *Tetrahedron Lett.*, 1973, **14**, 4759–4762.
556 For a related report on the intramolecular oxidative coupling of phenols, see: M. A. Schwartz, R. A. Holton and S. W. Scott, *J. Am. Chem. Soc.*, 1969, **91**, 2800.
557 I. M. Malkowsky, C. E. Rommel, K. Wedeking, R. Fröhlich, K. Bergander, M. Nieger, C. Quaiser, U. Griesbach, H. Pütter and S. R. Waldvogel, *Eur. J. Org. Chem.*, 2006, 241–245.
558 I. M. Malkowsky, C. E. Rommel, R. Fröhlich, U. Griesbach, H. Pütter and S. R. Waldvogel, *Chem. – Eur. J.*, 2006, **12**, 7482–7488.
559 I. M. Malkowsky, R. Fröhlich, U. Griesbach, H. Pütter and S. R. Waldvogel, *Eur. J. Inorg. Chem.*, 2006, 1690–1697.
560 J. Barjau, P. Königs, O. Kataeva and S. R. Waldvogel, *Synlett*, 2008, 2309–2312.
561 A. Kirste, S. Hayashi, G. Schnakenburg, I. M. Malkowsky, F. Stecker, A. Fischer, T. Fuchigami and S. R. Waldvogel, *Chem. – Eur. J.*, 2011, **17**, 14164–14169.
562 J. Barjau, G. Schnakenburg and S. R. Waldvogel, *Synthesis*, 2011, 2054–2061.
563 M. Mirion, L. Andernach, C. Stobe, J. Barjau, D. Schollmeyer, T. Opatz, A. Lützen and S. R. Waldvogel, *Eur. J. Org. Chem.*, 2015, 4876–4882.
564 For diversity oriented synthesis of polycyclic scaffolds by modification of these motifs, see: J. Barjau, G. Schnakenburg and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2011, **50**, 1415–1419.
565 Anodic oxidation of ortho-halogenated phenols has been reported to produce diaryl ethers, see: (*a*) S. Nishiyama, M. H. Kim and S. Yamamura, *Tetrahedron Lett.*, 1994, **35**, 8397–8400; (*b*) K. Mori, M. Takahashi, S. Yamamura and S. Nishiyama, *Tetrahedron*, 2001, **57**, 5527–5532; (*c*) T. Tanabe, F. Doi, T. Ogamino and S. Nishiyama, *Tetrahedron Lett.*, 2004, **45**, 3477–3480; (*d*) Y. Naito, T. Tanabe, Y. Kawabata, Y. Ishikawa and S. Nishiyama, *Tetrahedron Lett.*, 2010, **51**, 4776–4778; (*e*) K. Uno, T. Tanabe and S. Nishiyama, *ECS Trans.*, 2010, **25**, 91–95; (*f*) Y. Kawabata, Y. Naito, T. Saitoh, K. Kawa, T. Fuchigami and S. Nishiyama, *Eur. J. Org. Chem.*, 2014, 99–104.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

566 I. M. Malkowsky, U. Griesbach, H. Pütter and S. R. Waldvogel, *Eur. J. Org. Chem.*, 2006, 4569–4572.
567 For dehydrogenative coupling reactions using a molybdenum-based anode, see: S. B. Beil, T. Müller, S. B. Sillart, P. Franzmann, A. Bomm, M. Holtkamp, U. Karst, W. Schade and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2018, **57**, 2450–2454.
568 B. Marselli, J. Garcia-Gomez, P.-A. Michaud, M. A. Rodrigo and C. Comninellis, *J. Electrochem. Soc.*, 2003, **150**, D79–D83.
569 R. Fardel, U. Griesbach, H. Pütter and C. Comninellis, *J. Appl. Electrochem.*, 2006, **36**, 249–253.
570 Alkoxy radicals generated through anodic oxidation have also been reported to add to bisdimethoxybenzene and related derivatives, see: (*a*) M. G. Dolson and J. S. Swenton, *J. Am. Chem. Soc.*, 1981, **103**, 2361–2371; (*b*) J. S. Swenton, *Acc. Chem. Res.*, 1983, **16**, 74–81; (*c*) M. C. Carreño and M. Ribagorda, *J. Org. Chem.*, 2000, **65**, 1231–1234; (*d*) C.-C. Zeng and J. Y. Becker, *J. Org. Chem.*, 2004, **69**, 1053–1059; (*e*) S. Quideau, I. Fabre and D. Deffieux, *Org. Lett.*, 2004, **6**, 4571–4573; (*f*) T. Saitoh, E. Suzuki, A. Takasugi, R. Obata, Y. Ishikawa, K. Umezawa and S. Nishiyama, *Bioorg. Med. Chem. Lett.*, 2009, **19**, 5383–5386; (*g*) S. Yajima, T. Saitoh, K. Kawa, K. Nakamura, H. Nagase, Y. Einaga and S. Nishiyama, *Tetrahedron*, 2016, **72**, 8428–8435.
571 A. Kirste, M. Nieger, I. M. Malkowsky, F. Stecker, A. Fischer and S. R. Waldvogel, *Chem. – Eur. J.*, 2009, **15**, 2273–2277.
572 A. Kirste, G. Schnakenburg and S. R. Waldvogel, *Org. Lett.*, 2011, **13**, 3126–3129.
573 T. Yamamoto, B. Riehl, K. Naba, K. Nakahara, A. Wiebe, T. Saitoh, S. R. Waldvogel and Y. Einaga, *Chem. Commun.*, 2018, **54**, 2771–2773.
574 Guaiacol derivatives are potentially attainable aromatic monomers from depolymerization of lignin. For recent reviews on approaches to valorization of lignin, see: (*a*) C. Li, X. Zhao, A. Wang, G. W. Huber and T. Zhang, *Chem. Rev.*, 2015, **115**, 11559–11624; (*b*) M. D. Kärkäs, B. S. Matsuura, T. M. Monos, G. Magallanes and C. R. J. Stephenson, *Org. Biomol. Chem.*, 2016, **14**, 1853–1914; (*c*) M. V. Galkin and J. S. M. Samec, *ChemSusChem*, 2016, **9**, 1544–1558; (*d*) R. Rinaldi, R. Jastrzebski, M. T. Clough, J. Ralph, M. Kennema, P. C. A. Bruijnincx and B. M. Weckhuysen, *Angew. Chem., Int. Ed.*, 2016, **55**, 8164–8215; (*e*) R. E. Key and J. J. Bozell, *ACS Sustainable Chem. Eng.*, 2016, **4**, 5123–5135; (*f*) M. D. Kärkäs, *ChemSusChem*, 2017, **10**, 2111–2115; (*g*) S. Gillet, M. Aguedo, L. Petitjean, A. R. C. Morais, A. M. da Costa Lopes, R. M. Łukasik and P. T. Anastas, *Green Chem.*, 2017, **19**, 4200–4233.
575 B. Elsler, D. Schollmeyer, K. M. Dyballa, R. Franke and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2014, **53**, 5210–5213.
576 A. Wiebe, D. Schollmeyer, K. M. Dyballa, R. Franke and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2016, **55**, 11801–11805.
577 B. Elsler, A. Wiebe, D. Schollmeyer, K. M. Dyballa, R. Franke and S. R. Waldvogel, *Chem. – Eur. J.*, 2015, **21**, 12321–12325.
578 B. Riehl, K. M. Dyballa, R. Franke and S. R. Waldvogel, *Synthesis*, 2017, **49**, 252–259.
579 (*a*) L. Schulz, M. Enders, B. Elsler, D. Schollmeyer, K. M. Dyballa, R. Franke and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2017, **56**, 4877–4881; (*b*) L. Schulz, R. Franke and S. R. Waldvogel, *ChemElectroChem*, DOI: 10.1002/celc.201800422.
580 H. J. Schäfer, *Angew. Chem., Int. Ed.*, 2017, **56**, 15502–15503.
581 For dehydrogenative cross-coupling of naphthylamines with phenols, see: B. Dahms, R. Franke and S. R. Waldvogel, *ChemElectroChem*, 2018, **5**, 1249–1252.
582 A. Kirste, G. Schnakenburg, F. Stecker, A. Fischer and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2010, **49**, 971–975.
583 L. Eberson, M. P. Hartshorn and O. Persson, *Angew. Chem., Int. Ed. Engl.*, 1995, **34**, 2268–2269.
584 L. Eberson, M. P. Hartshorn and O. Persson, *J. Chem. Soc., Perkin Trans. 2*, 1995, 1735–1744.
585 O. Hollóczki, A. Berkessel, J. Mars, M. Mezger, A. Wiebe, S. R. Waldvogel and B. Kirchner, *ACS Catal.*, 2017, **7**, 1846–1852.
586 S. Lips, A. Wiebe, B. Elsler, D. Schollmeyer, K. M. Dyballa, R. Franke and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2016, **55**, 10872–10876.
587 A. Wiebe, B. Riehl, S. Lips, R. Franke and S. R. Waldvogel, *Sci. Adv.*, 2017, **3**, eaao3920.
588 A. Wiebe, S. Lips, D. Schollmeyer, R. Franke and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2017, **56**, 14727–14731.
589 B. Elsler, D. Schollmeyer and S. R. Waldvogel, *Faraday Discuss.*, 2014, **172**, 413–420.
590 For additional electrochemical strategies for synthesis of benzofuran derivatives, see: (*a*) A. R. Fakhari, H. Ahmar, S. S. Hosseiny Davarani, A. Shaabani, S. Nikjah and A. Maleki, *Synth. Commun.*, 2011, **41**, 561–568; (*b*) M. Ameri, A. Asghari, A. Amoozadeh, M. Bakherad and D. Nematollahi, *J. Electrochem. Soc.*, 2014, **161**, G75–G80.
591 G. Moutiers, J. Pinson, F. Terrier and R. Goumont, *Chem. – Eur. J.*, 2001, **7**, 1712–1719.
592 I. Gallardo and G. Guirado, *Eur. J. Org. Chem.*, 2008, 2463–2472.
593 For reviews on nucleophilic aromatic substitution involving electron-deficient arenes, see: (*a*) M. Mąkosza and K. Wojciechowski, *Chem. Rev.*, 2004, **104**, 2631–2666; (*b*) M. Mąkosza, *Chem. Soc. Rev.*, 2010, **39**, 2855–2868; (*c*) A. V. Gulevskaya and I. N. Tyaglivaya, *Russ. Chem. Bull.*, 2012, **61**, 1321–1341; (*d*) M. Mąkosza, *Chem. – Eur. J.*, 2014, **20**, 5536–5545; (*e*) M. Mąkosza and K. Wojciechowski, *Heterocycles*, 2014, **88**, 75–101; (*f*) O. N. Chupakhin and V. N. Charushin, *Tetrahedron Lett.*, 2016, **57**, 2665–2672; (*g*) O. N. Chupakhin and V. N. Charushin, *Pure Appl. Chem.*, 2017, **89**, 1195–1208.
594 H. Cruz, I. Gallardo and G. Guirado, *Green Chem.*, 2011, **13**, 2531–2542.
595 I. Gallardo, G. Guirado and J. Marquet, *J. Org. Chem.*, 2002, **67**, 2548–2555.
596 I. Gallardo, G. Guirado and J. Marquet, *J. Org. Chem.*, 2003, **68**, 631–633.
597 I. Gallardo, G. Guirado and J. Marquet, *Eur. J. Org. Chem.*, 2002, 251–259.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

598 I. Gallardo, G. Guirado and J. Marquet, *Eur. J. Org. Chem.*, 2002, 261–267.
599 I. Gallardo, G. Guirado and J. Marquet, *Chem. – Eur. J.*, 2001, **7**, 1759–1765.
600 H. Cruz, I. Gallardo and G. Guirado, *Eur. J. Org. Chem.*, 2011, 7378–7389.
601 I. Gallardo, G. Guirado and J. Marquet, *J. Org. Chem.*, 2003, **68**, 7334–7341.
602 A. V. Shchepochkin, O. N. Chupakhin, V. N. Charushin, D. V. Steglenko, V. I. Minkin, G. L. Rusinov and A. I. Matern, *RSC Adv.*, 2016, **6**, 77834–77840.
603 O. N. Chupakhin, A. V. Shchepochkin and V. N. Charushin, *Green Chem.*, 2017, **19**, 2931–2935.
604 T. Morofuji, A. Shimizu and J.-i. Yoshida, *Angew. Chem., Int. Ed.*, 2012, **51**, 7259–7262.
605 For trapping of arene radical cations with N-vinylamides using indirect electrolysis, see: L.-J. Li, Y.-Y. Jiang, C. M. Lam, C.-C. Zeng, L.-M. Hu and R. D. Little, *J. Org. Chem.*, 2015, **80**, 11021–11030.
606 T. Arai, H. Tateno, K. Nakabayashi, T. Kashiwagi and M. Atobe, *Chem. Commun.*, 2015, **51**, 4891–4894.
607 P. Wang, S. Tang, P. Huang and A. Lei, *Angew. Chem., Int. Ed.*, 2017, **56**, 3009–3013.
608 For indirect electrochemical arylation of pyrroles, see: G. Sun, S. Ren, X. Zhu, M. Huang and Y. Wan, *Org. Lett.*, 2016, **18**, 544–547.
609 P. Wang, S. Tang and A. Lei, *Green Chem.*, 2017, **19**, 2092–2095.
610 X.-Y. Qian, S.-Q. Li, J. Song and H.-C. Xu, *ACS Catal.*, 2017, **7**, 2730–2734.
611 For early studies on electrochemically-mediated cyclization of N-arylthioamides, see: (*a*) M. Lać an, K. Jakopčić, V. Rogić, S. Damoni, O. Rogić and I. Tabaković, *Synth. Commun.*, 1974, **4**, 219–224; (*b*) I. Tabaković, M. Trkovnik, M. Batušić and K. Tabaković, *Synthesis*, 1979, 590–592.
612 For a related electrochemical approach in flow, see: A. A. Folgueiras-Amador, X.-Y. Qian, H.-C. Xu and T. Wirth, *Chem. – Eur. J.*, 2018, **24**, 487–491.
613 Z.-J. Wu and H.-C. Xu, *Angew. Chem., Int. Ed.*, 2017, **56**, 4734–4738.
614 Z.-J. Wu, S.-R. Li, H. Long and H.-C. Xu, *Chem. Commun.*, 2018, **54**, 4601–4604.
615 U. Jahn and P. Hartmann, *Chem. Commun.*, 1998, 209–210.
616 U. Jahn, M. Müller and S. Aussieker, *J. Am. Chem. Soc.*, 2000, **122**, 5212–5213.
617 U. Jahn, P. Hartmann and E. Kaasalainen, *Org. Lett.*, 2004, **6**, 257–260.
618 F. Kafka, M. Holan, D. Hidasová, R. Pohl, I. Císařová, B. Klepetářová and U. Jahn, *Angew. Chem., Int. Ed.*, 2014, **53**, 9944–9948.
619 For an alternative electrochemical approach to access oxindoles, see: Y.-Y. Jiang, S. Liang, C.-C. Zeng, L.-M. Hu and B.-G. Sun, *Green Chem.*, 2016, **18**, 6311–6319.
620 For synthesis of oxindoles involving cathodic reduction, see: R. Munusamy, K. S. Dhathathreyan, K. K. Balasubramanian and C. S. Venkatachalam, *J. Chem. Soc., Perkin Trans. 2*, 2001, 1154–1166.
621 H. Ding, P. L. DeRoy, C. Perreault, A. Larivée, A. Siddiqui, C. G. Caldwell, S. Harran and P. G. Harran, *Angew. Chem., Int. Ed.*, 2015, **54**, 4818–4822.
622 A. W. G. Burgett, Q. Li, Q. Wei and P. G. Harran, *Angew. Chem., Int. Ed.*, 2003, **42**, 4961–4966.
623 J.-C. Zhao, S.-M. Yu, H.-B. Qiu and Z.-J. Yao, *Tetrahedron*, 2014, **70**, 3197–3210.
624 K. Liu, S. Tang, P. Huang and A. Lei, *Nat. Commun.*, 2017, **8**, 775.
625 A. Ronlán and V. D. Parker, *J. Chem. Soc. C*, 1971, 3214–3218.
626 A. Ronlán, *J. Chem. Soc. D*, 1971, 1643–1645.
627 J. A. Richards, P. E. Whitson and D. H. Evans, *J. Electroanal. Chem.*, 1975, **63**, 311–327.
628 A. Nilsson, U. Palmquist, T. Pettersson and A. Ronlán, *J. Chem. Soc., Perkin Trans. 1*, 1978, 696–707.
629 K. Ponsold and H. Kasch, *Tetrahedron Lett.*, 1979, **20**, 4465–4466.
630 H. Eickhoff, G. Jung and A. Rieker, *Tetrahedron*, 2001, **57**, 353–364.
631 K. Chiba, M. Fukuda, S. Kim, Y. Kitano and M. Tada, *J. Org. Chem.*, 1999, **64**, 7654–7656.
632 See also: Y. Imada, Y. Yamaguchi, N. Shida, Y. Okada and K. Chiba, *Chem. Commun.*, 2017, **53**, 3960–3963.
633 For seminal studies on the [3+2] cycloaddition between phenoxonium ions and electron-rich alkenes through anodic oxidation, see: (*a*) Y. Shizuri, K. Nakamura and S. Yamamura, *J. Chem. Soc., Chem. Commun.*, 1985, 530–531; (*b*) B. D. Gates, P. Dalidowicz, A. Tebben, S. Wang and J. S. Swenton, *J. Org. Chem.*, 1992, **57**, 2135–2143.
634 For a seminal study on the [3+2] cycloaddition between phenoxonium ions and vinyl sulfides through anodic oxidation, see: M. L. Kerns, S. M. Conroy and J. S. Swenton, *Tetrahedron Lett.*, 1994, **35**, 7529–7532.
635 For a related [3+2] cycloaddition of anodically generated radical intermediates, see: (*a*) A. Nishiyama, H. Eto, Y. Terada, M. Iguchi and S. Yamamura, *Chem. Pharm. Bull.*, 1983, **31**, 2834–2844; (*b*) T. Sumi, T. Saitoh, K. Natsui, T. Yamamoto, M. Atobe, Y. Einaga and S. Nishiyama, *Angew. Chem., Int. Ed.*, 2012, **51**, 5443–5446.
636 K. Chiba, M. Jinno, R. Kuramoto and M. Tada, *Tetrahedron Lett.*, 1998, **39**, 5527–5530.
637 S. Kim, S. Noda, K. Hayashi and K. Chiba, *Org. Lett.*, 2008, **10**, 1827–1829.
638 Y. Okada, M. Sugai and K. Chiba, *J. Org. Chem.*, 2016, **81**, 10922–10929.
639 H. Wakamatsu, Y. Okada, M. Sugai, S. R. Hussaini and K. Chiba, *Asian J. Org. Chem.*, 2017, **6**, 1584–1588.
640 S. Kim, K. Hirose, J. Uematsu, Y. Mikami and K. Chiba, *Chem. – Eur. J.*, 2012, **18**, 6284–6288.
641 Y. Okada, K. Shimada, Y. Kitano and K. Chiba, *Eur. J. Org. Chem.*, 2014, 1371–1375.
642 M. Iguchi, A. Nishiyama, Y. Terada and S. Yamamura, *Chem. Lett.*, 1978, 451–454.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

643 A. Nishiyama, H. Eto, Y. Terada, M. Iguchi and S. Yamamura, *Chem. Pharm. Bull.*, 1983, **31**, 2820–2833.
644 Y. Shizuri and S. Yamamura, *Tetrahedron Lett.*, 1983, **24**, 5011–5012.
645 S. Yamamura, Y. Shizuri, H. Shigemori, Y. Okuno and M. Ohkubo, *Tetrahedron*, 1991, **47**, 635–644.
646 D. Deffieux, I. Fabre, C. Courseille and S. Quideau, *J. Org. Chem.*, 2002, **67**, 4458–4465.
647 D. Deffieux, I. Fabre, A. Titz, J.-M. Léger and S. Quideau, *J. Org. Chem.*, 2004, **69**, 8731–8738.
648 L. Rycek, J. J. Hayward, M. A. Latif, J. Tanko, R. Simionescu and T. Hudlicky, *J. Org. Chem.*, 2016, **81**, 10930–10941.
649 H. Takakura and S. Yamamura, *Tetrahedron Lett.*, 1998, **39**, 3717–3720.
650 H. Takakura and S. Yamamura, *Tetrahedron Lett.*, 1999, **40**, 299–302.
651 H. R. El-Seedi, S. Yamamura and S. Nishiyama, *Tetrahedron*, 2002, **58**, 7485–7489.
652 H. R. El-Seedi, S. Yamamura and S. Nishiyama, *Tetrahedron Lett.*, 2002, **43**, 3301–3304.
653 S. Yamamura and S. Nishiyama, *Synlett*, 2002, 533–543.
654 U. Palmquist, A. Ronlán and V. D. Parker, *Acta Chem. Scand., Ser. B*, 1974, **28**, 267–269.
655 I. G. C. Coutts, N. J. Culbert, M. Edwards, J. A. Hadfield, D. R. Musto, V. H. Pavlidis and D. J. Richards, *J. Chem. Soc., Perkin Trans. 1*, 1985, 1829–1836.
656 G. W. Morrow and J. S. Swenton, *Tetrahedron Lett.*, 1987, **28**, 5445–5448.
657 G. W. Morrow, Y. Chen and J. S. Swenton, *Tetrahedron*, 1991, **47**, 655–664.
658 J. S. Swenton, K. Carpenter, Y. Chen, M. L. Kerns and G. W. Morrow, *J. Org. Chem.*, 1993, **58**, 3308–3316.
659 J. S. Swenton, A. Callinan, Y. Chen, J. J. Rohde, M. L. Kerns and G. W. Morrow, *J. Org. Chem.*, 1996, **61**, 1267–1274.
660 A. Hutinec, A. Ziogas, M. El-Mobayed and A. Rieker, *J. Chem. Soc., Perkin Trans. 1*, 1998, 2201–2208.
661 K. Mori, S. Yamamura and S. Nishiyama, *Tetrahedron*, 2001, **57**, 5533–5542.
662 T. Ogamino and S. Nishiyama, *Tetrahedron*, 2003, **59**, 9419–9423.
663 T. Ogamino, Y. Ishikawa and S. Nishiyama, *Heterocycles*, 2003, **61**, 73–78.
664 T. Ogamino, R. Obata and S. Nishiyama, *Tetrahedron Lett.*, 2006, **47**, 727–731.
665 See also: T. Ogamino and S. Nishiyama, *Tetrahedron Lett.*, 2005, **46**, 1083–1086.
666 J.-F. Cheng, S. Nishiyama and S. Yamamura, *Chem. Lett.*, 1990, 1591–1594.
667 S. Nishiyama, J.-F. Cheng, X. L. Tao and S. Yamamura, *Tetrahedron Lett.*, 1991, **32**, 4151–4154.
668 X. L. Tao, J.-F. Cheng, S. Nishiyama and S. Yamamura, *Tetrahedron*, 1994, **50**, 2017–2028.
669 Y. Wada, H. Fujioka and Y. Kita, *Mar. Drugs*, 2010, **8**, 1394–1416.
670 T. Ogamino, S. Ohnishi, Y. Ishikawa, T. Sugai, R. Obata and S. Nishiyama, *Sci. Technol. Adv. Mater.*, 2006, **7**, 175–183.
671 F. Doi, T. Ogamino, T. Sugai and S. Nishiyama, *Tetrahedron Lett.*, 2003, **44**, 4877–4880.
672 F. Doi, T. Ogamino, T. Sugai and S. Nishiyama, *Synlett*, 2003, 411–413.
673 F. Doi, T. Ohara, T. Ogamino, K. Higashinakasu, K. Hasegawa and S. Nishiyama, *Bull. Chem. Soc. Jpn.*, 2004, **77**, 2257–2263.
674 F. Doi, T. Ohara, T. Ogamino, T. Sugai, K. Higashinakasu, K. Yamada, H. Shigemori, K. Hasegawa and S. Nishiyama, *Phytochemistry*, 2004, **65**, 1405–1411.
675 A. Maleki and D. Nematollahi, *Org. Lett.*, 2011, **13**, 1928–1931.
676 D. Nematollahi and B. Dadpou, *Chin. Chem. Lett.*, 2011, **22**, 1067–1070.
677 F. Varmaghani and D. Nematollahi, *J. Phys. Org. Chem.*, 2012, **25**, 511–514.
678 A. Maleki, D. Nematollahi, F. Rasouli and A. Zeinodini-Meimand, *Green Chem.*, 2016, **18**, 672–675.
679 D. Nematollahi, F. Ghasemi, M. Sharafi-Kolkeshvandi and F. Varmaghani, *J. Electroanal. Chem.*, 2016, **775**, 299–305.
680 V. M. Shanmugam, K. Kulangiappar, M. Ramaprakash, D. Vasudevan, R. S. Kumar, D. Velayutham and T. Raju, *Tetrahedron Lett.*, 2017, **58**, 2294–2297.
681 S. B. H. Lang, D. Nematollahi, A. R. Massah, M. Sharafi-Kolkeshvandi and H. Alizadeh, *J. Electrochem. Soc.*, 2017, **164**, G87–G91.
682 D. Nematollahi and M. Hesari, *J. Electroanal. Chem.*, 2005, **577**, 197–203.
683 S. Osati, S. S. Hosseiny Davarani, N. Safari and M. H. Banitaba, *Electrochim. Acta*, 2011, **56**, 9426–9432.
684 D. Nematollahi, Z. Zohdijamil and H. Salehzadeh, *J. Electroanal. Chem.*, 2014, **720–721**, 156–161.
685 S. Khazalpour, D. Nematollahi and M. R. Pajohi-Alamoti, *New J. Chem.*, 2015, **39**, 6734–6737.
686 M. Sharafi-Kolkeshvandi, D. Nematollahi, F. Nikpour, M. Bayat and E. Soltani, *Electrochim. Acta*, 2016, **222**, 845–855.
687 D. Nematollahi, H. Hesari, H. Salehzadeh, M. Hesari and S. Momeni, *C. R. Chim.*, 2016, **19**, 357–362.
688 M. Jamshidi and D. Nematollahi, *ACS Sustainable Chem. Eng.*, 2017, **5**, 9423–9430.
689 S. Dadkhah-Tehrani, M. Shabani-Nooshabadi and M. Moradian, *J. Iran. Chem. Soc.*, 2018, **15**, 171–179.
690 D. Nematollahi, A. Afkhami, E. Tammari, T. Shariatmanesh, M. Hesariac and M. Shojaeifard, *Chem. Commun.*, 2007, 162–164.
691 D. Nematollahi and H. Khoshsafar, *Tetrahedron*, 2009, **65**, 4742–4750.
692 D. Nematollahi and E. Tammari, *Electrochim. Acta*, 2005, **50**, 3648–3654.
693 A. B. Moghaddam, M. R. Ganjali, P. Norouzi and M. Latifi, *Chem. Pharm. Bull.*, 2006, **54**, 1391–1396.
694 M. Rafiee and D. Nematollahi, *Chem. Pharm. Bull.*, 2007, **55**, 915–917.
695 D. Nematollahi, M. Alimoradi and M. Rafiee, *J. Phys. Org. Chem.*, 2007, **20**, 49–54.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

696 D. Nematollahi, M. Rafiee and H. R. Khavasi, *Bull. Chem. Soc. Jpn.*, 2008, **81**, 1505–1511.
697 D. Nematollahi and H. Shayani-jam, *J. Org. Chem.*, 2008, **73**, 3428–3434.
698 D. Nematollahi, S. Dehdashtian and A. Niazi, *J. Electroanal. Chem.*, 2008, **616**, 79–86.
699 D. Nematollahi and S. Dehdashtian, *Tetrahedron Lett.*, 2008, **49**, 645–649.
700 S. S. Hosseiny Davarani, N. Sheijooni Fumani, H. Arvin-Nezhad and F. Moradi, *Tetrahedron Lett.*, 2008, **49**, 710–714.
701 C.-C. Zeng, D.-W. Ping, Y.-S. Xu, L.-M. Hu and R.-G. Zhong, *Eur. J. Org. Chem.*, 2009, 5832–5840.
702 S. S. Hosseiny Davarani, N. Sheijooni Fumani, S. Vahidi, M.-A. Tabatabaei and H. Arvin-Nezhad, *J. Heterocycl. Chem.*, 2010, **47**, 40–45.
703 C.-C. Zeng, D.-W. Ping, L.-M. Hu, X.-Q. Song and R.-G. Zhong, *Org. Biomol. Chem.*, 2010, **8**, 2465–2472.
704 Y.-X. Bai, D.-W. Ping, R. D. Little, H.-Y. Tian, L.-M. Hu and C.-C. Zeng, *Tetrahedron*, 2011, **67**, 9334–9341.
705 H. Salehzadeh, D. Nematollahi and M. Rafiee, *J. Electroanal. Chem.*, 2011, **650**, 226–232.
706 N.-T. Zhang, X.-G. Gao, C.-C. Zeng, L.-M. Hu, H.-Y. Tian and Y.-B. She, *RSC Adv.*, 2012, **2**, 298–306.
707 A. Amani, S. Khazalpour and D. Nematollahi, *J. Electroanal. Chem.*, 2012, **670**, 36–41.
708 D. Nematollahi, S. S. Hosseiny Davarani and P. Mirahmadpour, *ACS Sustainable Chem. Eng.*, 2014, **2**, 579–583.
709 M. Ameri, A. Asghari, A. Amoozadeh, M. Bakherad and D. Nematollahi, *Chin. Chem. Lett.*, 2014, **25**, 1607–1610.
710 D. Nematollahi, S. Momeni and S. Khazalpour, *Electrochim. Acta*, 2014, **147**, 310–318.
711 A. Asghari, M. Ameri, A. A. Ziarati, S. Radmannia, A. Amoozadeh, B. Barfi and L. Boutorabi, *Chin. Chem. Lett.*, 2015, **26**, 681–684.
712 E. Salahifar, D. Nematollahi, M. Bayat, A. Mahyari and H. Amiri Rudbari, *Org. Lett.*, 2015, **17**, 4666–4669.
713 C.-C. Zeng, C.-F. Liu, J. Zeng and R.-G. Zhong, *J. Electroanal. Chem.*, 2007, **608**, 85–90.
714 D. Nematollahi and F. Varmaghani, *Electrochim. Acta*, 2008, **53**, 3350–3355.
715 D. Nematollahi and R. Esmaili, *Tetrahedron Lett.*, 2010, **51**, 4862–4865.
716 H.-L. Xiao, C.-W. Yang, N.-T. Zhang, C.-C. Zeng, L.-M. Hu, H.-Y. Tian and R. D. Little, *Tetrahedron*, 2013, **69**, 658–663.
717 H. Salehzadeh, M. Rafiee, D. Nematollahi and H. Beiginejad, *Curr. Org. Chem.*, 2013, **17**, 848–852.
718 H. Beiginejad and D. Nematollahi, *J. Org. Chem.*, 2014, **79**, 6326–6329.
719 A. Alizadeh, M. M. Khodaei, M. Fakhari and M. Shamsuddin, *RSC Adv.*, 2014, **4**, 20781–20788.
720 S. Kaihani, H. Salehzadeh and D. Nematollahi, *Electrochim. Acta*, 2015, **157**, 166–174.
721 E. Salahifar and D. Nematollahi, *New J. Chem.*, 2015, **39**, 3852–3858.
722 S. Khazalpour and D. Nematollahi, *Green Chem.*, 2015, **17**, 3508–3514.
723 M. Sharafi-Kolkeshvandi, D. Nematollahi, F. Nikpour and E. Salahifar, *New J. Chem.*, 2016, **40**, 5442–5447.
724 M. Jamshidi, D. Nematollahi, M. Bayat and E. Salahifar, *J. Electrochem. Soc.*, 2016, **163**, G211–G218.
725 D. Nematollahi, S. Khazalpour, M. Ranjbar and S. Momeni, *Sci. Rep.*, 2017, **7**, 4436.
726 S. Momeni and D. Nematollahi, *Sci. Rep.*, 2017, **7**, 41963.
727 M. Sharafi-Kolkeshvandi, D. Nematollahi and F. Nikpour, *Synthesis*, 2017, 1555–1560.
728 D. Nematollahi and E. Tammari, *J. Org. Chem.*, 2005, **70**, 7769–7772.
729 M. Shamsipur, S. S. Hosseiny Davarani, M. Nasiri-Aghdam and D. Nematollahi, *Electrochim. Acta*, 2006, **51**, 3327–3331.
730 C.-C. Zeng, D.-W. Ping, S.-C. Zhang, R.-G. Zhong and J. Y. Becker, *J. Electroanal. Chem.*, 2008, **622**, 90–96.
731 F.-J. Liu, C.-C. Zeng, D.-W. Ping, Y.-L. Cai and R.-G. Zhong, *Chin. J. Chem.*, 2008, **26**, 1651–1655.
732 D. Nematollahi, J. Azizian, M. Sargordan-Arani, M. Hesari, S. Jameh-Bozorghi, A. Alizadeh, L. Fotouhi and B. Mirza, *Chem. Pharm. Bull.*, 2008, **56**, 1562–1566.
733 M. M. Khodaei, A. Alizadeh and N. Pakravan, *J. Org. Chem.*, 2008, **73**, 2527–2532.
734 A. R. Fakhari, S. S. Hosseiny Davarani, H. Ahmar and S. Makarem, *J. Appl. Electrochem.*, 2008, **38**, 1743–1747.
735 P. R. Jamaat, S. S. Hosseiny Davarani, N. Sheijooni Fumani, L. Masoumi and N. Safari, *J. Porphyrins phthalocyanines*, 2008, **12**, 85–93.
736 C.-C. Zeng, F.-J. Liu, D.-W. Ping, L.-M. Hu, Y.-L. Cai and R.-G. Zhong, *Tetrahedron*, 2009, **65**, 4505–4512.
737 C.-C. Zeng, F.-J. Liu, D.-W. Ping, Y.-L. Cai, R.-G. Zhong and J. Y. Becker, *J. Electroanal. Chem.*, 2009, **625**, 131–137.
738 A. R. Fakhari, S. S. Hosseiny Davarani, H. Ahmar, K. Hasheminasab and H. R. Khavasi, *J. Heterocycl. Chem.*, 2009, **46**, 443–446.
739 D. Nematollahi, J. Azizian, M. Sargordan-Arani, M. Hesari and B. Mirza, *J. Heterocycl. Chem.*, 2009, **46**, 1000–1002.
740 A. Amani and D. Nematollahi, *J. Org. Chem.*, 2012, **77**, 11302–11306.
741 M. Ameri, A. Asghari, A. Amoozadeh, H. Daneshinejad and D. Nematollahi, *Chin. Chem. Lett.*, 2014, **25**, 797–801.
742 D. Nematollahi, S. Mahdinia, P. Karimi, H. Salehzadeh and S. Kaihani, *RSC Adv.*, 2015, **5**, 29209–29213.
743 E. Tammari, A. Nezhadali and S. Lotfi, *Electroanalysis*, 2015, **27**, 1693–1698.
744 O. Ghaderi, A. Asghari, M. Ameri and M. Rajabi, *Chem. Lett.*, 2016, **45**, 430–432.
745 I. Tabaković, Z. Grujić and Z. Bejtović, *J. Heterocycl. Chem.*, 1983, **20**, 635–638.
746 D. Nematollahi, D. Habibi, M. Rahmati and M. Rafiee, *J. Org. Chem.*, 2004, **69**, 2637–2640.
747 D. Nematollahi and M. Rafiee, *Green Chem.*, 2005, **7**, 638–644.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

748 S. S. Hosseiny Davarani, D. Nematollahi, M. Shamsipur, N. M. Najafi, L. Masoumi and S. Ramyar, *J. Org. Chem.*, 2006, **71**, 2139–2142.
749 S. S. Hosseiny Davarani, N. Mashkouri Najafi, S. Ramyar, L. Masoumi and M. Shamsipur, *Chem. Pharm. Bull.*, 2006, **54**, 959–962.
750 M. Shamsipur, S. S. Hosseiny Davarani and D. Nematollahi, *J. Heterocycl. Chem.*, 2006, **43**, 1673–1677.
751 A. Bayandori Moghaddam, F. Kobarfard, S. S. Hosseiny Davarani, D. Nematollahi, M. Shamsipur and A. R. Fakhari, *J. Electroanal. Chem.*, 2006, **586**, 161–166.
752 D. Nematollahi, A. Amani and E. Tammari, *J. Org. Chem.*, 2007, **72**, 3646–3651.
753 S. S. Hosseiny Davarani, M. Shamsipur, D. Nematollahi, S. Ramyar and L. Masoumi, *Chem. Pharm. Bull.*, 2007, **55**, 1198–1202.
754 S. Osati, N. Safari, M. Kalate-Bojdi and S. S. Hosseiny Davarani, *J. Electroanal. Chem.*, 2011, **655**, 120–127.
755 M. Ameri, A. Asghari, A. Amoozadeh, M. Bakherad and D. Nematollahi, *Tetrahedron Lett.*, 2015, **56**, 2141–2144.
756 N. Pakravan, D. Habibi, F. Varmaghani and M. Rahmati, *J. Electroanal. Chem.*, 2017, **801**, 206–214.
757 D. Nematollahi, A. R. Atlasi-Pak and R. Esmaili, *Helv. Chim. Acta*, 2012, **95**, 1605–1612.
758 S. S. Hosseiny Davarani, A. R. Fakhari, N. Sheijooni Fumani and M. Kalate-bojdi, *Electrochem. Commun.*, 2008, **10**, 1765–1768.
759 L. Fotouhi, M. Mosavi, M. M. Heravi and D. Nematollahi, *Tetrahedron Lett.*, 2006, **47**, 8553–8557.
760 E. Blattes, M.-B. Fleury and M. Largeron, *Electrochim. Acta*, 2005, **50**, 4902–4910.
761 D. Xu, A. Chiaroni, M.-B. Fleury and M. Largeron, *J. Org. Chem.*, 2006, **71**, 6374–6381.
762 H. Salehzadeh, D. Nematollahi and H. Hesari, *Green Chem.*, 2013, **15**, 2441–2446.
763 See also: Y. Shih, C. Ke, C. Pan and Y. Huang, *RSC Adv.*, 2013, **3**, 7330–7336.
764 Z. Grujić, I. Tabaković and M. Trkovnik, *Tetrahedron Lett.*, 1976, **17**, 4823–4824.
765 S. M. Golabi and D. Nematollahi, *J. Electroanal. Chem.*, 1997, **420**, 127–134.
766 H. Ghadimi, A. S. M. Ali, N. Mohamed and S. Ab Ghani, *J. Electrochem. Soc.*, 2012, **159**, E127–E131.
767 C.-C. Zeng, F.-J. Liu, D.-W. Ping, L.-M. Hu, Y.-L. Cai and R.-G. Zhong, *J. Org. Chem.*, 2009, **74**, 6386–6389.
768 A. Asghari, M. Ameri, S. Radmannia, M. Rajabi, M. Bakherad and D. Nematollahi, *J. Electroanal. Chem.*, 2014, **733**, 47–52.
769 M. Shabani-Nooshabadi, M. Moradian and S. Dadkhah-Tehrani, *Bull. Chem. Soc. Jpn.*, 2017, **90**, 68–73.
770 S. S. Hosseiny Davarani, A. R. Fakhari, A. Shaabani, H. Ahmar, A. Maleki and N. Sheijooni Fumani, *Tetrahedron Lett.*, 2008, **49**, 5622–5624.
771 M. Sharafi-Kolkeshvandi, D. Nematollahi, F. Nikpour, B. Dadpou and H. Alizadeh, *Electrochim. Acta*, 2016, **214**, 147–155.
772 S. S. Hosseiny Davarani, M. Kalate Bojdi and A. Mehdinia, *Chem. Pharm. Bull.*, 2011, **59**, 1209–1213.
773 D. Habibi, D. Nematollahi, H. M. Pordanjani and M. R. Sadeghi, *J. Heterocycl. Chem.*, 2015, **52**, 197–200.
774 S. Shahrokhian and A. Hamzehloei, *Electrochem. Commun.*, 2003, **5**, 706–710.
775 D. Nematollahi, M. S. Workentin and E. Tammari, *Chem. Commun.*, 2006, 1631–1633.
776 D. Nematollahi and A. Ghorbani, *J. Electroanal. Chem.*, 2008, **624**, 310–314.
777 K. Bechgaard and V. D. Parker, *J. Am. Chem. Soc.*, 1972, **94**, 4749–4750.
778 J.-M. Chapuzet and J. Simonet, *Tetrahedron*, 1991, **47**, 791–798.
779 S. R. Waldvogel and D. Mirk, *Tetrahedron Lett.*, 2000, **41**, 4769–4772.
780 M. Largeron, A. Neudorffer and M.-B. Fleury, *Angew. Chem., Int. Ed.*, 2003, **42**, 1026–1029.
781 M. Largeron, A. Chiaroni and M.-B. Fleury, *Chem. – Eur. J.*, 2008, **14**, 996–1003.
782 M. Largeron and M.-B. Fleury, *Org. Lett.*, 2009, **11**, 883–886.
783 K. L. Jensen, P. T. Franke, L. T. Nielsen, K. Daasbjerg and K. A. Jørgensen, *Angew. Chem., Int. Ed.*, 2010, **49**, 129–133.
784 For a related enantioselective approach for α-alkylation of aldehydes through the merger of electro- and organo-catalysis, see: X.-H. Ho, S.-i. Mho, H. Kang and H.-Y. Jang, *Eur. J. Org. Chem.*, 2010, 4436–4441.
785 For electrochemical synthesis of enaminones, see: (*a*) K. Xu, Z. Zhang, P. Qian, Z. Zha and Z. Wang, *Chem. Commun.*, 2015, **51**, 11108–11111; (*b*) Y. Li, H. Gao, Z. Zhang, P. Qian, M. Bi, Z. Zha and Z. Wang, *Chem. Commun.*, 2016, **52**, 8600–8603.
786 D. Koch and H. Schäfer, *Angew. Chem., Int. Ed. Engl.*, 1973, **12**, 245–246.
787 See also: (*a*) Ž. Saničanin and I. Tabaković, *Electrochim. Acta*, 1988, **33**, 1601–1604; (*b*) I. Tabaković, *Electrochim. Acta*, 1995, **40**, 2809–2813.
788 W. Eilenberg and H. J. Schäfer, *Tetrahedron Lett.*, 1984, **25**, 5023–5026.
789 S. Tang, X. Gao and A. Lei, *Chem. Commun.*, 2017, **53**, 3354–3356.
790 Z. He, W. Liu and Z. Li, *Chem. – Asian J.*, 2011, **6**, 1340–1343.
791 For preparation of polycyclic indoles through anodic oxidation, see: (*a*) X.-G. Gao, N.-T. Zhanga, C.-C. Zeng, Y.-D. Liu, L.-M. Hu and H.-Y. Tian, *Curr. Org. Synth.*, 2014, **11**, 141–148; (*b*) M. Ameri, A. Asghari, A. Amoozadeh and M. Bakherad, *Chin. Chem. Lett.*, 2017, **28**, 1031–1034.
792 For synthetic approaches to indoles involving electro-reduction, see: (*a*) S. C. Mishra and R. A. Mishra, *J. Electrochem. Soc. India*, 1990, **39**, 51–53; (*b*) F. LeStrat, J. A. Murphy and M. Hughes, *Org. Lett.*, 2002, **4**, 2735–2738; (*c*) L. Bouchard, I. Marcotte, J. M. Chapuzet and J. Lessard, *Can. J. Chem.*, 2003, **81**, 1108–1118; (*d*) P. Du and D. G. Peters, *J. Electrochem. Soc.*, 2010,

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

157, F167–F172; (*e*) P. Du, J. L. Brosmer and D. G. Peters, *Org. Lett.*, 2011, **13**, 4072–4075.
793 K. D. Moeller and D. G. New, *Tetrahedron Lett.*, 1994, **35**, 2857–2860.
794 D. G. New, Z. Tesfai and K. D. Moeller, *J. Org. Chem.*, 1996, **61**, 1578–1598.
795 D. A. Frey, N. Wu and K. D. Moeller, *Tetrahedron Lett.*, 1996, **37**, 8317–8320.
796 C. R. Whitehead, E. H. Sessions, I. Ghiviriga and D. L. Wright, *Org. Lett.*, 2002, **4**, 3763–3765.
797 J. B. Sperry and D. L. Wright, *Tetrahedron Lett.*, 2005, **46**, 411–414.
798 J. B. Sperry, C. R. Whitehead, I. Ghiviriga, R. M. Walczak and D. L. Wright, *J. Org. Chem.*, 2004, **69**, 3726–3734.
799 A. Redden and K. D. Moeller, *Org. Lett.*, 2011, **13**, 1678–1681.
800 J. B. Sperry and D. L. Wright, *J. Am. Chem. Soc.*, 2005, **127**, 8034–8035.
801 I. Kuwajima and H. Urabe, *Tetrahedron Lett.*, 1981, **22**, 5191–5194.
802 A. Padwa and S. S. Murphree, *Org. Prep. Proced. Int.*, 1991, **23**, 545–568.
803 C. O. Kappe, S. S. Murphree and A. Padwa, *Tetrahedron*, 1997, **53**, 14179–14233.
804 D. L. Wright, C. R. Whitehead, E. H. Sessions, I. Ghiviriga and D. A. Frey, *Org. Lett.*, 1999, **1**, 1535–1538.
805 D. L. Wright, *Prog. Heterocycl. Chem.*, 2005, **17**, 1–32.
806 M. A. Battiste, P. M. Pelphrey and D. L. Wright, *Chem. – Eur. J.*, 2006, **12**, 3438–3447.
807 M. Harmata, *Chem. Commun.*, 2010, **46**, 8904–8922.
808 J. B. Sperry and D. L. Wright, *Tetrahedron*, 2006, **62**, 6551–6557.
809 J. B. Sperry, I. Ghiviriga and D. L. Wright, *Chem. Commun.*, 2006, 194–196.
810 See also: C. Adouama, R. Keyrouz, G. Pilet, C. Monnereau, D. Gueyrard, T. Noël and M. Médebielle, *Chem. Commun.*, 2017, **53**, 5653–5656.
811 H. Wu and K. D. Moeller, *Org. Lett.*, 2007, **9**, 4599–4602.
812 K. Chiba, T. Miura, S. Kim, Y. Kitano and M. Tada, *J. Am. Chem. Soc.*, 2001, **123**, 11314–11315.
813 T. Miura, S. Kim, Y. Kitano, M. Tada and K. Chiba, *Angew. Chem., Int. Ed.*, 2006, **45**, 1461–1463.
814 M. Arata, T. Miura and K. Chiba, *Org. Lett.*, 2007, **9**, 4347–4350.
815 Y. Okada, R. Akaba and K. Chiba, *Org. Lett.*, 2009, **11**, 1033–1035.
816 Y. Okada and K. Chiba, *Electrochim. Acta*, 2010, **55**, 4112–4119.
817 Y. Okada and K. Chiba, *Electrochim. Acta*, 2011, **56**, 1037–1042.
818 Y. Okada, A. Nishimoto, R. Akaba and K. Chiba, *J. Org. Chem.*, 2011, **76**, 3470–3476.
819 Y. Yamaguchi, Y. Okada and K. Chiba, *J. Org. Chem.*, 2013, **78**, 2626–2638.
820 J. C. Moore, E. S. Davies, D. A. Walsh, P. Sharma and J. E. Moses, *Chem. Commun.*, 2014, **50**, 12523–12525.
821 Y. Okada, Y. Yamaguchi, A. Ozaki and K. Chiba, *Chem. Sci.*, 2016, **7**, 6387–6393.
822 A. Ozaki, Y. Yamaguchi, Y. Okada and K. Chiba, *ChemElectroChem*, 2017, **4**, 1852–1855.
823 Y. Okada, R. Akaba and K. Chiba, *Tetrahedron Lett.*, 2009, **50**, 5413–5416.
824 For a review on redox-tag processes, see: Y. Okada and K. Chiba, *Chem. Rev.*, 2018, **118**, 4592–4630.
825 M. A. Cismesia and T. P. Yoon, *Chem. Sci.*, 2015, **6**, 5426–5434.
826 For an overview on metal-catalyzed C–N bond forming cross-coupling reactions, see: (*a*) J. Bariwal and E. Van der Eycken, *Chem. Soc. Rev.*, 2013, **42**, 9283–9303; (*b*) C. Sambiagio, S. P. Marsden, A. J. Blacker and P. C. McGowan, *Chem. Soc. Rev.*, 2014, **43**, 3525–3550; (*c*) P. Ruiz-Castillo and S. L. Buchwald, *Chem. Rev.*, 2016, **116**, 12564–12649; (*d*) S. Bhunia, G. G. Pawar, S. V. Kumar, Y. Jiang and D. Ma, *Angew. Chem., Int. Ed.*, 2017, **56**, 16136–16179; (*e*) J. Schranck and A. Tlili, *ACS Catal.*, 2018, **8**, 405–418.
827 For recent reviews on catalytic methods for C–H amination, see: (*a*) K. Shin, H. Kim and S. Chang, *Acc. Chem. Res.*, 2015, **48**, 1040–1052; (*b*) J. Jiao, K. Murakami and K. Itami, *ACS Catal.*, 2016, **6**, 610–633; (*c*) H. Kim and S. Chang, *ACS Catal.*, 2016, **6**, 2341–2351; (*d*) Y. Zhou, J. Yuan, Q. Yang, Q. Xiao and Y. Peng, *ChemCatChem*, 2016, **8**, 2178–2192; (*e*) Y. Park, Y. Kim and S. Chang, *Chem. Rev.*, 2017, **117**, 9247–9301; (*f*) H. Kim and S. Chang, *Acc. Chem. Res.*, 2017, **50**, 482–486.
828 R. I. McDonald, G. Liu and S. S. Stahl, *Chem. Rev.*, 2011, **111**, 2981–3019.
829 T. J. Donohoe, C. K. A. Callens, A. Flores, A. R. Lacy and A. H. Rathi, *Chem. – Eur. J.*, 2011, **17**, 58–76.
830 S. E. Denmark, W. E. Kuester and M. T. Burk, *Angew. Chem., Int. Ed.*, 2012, **51**, 10938–10953.
831 S. Pan and T. Shibata, *ACS Catal.*, 2013, **3**, 704–712.
832 S. R. Chemler and M. T. Bovino, *ACS Catal.*, 2013, **3**, 1076–1091.
833 M. Chiarucci and M. Bandini, *Beilstein J. Org. Chem.*, 2013, **9**, 2586–2614.
834 A. M. Beauchemin, *Org. Biomol. Chem.*, 2013, **11**, 7039–7050.
835 J. C.-H. Yim and L. L. Schafer, *Eur. J. Org. Chem.*, 2014, 6825–6840.
836 L. Huang, M. Arndt, K. Gooßen, H. Heydt and L. J. Gooßen, *Chem. Rev.*, 2015, **115**, 2596–2697.
837 E. Bernoud, C. Lepori, M. Mellah, E. Schulz and J. Hannedouche, *Catal. Sci. Technol.*, 2015, **5**, 2017–2037.
838 S. M. Coman and V. I. Parvulescu, *Org. Process Res. Dev.*, 2015, **19**, 1327–1355.
839 V. Rodriguez-Ruiz, R. Carlino, S. Bezzenine-Lafollée, R. Gil, D. Prim, E. Schulz and J. Hannedouche, *Dalton Trans.*, 2015, **44**, 12029–12059.
840 M. T. Pirnot, Y.-M. Wang and S. L. Buchwald, *Angew. Chem., Int. Ed.*, 2016, **55**, 48–57.
841 J. R. Coombs and J. P. Morken, *Angew. Chem., Int. Ed.*, 2016, **55**, 2636–2649.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

842 G. E. M. Crisenza and J. F. Bower, *Chem. Lett.*, 2016, **45**, 2–9.
843 Z. Sorádová and R. Šebesta, *ChemCatChem*, 2016, **8**, 2581–2588.
844 Z. Dong, Z. Ren, S. J. Thompson, Y. Xu and G. Dong, *Chem. Rev.*, 2017, **117**, 9333–9403.
845 M. Patel, R. K. Saunthwal and A. K. Verma, *Acc. Chem. Res.*, 2017, **50**, 240–254.
846 X.-W. Lan, N.-X. Wang and Y. Xing, *Eur. J. Org. Chem.*, 2017, 5821–5851.
847 C. Lepori and J. Hannedouche, *Synthesis*, 2017, 1158–1167.
848 P. Kalck and M. Urrutigoïty, *Chem. Rev.*, 2018, **118**, 3833–3861.
849 K. D. Moeller, M. R. Marzabadi, D. G. New, M. Y. Chiang and S. Keith, *J. Am. Chem. Soc.*, 1990, **112**, 6123–6124.
850 C. M. Hudson, M. R. Marzabadi, K. D. Moeller and D. G. New, *J. Am. Chem. Soc.*, 1991, **113**, 7372–7385.
851 K. D. Moeller and L. V. Tinao, *J. Am. Chem. Soc.*, 1992, **114**, 1033–1041.
852 L. V. Tinao-Wooldridge, K. D. Moeller and C. M. Hudson, *J. Org. Chem.*, 1994, **59**, 2381–2389.
853 D. A. Frey, S. H. Krishna Reddy and K. D. Moeller, *J. Org. Chem.*, 1999, **64**, 2805–2813.
854 S. H. Krishna Reddy, K. Chiba, Y. Sun and K. D. Moeller, *Tetrahedron*, 2001, **57**, 5183–5197.
855 K. D. Moeller and C. M. Hudson, *Tetrahedron Lett.*, 1991, **32**, 2307–2310.
856 D. A. Frey, J. A. Marx and K. D. Moeller, *Electrochim. Acta*, 1997, **42**, 1967–1970.
857 S. H. Krishna Reddy and K. D. Moeller, *Tetrahedron Lett.*, 1998, **39**, 8027–8030.
858 Y. Sun and K. D. Moeller, *Tetrahedron Lett.*, 2002, **43**, 7159–7161.
859 Y.-t. Huang and K. D. Moeller, *Org. Lett.*, 2004, **6**, 4199–4202.
860 Y.-t. Huang and K. D. Moeller, *Tetrahedron*, 2006, **62**, 6536–6550.
861 F. Tang and K. D. Moeller, *J. Am. Chem. Soc.*, 2007, **129**, 12414–12415.
862 A. Redden, R. J. Perkins and K. D. Moeller, *Angew. Chem., Int. Ed.*, 2013, **52**, 12865–12868.
863 A. Sutterer and K. D. Moeller, *J. Am. Chem. Soc.*, 2000, **122**, 5636–5637.
864 Y. Sun, B. Liu, J. Kao, D. A. d'Avignon and K. D. Moeller, *Org. Lett.*, 2001, **3**, 1729–1732.
865 B. Liu and K. D. Moeller, *Tetrahedron Lett.*, 2001, **42**, 7163–7165.
866 S. Duan and K. D. Moeller, *J. Am. Chem. Soc.*, 2002, **124**, 9368–9369.
867 B. Liu, S. Duan, A. C. Sutterer and K. D. Moeller, *J. Am. Chem. Soc.*, 2002, **124**, 10101–10111.
868 J. D. Brandt and K. D. Moeller, *Org. Lett.*, 2005, **7**, 3553–3556.
869 J. D. Brandt and K. D. Moeller, *Heterocycles*, 2006, **67**, 621–628.
870 H.-C. Xu, J. D. Brandt and K. D. Moeller, *Tetrahedron Lett.*, 2008, **49**, 3868–3871.
871 G. Xu and K. D. Moeller, *Org. Lett.*, 2010, **12**, 2590–2593.
872 J. A. Smith and K. D. Moeller, *Org. Lett.*, 2013, **15**, 5818–5821.
873 R. J. Perkins, H.-C. Xu, J. M. Campbell and K. D. Moeller, *Beilstein J. Org. Chem.*, 2013, **9**, 1630–1636.
874 J. A. Smith, G. Xu, R. Feng, J. W. Janetka and K. D. Moeller, *Electroanalysis*, 2016, **28**, 2808–2817.
875 For a different approach to electrochemical oxyfunctionalization of alkenes, see: P. Röse, S. Emge, J.-i. Yoshida and G. Hilt, *Beilstein J. Org. Chem.*, 2015, **11**, 174–183.
876 H.-C. Xu and K. D. Moeller, *J. Am. Chem. Soc.*, 2008, **130**, 13542–13543.
877 H.-C. Xu and K. D. Moeller, *J. Am. Chem. Soc.*, 2010, **132**, 2839–2844.
878 For an approach to access six-membered ring systems, see: H.-C. Xu and K. D. Moeller, *Org. Lett.*, 2010, **12**, 5174–5177.
879 H.-C. Xu and K. D. Moeller, *Org. Lett.*, 2010, **12**, 1720–1723.
880 J. M. Campbell, H.-C. Xu and K. D. Moeller, *J. Am. Chem. Soc.*, 2012, **134**, 18338–18344.
881 For early work on kinetic studies of enol ether radical cations, see: J. H. Horner, E. Taxil and M. Newcomb, *J. Am. Chem. Soc.*, 2002, **124**, 5402–5410.
882 L. A. Anderson, A. Redden and K. D. Moeller, *Green Chem.*, 2011, **13**, 1652–1654.
883 H.-C. Xu and K. D. Moeller, *Angew. Chem., Int. Ed.*, 2010, **49**, 8004–8007.
884 Y. Ashikari, T. Nokami and J.-i. Yoshida, *Org. Biomol. Chem.*, 2013, **11**, 3322–3331.
885 N. Fu, G. S. Sauer, A. Saha, A. Loo and S. Lin, *Science*, 2017, **357**, 575–579.
886 J. B. Parry, N. Fu and S. Lin, *Synlett*, 2018, 257–265.
887 M. Zanda, *SYNFORM*, 2017, A202–A204.
888 For seminal work on electrochemical diazidation of alkenes, see: H. Schäfer, *Angew. Chem., Int. Ed. Engl.*, 1970, **9**, 158–159.
889 N. Fu, G. S. Sauer and S. Lin, *J. Am. Chem. Soc.*, 2017, **139**, 15548–15553.
890 K.-Y. Ye, G. Pombar, N. Fu, G. S. Sauer, I. Keresztes and S. Lin, *J. Am. Chem. Soc.*, 2018, **140**, 2438–2441.
891 For a recent review on radical difunctionalization of alkenes, see: G. S. Sauer and S. Lin, *ACS Catal.*, 2018, **8**, 5175–5187.
892 C. Tian, L. Massignan, T. H. Meyer and L. Ackermann, *Angew. Chem., Int. Ed.*, 2018, **57**, 2383–2387.
893 S. Tang, D. Wang, Y. Liu, L. Zeng and A. Lei, *Nat. Commun.*, 2018, **9**, 798.
894 N. Sauermann, R. Mei and L. Ackermann, *Angew. Chem., Int. Ed.*, 2018, **57**, 5090–5094.
895 X. Gao, P. Wang, L. Zeng, S. Tang and A. Lei, *J. Am. Chem. Soc.*, 2018, **140**, 4195–4199.
896 F. Xu, Y.-J. Li, C. Huang and H.-C. Xu, *ACS Catal.*, 2018, **8**, 3820–3824.

 This journal is © The Royal Society of Chemistry 2018

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

897 M. E. Wolff, *Chem. Rev.*, 1963, **63**, 55–64.
898 R. S. Neale, *Synthesis*, 1971, 1–15.
899 L. Stella, *Angew. Chem., Int. Ed. Engl.*, 1983, **22**, 337–350.
900 G. Majetich and K. Wheless, *Tetrahedron*, 1995, **51**, 7095–7129.
901 S. Z. Zard, *Synlett*, 1996, 1148–1154.
902 W. R. Bowman, A. J. Fletcher and G. B. S. Potts, *J. Chem. Soc., Perkin Trans. 1*, 2002, 2747–2762.
903 S. Z. Zard, *Chem. Soc. Rev.*, 2008, **37**, 1603–1618.
904 J. L. Jeffrey and R. Sarpong, *Chem. Sci.*, 2013, **4**, 4092–4106.
905 J.-R. Chen, X.-Q. Hu, L.-Q. Lu and W.-J. Xiao, *Chem. Soc. Rev.*, 2016, **45**, 2044–2056.
906 T. Xiong and Q. Zhang, *Chem. Soc. Rev.*, 2016, **45**, 3069–3087.
907 R. K. Rit, M. Shankar and A. K. Sahoo, *Org. Biomol. Chem.*, 2017, **15**, 1282–1293.
908 K. Murakami, G. J. P. Perry and K. Itami, *Org. Biomol. Chem.*, 2017, **15**, 6071–6075.
909 T. Siu and A. K. Yudin, *J. Am. Chem. Soc.*, 2002, **124**, 530–531.
910 T. Siu, C. J. Picard and A. K. Yudin, *J. Org. Chem.*, 2005, **70**, 932–937.
911 For electrochemical aziridination proceeding *via* initial alkene oxidation, see: J. Li, W. Huang, J. Chen, L. He, X. Cheng and G. Li, *Angew. Chem., Int. Ed.*, 2018, **57**, 5695–5698.
912 T. Siu and A. K. Yudin, *Org. Lett.*, 2002, **4**, 1839–1842.
913 H.-C. Xu, J. M. Campbell and K. D. Moeller, *J. Org. Chem.*, 2014, **79**, 379–391.
914 P. Xiong, H.-H. Xu and H.-C. Xu, *J. Am. Chem. Soc.*, 2017, **139**, 2956–2959.
915 L. Zhu, P. Xiong, Z.-Y. Mao, Y.-H. Wang, X. Yan, X. Lu and H.-C. Xu, *Angew. Chem., Int. Ed.*, 2016, **55**, 2226–2229.
916 F. Xu, L. Zhu, S. Zhu, X. Yan and H.-C. Xu, *Chem. – Eur. J.*, 2014, **20**, 12740–12744.
917 A. A. Folgueiras-Amador, K. Philipps, S. Guilbaud, J. Poelakker and T. Wirth, *Angew. Chem., Int. Ed.*, 2017, **56**, 15446–15450.
918 T. Gieshoff, A. Kehl, D. Schollmeyer, K. D. Moeller and S. R. Waldvogel, *Chem. Commun.*, 2017, **53**, 2974–2977.
919 Z.-W. Hou, Z.-Y. Mao, H.-B. Zhao, Y. Y. Melcamu, X. Lu, J. Song and H.-C. Xu, *Angew. Chem., Int. Ed.*, 2016, **55**, 9168–9172.
920 Z.-W. Hou, Z.-Y. Mao and H.-C. Xu, *Synlett*, 2017, 1867–1872.
921 Z.-W. Hou, Z.-Y. Mao, Y. Y. Melcamu, X. Lu and H.-C. Xu, *Angew. Chem., Int. Ed.*, 2018, **57**, 1636–1639.
922 For electrochemical homo-coupling of imidazopyridine-based heterocycles, see: Y. Gao, Y. Wang, J. Zhou, H. Mei and J. Han, *Green Chem.*, 2018, **20**, 583–587.
923 Z.-W. Hou, Z.-Y. Mao, J. Song and H.-C. Xu, *ACS Catal.*, 2017, **7**, 5810–5813.
924 For ferrocene-mediated electrochemical difluoromethylarylation of alkynes, see: P. Xiong, H.-H. Xu, J. Song and H.-C. Xu, *J. Am. Chem. Soc.*, 2018, **140**, 2460–2464.
925 For ferrocene-mediated intermolecular dehydrogenative cross-coupling of N-alkyloxyamides with xanthenes, see: M.-Y. Lin, K. Xu, Y.-Y. Jiang, Y.-G. Liu, B.-G. Sun and C.-C. Zeng, *Adv. Synth. Catal.*, 2018, **360**, 1665–1672.
926 H.-B. Zhao, Z.-W. Hou, Z.-J. Liu, Z.-F. Zhou, J. Song and H.-C. Xu, *Angew. Chem., Int. Ed.*, 2017, **56**, 587–590.
927 For synthesis of benzimidazoles through the merger of electrochemistry and Co catalysis, see: Y.-L. Lai, J.-S. Ye and J.-M. Huang, *Chem. – Eur. J.*, 2016, **22**, 5425–5429.
928 H.-B. Zhao, Z.-J. Liu, J. Song and H.-C. Xu, *Angew. Chem., Int. Ed.*, 2017, **56**, 12732–12735.
929 K. Subramanian, S. L. Yedage and B. M. Bhanage, *J. Org. Chem.*, 2017, **82**, 10025–10032.
930 B. R. Rosen, E. W. Werner, A. G. O'Brien and P. S. Baran, *J. Am. Chem. Soc.*, 2014, **136**, 5571–5574.
931 S. R. Waldvogel and B. Janza, *Angew. Chem., Int. Ed.*, 2014, **53**, 7122–7123.
932 A. Kehl, T. Gieshoff, D. Schollmeyer and S. R. Waldvogel, *Chem. – Eur. J.*, 2018, **24**, 590–593.
933 T. Gieshoff, D. Schollmeyer and S. R. Waldvogel, *Angew. Chem., Int. Ed.*, 2016, **55**, 9437–9440.
934 T. Gieshoff, A. Kehl, D. Schollmeyer, K. D. Moeller and S. R. Waldvogel, *J. Am. Chem. Soc.*, 2017, **139**, 12317–12324.
935 H. Lund, *Acta Chem. Scand.*, 1957, **11**, 1323–1330.
936 G. Manning, V. D. Parker and R. N. Adams, *J. Am. Chem. Soc.*, 1969, **91**, 4584–4585.
937 C. V. Ristagno and H. J. Shine, *J. Org. Chem.*, 1971, **36**, 4050–4055.
938 H. N. Blount, *J. Electroanal. Chem.*, 1973, **42**, 271–274.
939 M. Masui, H. Ohmori, H. Sayo, A. Ueda and C. Ueda, *J. Chem. Soc., Perkin Trans. 2*, 1976, 1180–1183.
940 J. F. Evans and H. N. Blount, *J. Org. Chem.*, 1976, **41**, 516–519.
941 B. Reitstöen and V. D. Parker, *Acta Chem. Scand.*, 1992, **46**, 464–468.
942 Y. Li, S. Asaoka, T. Yamagishi and T. Iyoda, *Electrochemistry*, 2004, **72**, 171–174.
943 See also: C. J. Schlesener and J. K. Kochi, *J. Org. Chem.*, 1984, **49**, 3142–3150.
944 S. R. Waldvogel and S. Möhle, *Angew. Chem., Int. Ed.*, 2015, **54**, 6398–6399.
945 For copper-promoted electrochemical amination of boronic acids, see: H.-L. Qi, D.-S. Chen, J.-S. Ye and J.-M. Huang, *J. Org. Chem.*, 2013, **78**, 7482–7487.
946 T. Zincke, *Justus Liebigs Ann. Chem.*, 1904, **330**, 361–374.
947 T. Zincke, G. Heuser and W. Möller, *Justus Liebigs Ann. Chem.*, 1904, **333**, 296–345.
948 T. Zincke and W. Würker, *Justus Liebigs Ann. Chem.*, 1904, **338**, 107–141.
949 S. Herold, S. Möhle, M. Zirbes, F. Richter, H. Nefzger and S. R. Waldvogel, *Eur. J. Org. Chem.*, 2016, 1274–1278.
950 S. Möhle, S. Herold, F. Richter, H. Nefzger and S. R. Waldvogel, *ChemElectroChem*, 2017, **4**, 2196–2210.
951 L. J. Wesenberg, S. Herold, A. Shimizu, J.-i. Yoshida and S. R. Waldvogel, *Chem. – Eur. J.*, 2017, **23**, 12096–12099.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

952 T. Morofuji, A. Shimizu and J.-i. Yoshida, *Chem. – Eur. J.*, 2015, **21**, 3211–3214.
953 For a related dehydrogenative intramolecular C–N cross-coupling reaction, see: Z. Ye, M. Ding, Y. Wu, Y. Li, W. Hua and F. Zhang, *Green Chem.*, 2018, **20**, 1732–1737.
954 T. Morofuji, A. Shimizu and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2014, **136**, 4496–4499.
955 For synthesis of pyren-1-yl azoliums *via* electrochemical oxidative C–N coupling, see: G. de Robillard, O. Makni, H. Cattey, J. Andrieu and C. H. Devillers, *Green Chem.*, 2015, **17**, 4669–4679.
956 For an example on anodic flow oxidation of toluene derivatives to benzyl amides, see: M. A. Kabeshov, B. Musio and S. V. Ley, *React. Chem. Eng.*, 2017, **2**, 822–825.
957 T. Morofuji, A. Shimizu and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2015, **137**, 9816–9819.
958 R. Hayashi, A. Shimizu, Y. Song, Y. Ashikari, T. Nokami and J.-i. Yoshida, *Chem. – Eur. J.*, 2017, **23**, 61–64.
959 R. Hayashi, A. Shimizu and J.-i. Yoshida, *J. Am. Chem. Soc.*, 2016, **138**, 8400–8403.
960 J. Wu, Y. Zhou, Y. Zhou, C.-W. Chiang and A. Lei, *ACS Catal.*, 2017, **7**, 8320–8323.
961 For electrochemical C(sp^3)–H functionalization of ethers with indoles, see: K.-S. Du and J.-M. Huang, *Org. Lett.*, 2018, **20**, 2911–2915.
962 E. A. Merritt and B. Olofsson, *Angew. Chem., Int. Ed.*, 2009, **48**, 9052–9070.
963 F. V. Singh and T. Wirth, *Chem. – Asian J.*, 2014, **9**, 950–971.
964 G. Maertens, C. L'Homme and S. Canesi, *Front. Chem.*, 2015, **2**, 115.
965 A. Yoshimura and V. V. Zhdankin, *Chem. Rev.*, 2016, **116**, 3328–3435.
966 Y. Li, D. P. Hari, M. V. Vita and J. Waser, *Angew. Chem., Int. Ed.*, 2016, **55**, 4436–4454.
967 L. Wang and J. Liu, *Eur. J. Org. Chem.*, 2016, 1813–1824.
968 A. Yoshimura, M. S. Yusubov and V. V. Zhdankin, *Org. Biomol. Chem.*, 2016, **14**, 4771–4781.
969 F. C. Sousa e Silva, A. F. Tierno and S. E. Wengryniuk, *Molecules*, 2017, **22**, 780.
970 D. Kajiyama, K. Inoue, Y. Ishikawa and S. Nishiyama, *Tetrahedron*, 2010, **66**, 9779–9784.
971 D. Kajiyama, T. Saitoh, S. Yamaguchi and S. Nishiyama, *Synthesis*, 2012, 1667–1671.
972 Y. Amano, K. Inoue and S. Nishiyama, *Synlett*, 2008, 134–136.
973 Y. Amano and S. Nishiyama, *Tetrahedron Lett.*, 2006, **47**, 6505–6507.
974 Y. Amano and S. Nishiyama, *Heterocycles*, 2008, **75**, 1997–2003.
975 K. Inoue, Y. Ishikawa and S. Nishiyama, *Org. Lett.*, 2010, **12**, 436–439.
976 See also: D. Kajiyama, T. Saitoh and S. Nishiyama, *Electrochemistry*, 2013, **81**, 319–324.
977 T. Broese and R. Francke, *Org. Lett.*, 2016, **18**, 5896–5899.
978 For the use of soluble polymediators and polyelectrolytes in electrosynthesis, see: B. Schille, N. O. Giltzau and R. Francke, *Angew. Chem., Int. Ed.*, 2018, **57**, 422–426.
979 O. Koleda, T. Broese, J. Noetzel, M. Roemelt, E. Suna and R. Francke, *J. Org. Chem.*, 2017, **82**, 11669–11681.
980 W.-C. Li, C.-C. Zeng, L.-M. Hu, H.-Y. Tian and R. D. Little, *Adv. Synth. Catal.*, 2013, **355**, 2884–2890.
981 W.-J. Gao, W.-C. Li, C.-C. Zeng, H.-Y. Tian, L.-M. Hu and R. D. Little, *J. Org. Chem.*, 2014, **79**, 9613–9618.
982 See also: E. O. Yurko, T. V. Gryaznova, K. V. Kholin, V. V. Khrizanforova and Y. H. Budnikova, *Dalton Trans.*, 2018, **47**, 190–196.
983 S. Liang, C.-C. Zeng, H.-Y. Tian, B.-G. Sun, X.-G. Luo and F.-z. Ren, *J. Org. Chem.*, 2016, **81**, 11565–11573.
984 W.-J. Gao, C. M. Lam, B.-G. Sun, R. D. Little and C.-C. Zeng, *Tetrahedron*, 2017, **73**, 2447–2454.
985 For synthesis of potential building blocks from lignin using electrochemistry, see: B. H. Nguyen, R. J. Perkins, J. A. Smith and K. D. Moeller, *J. Org. Chem.*, 2015, **80**, 11953–11962.
986 See also: D. T. Rensing, B. H. Nguyen and K. D. Moeller, *Org. Chem. Front.*, 2016, **3**, 1236–1240.
987 S. Liang, C.-C. Zeng, H.-Y. Tian, B.-G. Sun, X.-G. Luo and F.-z. Ren, *Adv. Synth. Catal.*, 2018, **360**, 1444–1452.
988 L.-S. Kang, M.-H. Luo, C. M. Lam, L.-M. Hu, R. D. Little and C.-C. Zeng, *Green Chem.*, 2016, **18**, 3767–3774.
989 X.-J. Pan, J. Gao and G.-Q. Yuan, *Tetrahedron*, 2015, **71**, 5525–5530.
990 For a review on iodine-catalyzed oxidative coupling reactions, see: D. Liu and A. Lei, *Chem. – Asian J.*, 2015, **10**, 806–823.
991 J. Chen, W.-Q. Yan, C. M. Lam, C.-C. Zeng, L.-M. Hu and R. D. Little, *Org. Lett.*, 2015, **17**, 986–989.
992 K. Fujimoto, Y. Tokuda, Y. Matsubara, H. Maekawa, T. Mizuno and I. Nishiguchi, *Tetrahedron Lett.*, 1995, **36**, 7483–7486.
993 W.-q. Yan, M.-y. Lin, R. D. Little and C.-C. Zeng, *Tetrahedron*, 2017, **73**, 764–770.
994 S. Liang, C.-C. Zeng, X.-G. Luo, F.-z. Ren, H.-Y. Tian, B.-G. Sun and R. D. Little, *Green Chem.*, 2016, **18**, 2222–2230.
995 A. Studer and D. P. Curran, *Nat. Chem.*, 2014, **6**, 765–773.
996 B. H. Nguyen, A. Redden and K. D. Moeller, *Green Chem.*, 2014, **16**, 69–72.
997 B. H. Nguyen, R. J. Perkins, J. A. Smith and K. D. Moeller, *Beilstein J. Org. Chem.*, 2015, **11**, 280–287.
998 K. N. Singh and A. K. Shukla, *J. Sci. Ind. Res.*, 1999, **58**, 327–331.
999 D. Pletcher, *Electrochem. Commun.*, 2018, **88**, 1–4.
1000 K.-P. Ho, T. H. Chan and K.-Y. Wong, *Green Chem.*, 2006, **8**, 900–905.
1001 S. Sengmany, A. Vitu-Thiebaud, E. Le Gall, S. Condon, E. Léonel, C. Thobie-Gautier, M. Pipelier, J. Lebreton and D. Dubreuil, *J. Org. Chem.*, 2013, **78**, 370–379.
1002 S. Sengmany, S. Vasseur, A. Lajnef, E. Le Gall and E. Léonel, *Eur. J. Org. Chem.*, 2016, 4865–4871.

Open Access Article. Published on 18 June 2018. Downloaded on 5/7/2026 10:36:06 AM.
This article is licensed under a Creative Commons Attribution 3.0 Unported Licence.

1003 H. Shimakoshi, Z. Luo, T. Inaba and Y. Hisaeda, *Dalton Trans.*, 2016, **45**, 10173–10180.
1004 C. Li, Y. Kawamata, H. Nakamura, J. C. Vantourout, Z. Liu, Q. Hou, D. Bao, J. T. Starr, J. Chen, M. Yan and P. S. Baran, *Angew. Chem., Int. Ed.*, 2017, **56**, 13088–13093.
1005 R. J. Perkins, D. J. Pedro and E. C. Hansen, *Org. Lett.*, 2017, **19**, 3755–3758.
1006 H. Li, C. P. Breen, H. Seo, T. F. Jamison, Y.-Q. Fang and M. M. Bio, *Org. Lett.*, 2018, **20**, 1338–1341.
1007 For a discussion on electrobiorefineries, see: F. Harnisch and C. Urban, *Angew. Chem., Int. Ed.*, DOI: 10.1002/anie.201711727.
1008 All electrochemical reactions can be divided into two half-reactions, oxidation and reduction. It is thus possible to combine two desirable half-reactions, allowing these reactions to be carried out simultaneously.
1009 For a review on paired electrolysis, see: C. A. Paddon, M. Atobe, T. Fuchigami, P. He, P. Watts, S. J. Haswell, G. J. Pritchard, S. D. Bull and F. Marken, *J. Appl. Electrochem.*, 2006, **36**, 617–634.
1010 For the use of paired electrolysis for reduction of CO_2, see: M. J. Llorente, B. H. Nguyen, C. P. Kubiak and K. D. Moeller, *J. Am. Chem. Soc.*, 2016, **138**, 15110–15113.
1011 For synthesis of diacids and diols in a divergent fashion using paired electrolysis, see: R. Matthessen, J. Fransaer, K. Binnemans and D. E. De Vos, *ChemElectroChem*, 2015, **2**, 73–76.
1012 For synthesis of protected homoallylic alcohols by combining a cathodic Barbier reaction with the Shono oxidation using convergent paired electrolysis, see: G. Hilt, *Angew. Chem., Int. Ed.*, 2003, **42**, 1720–1721.
1013 J.-i. Yoshida, *Chem. Commun.*, 2005, 4509–4516.
1014 F. J. del Campo, *Electrochem. Commun.*, 2014, **45**, 91–94.
1015 M. Atobe, H. Tateno and Y. Matsumura, *Chem. Rev.*, 2018, **118**, 4541–4572.
1016 D. Pletcher, R. A. Green and R. C. D. Brown, *Chem. Rev.*, 2018, **118**, 4573–4591.
1017 M. Atobe, *Curr. Opin. Electrochem.*, 2017, **2**, 1–6.
1018 M. Navarro, *Curr. Opin. Electrochem.*, 2017, **2**, 43–52.
1019 C. Gütz, A. Stenglein and S. R. Waldvogel, *Org. Process Res. Dev.*, 2017, **21**, 771–778.