


 Cite this: *RSC Adv.*, 2020, **10**, 10883

 Received 29th February 2020
 Accepted 4th March 2020

 DOI: 10.1039/d0ra02016h
rsc.li/rsc-advances

Solvent-free and room temperature microwave-assisted direct C7 allylation of indolines via sequential C–H and C–C activation†

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A Ru or Rh-catalyzed efficient and atom-economic C7 allylation of indolines with vinylcyclopropanes was developed via sequential C–H and C–C activation. A wide range of substrates were well tolerated to afford the corresponding allylated indolines in high yields and *E/Z* selectivities under microwave irradiation. The obtained allylated indolines could further undergo transformations to afford various value-added chemicals. Importantly, this reaction proceeded at room temperature under solvent-free conditions.

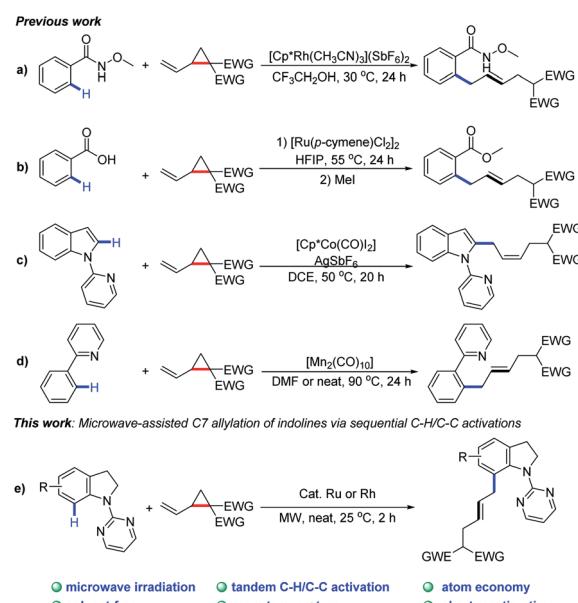
The development of sustainable methodologies is attractive for access to complex molecular architectures in organic chemistry.¹ In recent years, various non-conventional techniques, such as microwave irradiation, sonochemistry, mechanical grinding and photochemistry, have achieved remarkable success.² In particular, microwaves have shown unique advantages with regards to reaction times, energy efficiency, temperature, and reaction media.³ On the other hand, transition-metal-catalyzed activation of C–H⁴ and C–C⁵ bonds has been considered as an ideal method for the formation of C–C and C–X bonds. Nevertheless, transition-metal-catalyzed C–H or C–C bond activation under the above non-conventional techniques remains to be explored. It is thus highly imperative to develop a practical strategy in combination of C–H or C–C activation and microwave irradiation.⁶

Recently, there have significant advances in C–H activation technology by merging C–H functionalization with challenging C–C cleavage strategies.⁷ Since the pioneering work by Bergman and co-workers⁸ on the sequential C–H and C–C bond activation, many research groups, including Dong,⁹ Ackermann,¹⁰ Li,¹¹ Cramer,¹² and others¹³ have contributed to C–H/C–C activation. In this content, certain small strained rings are often utilized as an effective synthons to undergo ring-opening reactions driven by strain-release energy.¹⁴ Very recently, VCPs (vinylcyclopropanes) have been reported as allyl reagents to access various (hetero)aromatic derivatives through sequential C–H and C–C activation (Scheme 1a–d).¹⁵

As a continuation of our interest in chelation-directed reactions and novel methods for C–H functionalization,¹⁶ we herein report a Ru or Rh-catalyzed C–7 allylation of indolines under

microwave irradiation using VCPs as the allylating agents (Scheme 1e). This transformation possesses great synthetic potential from the viewpoint of green and sustainable chemistry. Notable features of our protocol include (1) C–H/C–C activation with VCPs by microwave irradiation, (2) broad substrate scope with good regio- and *E/Z* selectivities, (3) high atom economy, and (4) high efficiency (2 h) at room temperature under solvent-free conditions.

We initiated our investigation by choosing indoline **1a** and VCP **2a** as model substrates under microwave irradiation conditions (Table 1). To our delight, the allylated product **3aa**



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† Electronic supplementary information (ESI) available. See DOI: 10.1039/d0ra02016h

Scheme 1 Sequential C–H/C–C activations using VCPs.

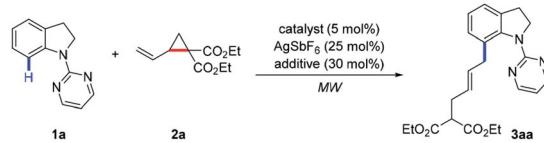


Table 1 Optimization of the reaction conditions^a

Entry	Catalyst (mol%)	Additive (mol%)	T (°C)	Yield (%)
1	[Ru(<i>p</i> -cymene)Cl ₂] ₂	AdCOOH	90	40
2	RuCl ₃ ·3H ₂ O	AdCOOH	90	N.R
3	[Cp*RuCl ₂] ₂	AdCOOH	90	N.R
4	[Ru(<i>p</i> -cymene)Cl ₂] ₂	MesCOOH	90	47
5	[Ru(<i>p</i> -cymene)Cl ₂] ₂	AcOH	90	40
6	[Ru(<i>p</i> -cymene)Cl ₂] ₂	NaOAc	90	20
7	[Ru(<i>p</i> -cymene)Cl ₂] ₂	PivONa·H ₂ O	90	21
8	[Ru(<i>p</i> -cymene)Cl ₂] ₂	DABCO	90	Trace
9 ^b	[Ru(<i>p</i> -cymene)Cl ₂] ₂	MesCOOH	90	57
10 ^b	[Ru(<i>p</i> -cymene)Cl ₂] ₂	MesCOOH	70	68
11 ^b	[Ru(<i>p</i> -cymene)Cl ₂] ₂	MesCOOH	50	83
12 ^b	[Ru(<i>p</i> -cymene)Cl ₂] ₂	MesCOOH	25	65
13 ^{b,c}	[Ru(<i>p</i> -cymene)Cl ₂] ₂	MesCOOH	25	87 (>20 : 1) ^e
14 ^{c,d}	[Cp*Rh(CH ₃ CN) ₃](SbF ₆) ₂	AdCOOH	80	78 (10 : 1) ^e

^a Reaction conditions: **1a** (0.2 mmol), **2a** (0.4 mmol), [Ru(*p*-cymene)Cl₂]₂ (5 mol%), additive (30 mol%), MW, 1 h, 90 °C. ^b MesCOOH (50 mol%).

^c *t* = 2 h. ^d [Cp*Rh(CH₃CN)₃](SbF₆)₂ (8 mol%). ^e The *E* : *Z* ratio was determined by ¹H NMR analysis. MW = microwave irradiation.

Table 2 Substrate scope of indolines^{a,b}

		[Ru(<i>p</i> -cymene)Cl ₂] ₂ (5 mol %), AgSbF ₆ (25 mol %), MesCOOH (50 mol %), MW, 25 °C, 2 h	
1	2a	DG = 2-Pyrimidine	3
3aa , 87% (>20:1) [Ru] 78% (10:1) [Rh]	3ba , 96% (>20:1) [Ru] 92% (11:1) [Rh]	3ca , 81% (10:1) [Ru] 68% (6:1) [Rh]	3da , 90% (7:1) [Ru] 81% (5:1) [Rh]
3ea , 65% (5:1) [Ru] 58% (4:1) [Rh]	3fa , 75% (3:1) [Ru] 52% (3:1) [Rh]	3ga , 91% (>20:1) [Ru] 90% (8:1) [Rh]	3ha , 90% (>20:1) [Ru] 91% (13:1) [Rh]
3ia , 92% (>20:1) [Ru] 86% (13:1) [Rh]	3ja , 93% (>20:1) [Ru] 74% (>20:1) [Rh]	3ka , 94% (>20:1) [Ru] 72% (>20:1) [Rh]	3la , 96% (>20:1) [Ru] 76% (>20:1) [Rh]
3ma , 54% (>20:1) [Ru] 56% (14:1) [Rh]	3na , 86% (11:1) [Ru] 84% (11:1) [Rh]	3oa , 90% (>20:1) [Ru] 76% (14:1) [Rh]	3pa , 80% (>20:1) [Ru] 87% (>17:1) [Rh]
3qa , 91% (10:1) [Ru] 81% (>20:1) [Rh]	3ra , 90% (>20:1) [Ru] 84% (>20:1) [Rh]	3sa , 73% (>20:1) [Ru] 54% (13:1) [Rh]	3ta , 60% (4:1) [Ru] 71% (>20:1) [Rh]
3ua , 82% (5:1) [Ru] 70% (6:1) [Rh]	3va , 89% (>20:1) [Ru] 72% (>20:1) [Rh]	3wa , 95% (5:1) [Ru] 80% (>20:1) [Rh]	3xa , 91% (14:1) [Ru] 71% (>20:1) [Rh]
3za , 78% (>20:1) [Ru] 65% (>20:1) [Rh]	3za , 71% (3:1) [Ru] 64% (4:1) [Rh]	3'a , 78% (10:1) [Ru] 59% (13:1) [Rh]	3'b'a , 84% (3:1) [Ru] 59% (5:1) [Rh]

^a Reaction conditions [Ru]: **1** (0.2 mmol), **2a** (0.4 mmol), [Ru(*p*-cymene)Cl₂]₂ (5 mol%), MesCOOH (50 mol%), AgSbF₆ (25 mol%), MW, 25 °C, 2 h; [Rh]: **1** (0.2 mmol), **2a** (0.4 mmol), [Cp*Rh(CH₃CN)₃](SbF₆)₂ (8 mol%), AdCOOH (30 mol%), AgSbF₆ (20 mol%), MW, 80 °C, 2 h. ^b The *E* : *Z* ratio was determined by ¹H NMR analysis. MW = microwave.

was isolated in 40% yield in the presence of [Ru(*p*-cymene)Cl₂]₂ (5 mol%), AgSbF₆ (25 mol%), and AdCOOH (30 mol%) at 25 °C for 1 h (Table 1, entry 1). Other ruthenium salts, such as [Cp*RuCl₂]₂ and RhCl₃·3H₂O, were also investigated, which showed no catalytic reactivity (Table 1, entries 2 and 3). Subsequently, the effects of various additives, including MesCOOH, AcOH, NaOAc, PivONa·H₂O, and DABCO, were screened, which indicates that MesCOOH was the best choice to afford **3aa** in 47% yield (Table 1, entry 4). Increasing the loading of MesCOOH from 30 mol% to 50 mol% resulted in an increased yield (57%) of **3aa** (Table 1, entry 9). To improve reaction efficiency, the temperature range was evaluated (Table 1, entries 10–12). It was found that product **3aa** was obtained in 87% yield with excellent *E*/Z selectivity (Table 1, entry 13). To demonstrate the unique reactivity of Ru catalyst in this protocol, [Cp*Rh(CH₃CN)₃](SbF₆)₂ was also tested to give a slightly lower yield with decreased *E*/Z selectivity at higher temperature (Table 1, entry 14).

Under the optimized reaction conditions with the ruthenium catalyst, the substrate scope of indolines **1** was investigated (Table 2). C2-alkyl and aryl substituted indolines were initially examined to give products **3ba**–**fa** in 65–96% yields. Indoline bearing methyl group at the C3 position was also tolerated to afford product **3ga** in 91% yield. For C4 and C5 substituted

indolines, both electron-donating (Me, OMe, and OBn) and electron-withdrawing (F and Cl) groups were well compatible to provide the corresponding allylated products **3ha**–**la** and **3na**–**ra** in 80–96% yields. The presence of bromo and ester functionalities on substrates led to products **3ma** (54%), **3sa** (73%), and **3ta** (60%) in slightly decreased yields. In addition, disubstituted indolines were also tested to deliver products **3ua** and **3va** in 82% and 89% yields, respectively. Notably, C6-halogen substituted indolines exhibited high activity to afford products **3wa**–**ya** in 78–95% yield. However, C6-methyl substituted substrate turned to be unsuccessful probably due to steric hindrance effect. Moreover, quinoline and carbazole substrates were also employed to afford the allylated products **3za** and **3'a** in 71% and 78% yields. Finally, a series of other directing groups such as acetyl, pyridyl, and others were also tried (see in

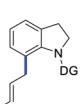
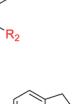
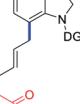


the ESI[†]). While **3b'a** was obtained in 84% yield, others failed to give the coupling products. Overall, indolines with a variety of functionalities ranging from C2 to C6 positions could react with VCPs **2** to afford the allylated products in good yields with high *E/Z* selectivities under the $[\text{Ru}(p\text{-cymene})\text{Cl}_2]_2$ catalytic system. On the other hand, when $[\text{RhCp}^*(\text{CH}_3\text{CN})_3](\text{SbF}_6)_2$ was employed as the catalyst instead of Ru analogue, decreased yields accompanied with lower *E/Z* selectivities was observed in most cases.

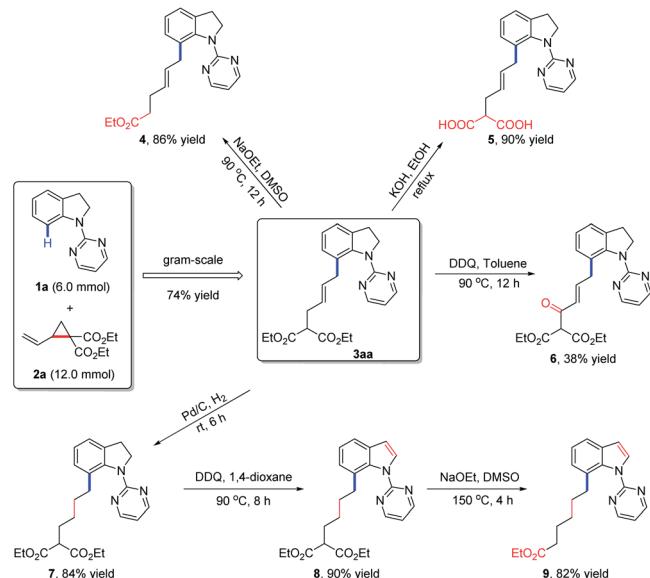
Encouraged by the above results, the scope of VCPs **2** was also explored to further examine the generality of the current C–C/H activation strategy with the ruthenium catalyst (Table 3). Various ester-substituted VCPs, including CO_2Me , $^t\text{BuCO}_2$, $^i\text{PrCO}_2$, and BnCO_2 groups, were successfully coupled with indoline **1a** to afford products **3ab–ad** in 74–90% yields with up to 20 : 1 *E/Z* selectivity. For VCPs having $\text{CF}_3\text{CH}_2\text{CO}_2$ and lactone activating groups, products **3af** and **3ag** were isolated in decreased yields. When CN and PhSO_2 functionalities were introduced into VCP moiety, the corresponding products **3ah–ak** were obtained in 56–78% yields with decreased *E/Z* values. Furthermore, the substituted indolines could also furnish the corresponding products (**3fe**, **3kc**, and **3sc**) in 74–90% yields. Finally, Rh catalytic system could give the corresponding products in comparable yields and *E/Z* ratios.

To evaluate the practical utility of the current methodology, a gram-scale reaction between **1a** (6.0 mmol) and **2a** (12.0 mmol) was performed under standard conditions, which delivered the allylated product **3aa** in 74% yield (Scheme 2).

Table 3 Substrate scope of VCPs^{a,b}

 + 		$[\text{Ru}(p\text{-cymene})\text{Cl}_2]_2$ (5 mol %) AgSbF_6 (25 mol %) MesCOOH (50 mol %) MW, 25 °C, 2 h				
		DG = 2-Pyrimidine				
	3ab , $R_1 = R_2 = \text{CO}_2\text{Me}$	3ae , $R_1 = R_2 = \text{CO}_2\text{Bn}$	3ai , $R_1 = R_2 = \text{SO}_2\text{Ph}$			
	86% (>20:1) [Ru] 78% (11:1) [Rh]	90% (7:1) [Ru] 82% (13:1) [Rh]	58% (8:1) [Ru] 52% (6:1) [Rh] ^d			
	3ac , $R_1 = R_2 = \text{CO}_2\text{Bu}$	3af , $R_1 = R_2 = \text{CO}_2\text{CH}_2\text{CF}_3$	3aj , $R_1 = \text{CN}, R_2 = \text{CO}_2\text{Me}$			
	87% (>20:1) [Ru] 76% (>20:1) [Rh]	76% (6:1) [Ru] 73% (5:1) [Rh]	78% (3:1) [Ru] ^c 65% (2:1) [Rh]			
	3ad , $R_1 = R_2 = \text{CO}_2\text{Pr}$	3ah , $R_1 = R_2 = \text{CN}$	3ak , $R_1 = \text{CN}, R_2 = \text{SO}_2\text{Ph}$			
	74% (>20:1) [Ru] 80% (>20:1) [Rh]	71% (4:1) [Ru] ^c 62% (4:1) [Rh]	56% (4:1) [Ru] ^d 61% (3:1) [Rh] ^d			
	3ag , 65% (>20:1) [Ru] 60% (15:1) [Rh]	3fe , 90% (>20:1) [Ru] 84% (4:1) [Rh]	3kc , 85% (4:1) [Ru] 82% (11:1) [Rh]	3sc , 74% (11:1) [Ru] 81% (>20:1) [Rh]		

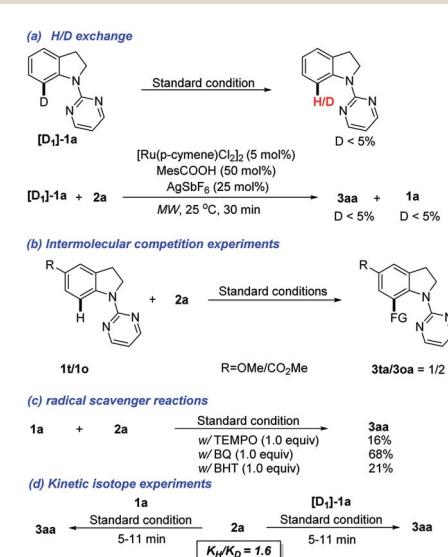
^a Reaction conditions [Ru]: **1** (0.2 mmol), **2a** (0.4 mmol), $[\text{Ru}(p\text{-cymene})\text{Cl}_2]_2$ (5 mol %), MesCOOH (50 mol %), AgSbF_6 (25 mol %), MW, 25 °C, 2 h; [Rh]: **1** (0.2 mmol), **2a** (0.4 mmol), $[\text{Cp}^*\text{Rh}(\text{CH}_3\text{CN})_3](\text{SbF}_6)_2$ (8 mol %), AdCOOH (30 mol %), AgSbF_6 (20 mol %), MW, 80 °C, 2 h. ^b The *E* : *Z* ratio was determined by ¹H NMR analysis. ^c 40 °C. ^d [Ru] CH_2Cl_2 (0.5 mL), 40 °C; [Rh]: MeOH (0.5 mL). MW = microwave.



Scheme 2 Gram-scale reaction and further derivatization of product **3aa**.

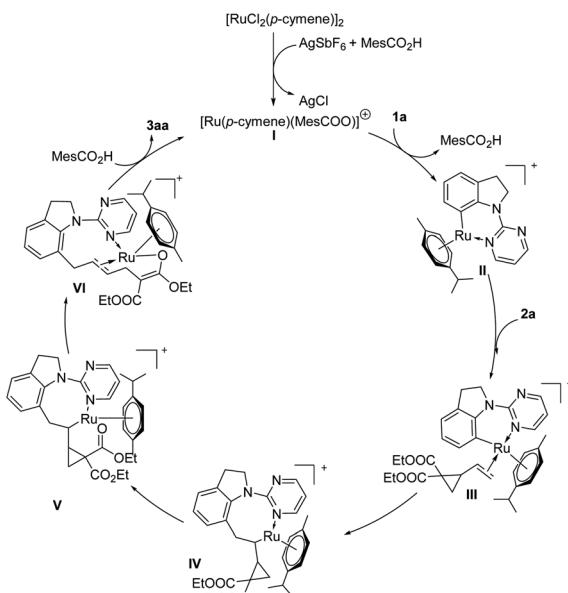
Meanwhile, the derivatizations of **3aa** were also conducted to highlight the synthetic importance of allylated indolines. Firstly, **3aa** could undergo decarboxylation in the presence of NaOEt in DMSO to afford compound **4** in 86% yield. Under the condition of KOH in EtOH , the hydrolysis product **5** was obtained in 90% yield. Secondly, in the presence of DDQ in Toluene , product **3aa** could be oxidized to compound **6** in 38% yield. Thirdly, hydrogenation of **3aa** under Pd-C/H_2 conditions would afford a reduced product **7**, which could further undergo oxidation and decarboxylation transformations to give indole derivatives **8** and **9** in 90% and 82% yields, respectively.

To gain insights into the reaction mechanism, a series of control experiments were conducted (Scheme 3). In the H/D



Scheme 3 Control experiments.





Scheme 4 Proposed reaction mechanism.

exchange experiment, the deuterated **[D]-1a** was prepared and subjected to the standard conditions. It was found that only negligible amount of deuterium was detected for retrieved **1a**. When **2a** reacted with **[D]-1a** for 30 min, a similar result with a distinct D/H exchange was also observed (Scheme 3a). Next, the intermolecular competition experiment between substituted indolines **1o** and **1t** was conducted. Methoxyl-substituted allylated indoline **3oa** was isolated as the major product, indicating that indolines bearing electron-donating groups are more reactive (Scheme 3b). When a radical quencher, such as 2,2,6,6-tetramethyl-1-piperidinyloxy (TEMPO), 2,6-di-*tert*-butyl-4-methylphenol (BHT), or benzoquinone (BQ), was added, the allylation reaction was suppressed with product **3aa** obtained in a decreased yield (Scheme 3c). These results could't indicate that a non-radical mediated reaction pathway. Finally, the KIE values observed in parallel reactions suggest that the C-H bond cleavage is not involved in the limiting step (Scheme 3d).

On the basis of above discussion and related reports,¹⁷ a plausible catalytic cycle was proposed (Scheme 4). A pyrimidine-directed C-H activation between indoline **1a** and an *in situ*-generated Ru cationic complex **I** gives a six-membered ruthenacycle **II**. Coordination of intermediate **II** with VCP **2a** provides intermediate **III**, which undergoes 1,2-migratory insertion to afford an ruthenium intermediate **IV**. Then oxygen coordinates to Ru metal center to form intermediate **V**. After C-C cleavage of VCP **2a**, intermediate **VI** will be generated, followed by protonation to deliver the desired product **3aa** and regenerate the active catalyst species **I**.

Conclusions

In summary, we have successfully demonstrated a $[\text{Ru}(p\text{-cymene})\text{Cl}_2]_2$ -catalyzed C7 allylation of indolines with VCPs as the

allylating reagents *via* a sequential C-H/C-C process. This protocol is applicable to broad substrate scope with good functional group compatibility and high *E/Z* selectivity. The practical applications of allylation transformation are highlighted through gram-scale production and multiple derivatizations. Notably, the use of microwave irradiation under room temperature and solvent-free conditions makes this protocol more valuable and operational convenience, which provides a new insight for the formation of indoline derivatives.

Conflicts of interest

There are no conflicts to declare.

Acknowledgements

Financial support from the National Natural Science Foundation of China (21672192, 21803059, U1904212, and 21929101) is gratefully appreciated.

Notes and references

- (a) B. M. Trost, *Acc. Chem. Res.*, 2002, **35**, 695; (b) P. T. Anastas and N. Eghbali, *Chem. Soc. Rev.*, 2010, **39**, 301; (c) C.-J. Li and P. T. Anastas, *Chem. Soc. Rev.*, 2012, **41**, 1413.
- (a) R. S. Varma, *Green Chem.*, 2014, **16**, 2027; (b) R. Luque and F. L.-Y. Lam, *Sustainable Catalysis Energy-Efficient Reactions and Applications*, Wiley-VCH Verlag GmbH & Co. KGaA, Germany, 2018; (c) W. Zhang and B. W. Cue, *Green Techniques for Organic Synthesis and Medicinal Chemistry*, John Wiley & Sons, Ltd, New York, 2nd edn, 2018; (d) A. Beillard, X. Bantrel, T.-X. Métro, J. Martinez and F. Lamaty, *Chem. Rev.*, 2019, **119**, 7529.
- (a) A. Daştan, A. Kulkarni and B. Török, *Green Chem.*, 2012, **14**, 17; (b) M. B. Gawande, S. N. Shelke, R. Zboril and R. S. Varma, *Acc. Chem. Res.*, 2014, **47**, 1338; (c) A. K. Rath, M. B. Gawande, R. Zboril and R. S. Varma, *Coord. Chem. Rev.*, 2015, **291**, 68; (d) M. Tsuji, *ChemistrySelect*, 2017, **2**, 805; (e) G. B. Dudley and A. E. Stiegman, *Chem. Rec.*, 2018, **18**, 381.
- (a) L. Yang and H. Huang, *Chem. Rev.*, 2015, **115**, 3468; (b) Z. Chen, B. Wang, J. Zhang, W. Yu, Z. Liu and Y. Zhang, *Org. Chem. Front.*, 2015, **2**, 1107; (c) L. Huang, M. Arndt, K. Gooßen, H. Heydt and L. J. Gooßen, *Chem. Rev.*, 2015, **115**, 2596; (d) T. Gensch, M. N. Hopkinson, F. Glorius and J. Wencel-Delord, *Chem. Soc. Rev.*, 2016, **45**, 2900; (e) W. Yu, W. Zhang, Y. Liu, Z. Liu and Y. Zhang, *Org. Chem. Front.*, 2017, **4**, 77; (f) J. He, M. Wasa, K. S. L. Chan, Q. Shao and J.-Q. Yu, *Chem. Rev.*, 2017, **117**, 8754; (g) P. Gandeepan, T. Müller, D. Zell, G. Cera, S. Warratz and L. Ackermann, *Chem. Rev.*, 2019, **119**, 2192; (h) Z. Chen, M.-Y. Rong, J. Nie, X.-F. Zhu, B.-F. Shi and J.-A. Ma, *Chem. Soc. Rev.*, 2019, **48**, 4921.
- (a) F. Chen, T. Wang and N. Jiao, *Chem. Rev.*, 2014, **114**, 8613; (b) L. Souillart and N. Cramer, *Chem. Rev.*, 2015, **115**, 9410; (c) D.-S. Kim, W.-J. Park and C.-H. Jun, *Chem. Rev.*, 2017, **117**,



8977; (d) F. Song, T. Gou, B.-Q. Wang and Z.-J. Shi, *Chem. Soc. Rev.*, 2018, **47**, 7078.

6 S. Laclef, M. Harari, J. Godeau, I. Schmitz-Afonso, L. Bischo, C. Hoarau, V. Levacher, C. Fruit and T. Besson, *Org. Lett.*, 2015, **17**, 1700.

7 (a) F. Wang, S. Yu and X. Li, *Chem. Soc. Rev.*, 2016, **45**, 6462; (b) Z. Nairoukh, M. Cormier and I. Marek, *Nat. Rev. Chem.*, 2017, **1**, 0035.

8 (a) R. A. PerianaRobert and G. Bergman, *J. Am. Chem. Soc.*, 1984, **106**, 7272; (b) R. A. PerianaRobert and G. Bergman, *J. Am. Chem. Soc.*, 1986, **108**, 7346.

9 (a) H. M. Ko and G. Dong, *Nat. Chem.*, 2014, **6**, 739; (b) Y. Xia, G. Lu, P. Liu and G. Dong, *Nature*, 2016, **539**, 546; (c) Y. Xia, J. Wang and G. Dong, *Angew. Chem., Int. Ed.*, 2017, **56**, 2376.

10 (a) A. Masarwa, D. Didier, T. Zabrodski, M. Schinkel, L. Ackermann and I. Marek, *Nature*, 2014, **505**, 199; (b) Y.-F. Liang, V. Müller, W. Liu, A. Münch, D. Stalke and L. Ackermann, *Angew. Chem., Int. Ed.*, 2017, **56**, 9415; (c) H. Wang, I. Choi, T. Rogge, N. Kaplaneris and L. Ackermann, *Nat. Catal.*, 2018, **1**, 993.

11 (a) S. Yu and X. Li, *Org. Lett.*, 2014, **16**, 1220; (b) X. Zhou, S. Yu, L. Kong and X. Li, *ACS Catal.*, 2016, **6**, 647; (c) L. Kong, S. Yu, G. Tang, H. Wang, X. Zhou and X. Li, *Org. Lett.*, 2016, **18**, 3802.

12 (a) T. Seiser, O. A. Roth and N. Cramer, *Angew. Chem., Int. Ed.*, 2009, **48**, 6320; (b) L. Souillart and N. Cramer, *Chem. Sci.*, 2014, **5**, 837; (c) L. Souillart and N. Cramer, *Angew. Chem., Int. Ed.*, 2014, **53**, 9640.

13 (a) Z. Chai and T. J. Rainey, *J. Am. Chem. Soc.*, 2012, **134**, 3615; (b) H. Zhang, K. Wang, B. Wang, H. Yi, F. Hu, C. Li, Y. Zhang and J. Wang, *Angew. Chem., Int. Ed.*, 2014, **53**, 13234; (c) A. Masarwa, M. Weber and R. Sarpong, *J. Am. Chem. Soc.*, 2015, **137**, 6327; (d) M. Li and F. Y. Kwong, *Angew. Chem., Int. Ed.*, 2018, **57**, 6512; (e) J.-L. Pan, C. Liu, C. Chen, T.-Q. Liu, M. Wang, Z. Sun and S.-Y. Zhang, *Org. Lett.*, 2019, **21**, 2823; (f) S. Li, P. Shi, R.-H. Liu, X.-H. Hu and T.-P. Loh, *Org. Lett.*, 2019, **21**, 1602.

14 (a) B. H. Rotstein, S. Zaretsky, V. Rai and A. K. Yudin, *Chem. Rev.*, 2014, **114**, 8323; (b) I. Marek, A. Masarwa, P.-O. Delaye and M. Leibeling, *Angew. Chem., Int. Ed.*, 2015, **54**, 414; (c) G. Fumagalli, S. Stanton and J. F. Bower, *Chem. Rev.*, 2017, **117**, 9404; (d) P.-H. Chen, B. A. Billett, T. Tsukamoto and G. Dong, *ACS Catal.*, 2017, **7**, 1340.

15 (a) J.-Q. Wu, Z.-P. Qiu, S.-S. Zhang, J.-G. Liu, Y.-X. Lao, L.-Q. Gu, Z.-S. Huang, J. Li and H. Wang, *Chem. Commun.*, 2015, **51**, 77; (b) Q. Lu, F. J. R. Klauck and F. Glorius, *Chem. Sci.*, 2017, **8**, 3379; (c) D. Zell, Q. Bu, M. Feldt and L. Ackermann, *Angew. Chem., Int. Ed.*, 2016, **55**, 7408; (d) T. H. Meyer, W. Liu, M. Feldt, A. Wuttke, R. A. Mata and L. Ackermann, *Chem.-Eur. J.*, 2017, **23**, 5443; (e) Z. Hu, X.-Q. Hu, G. Zhang and L. J. Gooßen, *Org. Lett.*, 2019, **21**, 6770.

16 (a) X. Zhu, X.-J. Shen, Z.-Y. Tian, S. Lu, L.-L. Tian, W.-B. Liu, B. Song and X.-Q. Hao, *J. Org. Chem.*, 2017, **82**, 6022; (b) Y.-J. Guo, S. Lu, L.-L. Tian, E.-L. Huang, X.-Q. Hao, X. Zhu, T. Shao and M.-P. Song, *J. Org. Chem.*, 2018, **83**, 338; (c) S. Lu, L.-L. Tian, T.-W. Cui, Y.-S. Zhu, X. Zhu, X.-Q. Hao and M.-P. Song, *J. Org. Chem.*, 2018, **83**, 13991; (d) X.-C. Li, C. Du, H. Zhang, J.-L. Niu and M.-P. Song, *Org. Lett.*, 2019, **21**, 2863; (e) L.-L. Tian, S. Lu, Z.-H. Zhang, E.-L. Huang, H.-T. Yan, X. Zhu, X.-Q. Hao and M.-P. Song, *J. Org. Chem.*, 2019, **84**, 5213; (f) Q. Wang, C.-L. Zhi, P. -P. Lu, S. Liu, X. Zhu, X.-Q. Hao and M.-P. Song, *Adv. Synth. Catal.*, 2019, **361**, 1253; (g) X.-M. Zhao, E.-L. Huang, Y.-S. Zhu, J. Li, B. Song, X. Zhu and X.-Q. Hao, *Org. Biomol. Chem.*, 2019, **17**, 4869; (h) C. Zhi, Q. Wang, S. Liu, Y. Xue, L. Shi, X. Zhu, X.-Q. Hao and M.-P. Song, *J. Org. Chem.*, 2020, **85**, 1022.

17 (a) G. Song, F. Wang and X. Li, *Chem. Soc. Rev.*, 2012, **41**, 3651; (b) J. Li and L. Ackermann, *Angew. Chem., Int. Ed.*, 2015, **54**, 3635.

