Aerobic dehydrogenation of amines to nitriles catalyzed by triazolylidene ruthenium complexes with O₂ as terminal oxidant

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Abstract

Pyridyl-substituted mesoionic triazolylidene ruthenium cymene complexes catalyze the oxidation of both aromatic and aliphatic amines to nitriles with high activity and selectivity under benign conditions using dioxygen as the terminal oxidant. Modification on the pyridyl moiety of the ligand scaffold has negligible effect on the catalytic performance, while substituents on the triazolylidene directly affect the catalytic fitness of the metal center, leading to distinct catalytic profiles. Pre-dissociation of the cymene ligand and formation of a solvento analogue further enhances the catalytic activity towards nitrile formation. Variation of reaction conditions provided valuable mechanistic insights and resulted in a highly efficient protocol for nitrile formation with maximum turnover numbers around 10,000. The turnover frequency reaches up to 400 h^{-1}, providing one of the fastest catalytic systems known to date for this transformation.
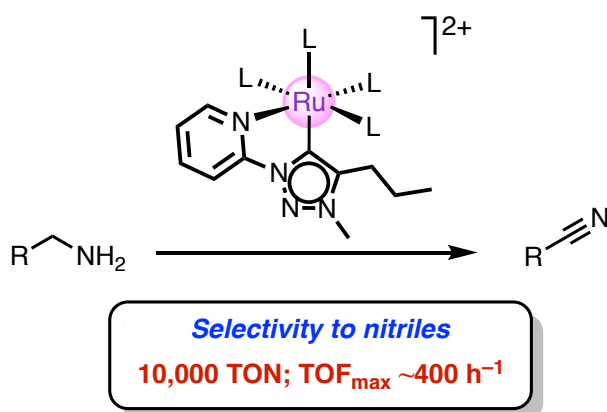
Introduction

Nitriles are a prominent class of organic molecules which serve as versatile intermediates in organic transformations and are included in a wide variety of natural products and biologically active compounds.[1,2] Thus, nitriles are valuable synthetic building blocks in the synthesis of fine chemicals.[3,4] They are prepared by conventional methods such as dehydration of amides/aldoximes,[5,6] through cyanation of alkyl or aryl halides,[7] or by the Sandmeyer and Schmidt-type reactions,[8,9] among many others.[10,11] These traditional synthetic routes to prepare nitriles proceed with low atom economy, produce stoichiometric waste, require toxic reagents like HCN, have limited selectivity and often require harsh reaction conditions.[12] In recent years, a substantial amount of research has been directed towards the development of direct oxidative dehydrogenation (ODH) of primary amines to nitriles by molecular oxygen in the presence of transition metal catalysts, a pathway that starts from abundant low-value precursors and avoids toxic reagents and harsh reaction conditions.[13,14] However, the ODH of primary amines to nitriles is a challenging task, as it requires the removal of four electrons and four protons and generally involves several competing pathways, such as dehydrogenation, coupling and transamination.[15] Amine dehydrogenation can hence lead to a broad range of products such as oximes, imines, amides, nitriles, amine oxides and azo compounds. For this reason, the design and development of efficient and selective catalytic systems is of great importance. McWhinnie *et al.* first reported the conversion of amine ligands coordinated to a ruthenium(II) center to nitriles upon exposure to oxygen under ambient conditions.[16,17] Later, Taube and co-workers studied the oxidation of different types of amine ligands coordinated to ruthenium centers.[18,19] Pioneering studies of Tang *et al.* reported the first catalytic reaction for the oxidative dehydrogenation of amines with high activity but poor selectivity when using 2–3 atm of oxygen at 100 ºC.[20] This study triggered the development of numerous ruthenium-based homogeneous catalytic systems for the aerobic oxidation of amines.[21–24] For example, James *et al.* reported the synthesis of a ruthenium-porphyrin complex that exhibited 100% selectivity under mild conditions (50 ºC, <16 h),[21] Later on Parvulescu introduced a faster novel ruthenium-terpyridyl complex that showed conversion of amines to nitriles after 2 h when using 5 atm of oxygen at 60 ºC,[25] while Maiti and co-workers developed new ruthenium hydrido complexes for the selective generation of nitriles and imines by varying the catalyst instead of the substrates.[26] More recently, different N-chelated ruthenium complexes[27–29] as well as simple [Ru(arene)Cl$_2$]$_2$ complexes have been demonstrated to be highly competent in amine dehydrogenation.[30,31]

Herein we describe new ruthenium(II) complexes featuring a range of functionalized triazolylidene ligands and their catalytic activity in amine oxidation. This subclass of N-heterocyclic carbene (NHC) ligands[32,33] is particularly attractive due to the strong σ-donor capability and easy synthetic accessibility through versatile "click chemistry".[34–38] Mesoionic NHCs can act as electron reservoirs, displaying a non-innocent behavior when coordinated to
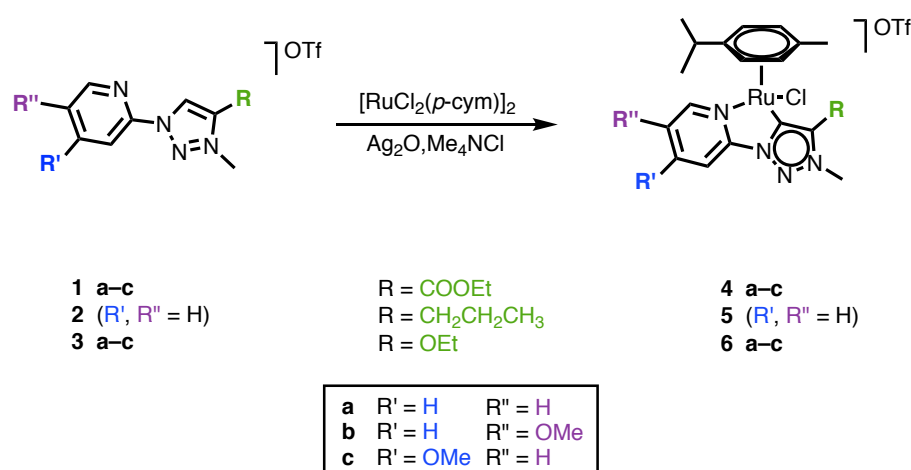
metal centers. Specifically in this work, we have included pendant ester, ether and aliphatic groups adjacent to the carbene carbon, as this creates opportunities to electronically modulate the catalytic properties of the metal center. Further modifications include the pyridyl moiety by incorporation of a methoxy group in different positions, as well as the ancillary ligands through replacing the *p*-cymene with MeCN ligands and formation of the corresponding solvento complex. Interestingly, these mesoionic triazolylidene ruthenium complexes display higher catalytic activity than other catalytic systems reported so far for the oxygen-mediated conversion of amines into the corresponding nitriles.

Result and discussion

### 1. Synthesis and characterization of a series of pyridyl-triazolylidene ruthenium complexes.

The differently functionalized triazolium salts **1**–**3** were prepared *via* a previously reported procedure.[39] Metalation of **1**–**3** and formation of the corresponding triazolylidene ruthenium(II) complexes **4**–**6** was achieved *via* a silver triazolylidene intermediate and *in situ* carbene transfer to [RuCl$_2$(*p*-cymene)]$_2$ in the presence of Me$_4$NCl (Scheme 1). The chloride salt serves both as a means to remove silver upon transmetalation and to prevent scrambling of the anionic ancillary ligand in the ruthenium complex (Cl$^-$ *vs*. OTf$^-$). Purification by column chromatography yielded complexes **4**–**6** as orange air-stable solids in 20–60% yield.

Scheme 1. Synthesis of 1,2,3-triazolylidene ruthenium(II) complexes **4**–**6**.

1 a–c	R = COOEt
2 (R', R'' = H)	R = CH$_2$CH$_2$CH$_3$
3 a–c	R = OEt

| 4 a–c |
| 5 (R', R'' = H) |
| 6 a–c |

a	R' = H	R'' = H
b	R' = H	R'' = OMe
c	R' = OMe	R'' = H

NMR characterization. All complexes were fully characterized by ^{1}H and ^{13}C NMR spectroscopy. Triazolylidene coordination to the ruthenium metal center was indicated by the absence of the low-field proton resonance of the triazolium salt and the appearance of the carbene carbon resonance in the ^{13}C NMR spectrum at around $\delta_C = 175$ ppm. This frequency is in agreement with previously reported chelating triazolylidene ruthenium complexes.[40,41] The absence of the triazolylidene proton as well as the splitting of the aromatic *p*-cymene protons into four distinct doublets indicate chelation and therefore *N,C*-bidentate coordination of the pyridyl-triazolylidene ligand in complexes **4**–**6**. Furthermore, the pyridyl α proton is considerably deshielded, *e.g.*, a doublet at $\delta_H = 8.81$ for complex **6b**, which is characteristic[41,42,43] for N-coordination of the pyridyl unit to the ruthenium center (*cf.* $\delta_H = 8.22$ ppm in the ligand precursor **3b**).

Structural characterization in the solid state. Unambiguous evidence for the proposed structures was obtained from single crystal X-ray diffraction analyses of complexes **4a**–**c** and **6a**. The molecular structures show the typical three-legged piano-stool arrangement around ruthenium(II) with the *C,N*-bidentate chelating triazolylidene pyridyl ligand occupying two of the three base positions (Fig. 1). The five-membered metallacycle features a typically acute C_{trz}–Ru–N_{py} bite angle of $77(\pm 1)°$ (Table 1).[41,42] The Ru–C_{trz} bond length is 2.04(2) Å, and is thus in the expected range when compared to related triazolylidene complexes.[44,45] All four complexes show similar bond distances suggesting that variation of the ligand substitution pattern has only minor influences on the ligand bonding.


4a

4b

4c

6a
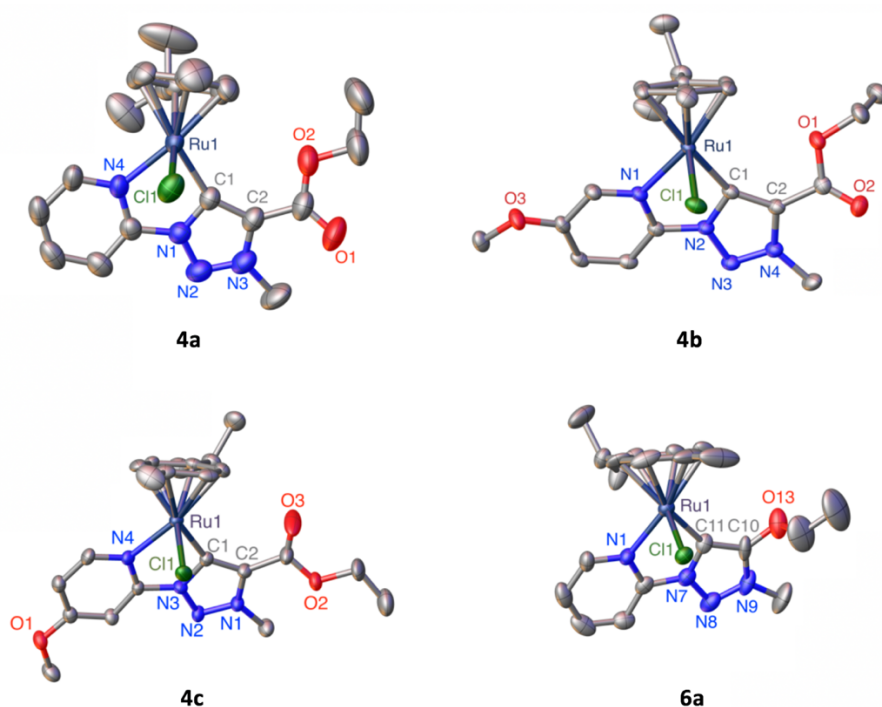
Figure 1. ORTEP presentation of complexes **4a–c** and **6a** (50% probability ellipsoids, non-coordinating OTf⁻ anions and co-crystallized solvent molecules omitted for clarity).

Table 1. Selected bond lengths (Å) and angles (°) for complexes **4a–c** and **6**.

	4a[a]	**4b**	**4c**[b]	**6a**[b]
Ru–N_{py}	2.0983(18)	2.1024(15)	2.108(4)	2.113(7)
Ru–C_{trz}	2.049(2)	2.0570(19)	2.034(6)	2.032(8)
Ru–Cg_{cym}	1.706(10)	1.705(8)	1.707(2)	1.692(3)
Ru–Cl	2.4004(6)	2.4091(5)	2.4022(13)	2.395(2)
C_{trz}–C_{trz}	1.399(3)	1.392(3)	1.399(8)	1.387(12)
C_{trz}–Ru–N_{py}	77.92(8)	77.60(7)	77.7(2)	76.4(3)

[a] data from ref. [41]; [b] unit cell contains two independent complex molecules; bond lengths (Å) and angles (°) only given for one of these molecules as the second features identical bond lengths and angles within esd's.

2. Catalytic oxidation of amines to nitriles. The catalytic activity of the novel ruthenium complexes **4**–**6** was evaluated in amine oxidation using 4-methylbenzylamine as model substrate and 1,2-dichlorobenzene as solvent. An initial run with complex **4b** as catalyst precursor at 5 mol% catalyst loading produced the corresponding nitrile as the major product in a high 86% yield after 3 h. The catalytic profile of the reaction reveals rapid conversion of the substrate that is almost complete within the first hour of reaction (94% conversion) to yield a mixture of benzonitrile (29%) and the corresponding imine (49%; Fig. 2). Nitriles are formed by a double-dehydrogenation of amines, while the secondary aldimine is a product from mono-dehydrogenation followed by condensation with a substrate amine. Although imine formation is favored over nitrile production in the first hour, the imine product is intermittent and disappeared over time. Consumption of the imine was accompanied with an increase of nitrile formation and the appearance of traces of tolylaldehyde, suggesting that the imine is formed as a transient species. After 3 h, all imine was consumed, and the corresponding nitrile was the main product together with low quantities (<15%) of aldehyde as a side-product. A blank reaction under identical conditions yet in the absence of complex **4b** led to slow formation of small quantities of imine (15% after 22 h), yet no nitrile nor aldehyde was observed, indicating a pivotal role of the triazolylidene ruthenium complex as catalyst for amine oxidation.
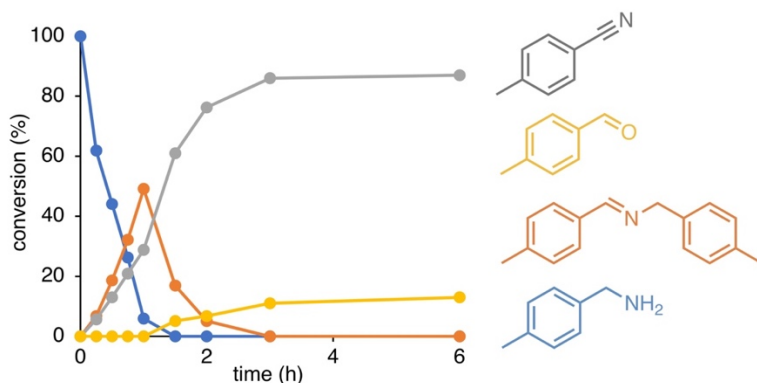
Figure 2. Time-dependent conversion profile for the catalytic oxidation of 4-methylbenzylamine with complex **4b**. General conditions: 4-methylbenzylamine (0.2 mmol), [Ru] (0.01 mmol, 5 mol%), 1,2-dichlorobenzene (2 mL), 150 ºC. Conversions were determined by ^{1}H NMR integration (1,3,5-trimethoxybenzene as internal standard) and are averaged over 2 runs.

To better understand the catalytic profile and optimize the reactions conditions, a series of experiments were monitored under strict control of the atmosphere (Table S1). The first set of experiments were directed towards enhancing the selectivity of the oxidation towards either nitrile or imine formation. Therefore, the reaction was run under inert atmosphere using dry solvent either in an open system with a condenser and argon atmosphere (1 bar) or in a closed Schlenk flask with a small headspace. Both experiments barely gave any products. However, the reaction proceeded well when performed under an atmosphere of air, both in an open and closed system. Under these conditions, a mixture of nitrile and imine formed within 1 h in about 30% and 50% yield, respectively, suggesting H_2O rather than H_2 as the potential by-product of the dehydrogenation of the amine to the aldimine. Moreover, the fact that both imine and nitrile are formed in similar ratios irrespective of whether the reactor is an open or closed system strongly suggests that the by-product does not inhibit the catalytic reaction. Further support for this conclusion was provided by the insensitivity of the reaction to H_2 atmosphere or moist solvent. Thus, when a reaction was first performed for 1 h in air followed by saturation with H_2, no hydrogenation of the imine or the nitrile back to the benzylamine was observed even after 1 h under H_2 atmosphere. Hence, the ruthenium catalyst derived from complex **4b** catalyzes the dehydrogenation of amines and imines, but not the revers, *i.e.* the hydrogenation of imines or nitriles. Continuous saturation of the reaction mixture with air did slow down the dehydrogenation, however the activity is substantially enhanced when the reaction was run under an atmosphere of molecular oxygen. These conditions reduced the time to completion from 3 to 1 h, and they also markedly enhanced the product selectivity to afford the nitrile in high yield (89%), with only 6% imine formed (15:1 nitrile/imine selectivity, *cf.* 1:1.7 selectivity in air after 1 h).

Attempts to detect the formation of water during the reaction by NMR spectroscopy were not conclusive. However, monitoring the gas phase of the catalytic reaction with anhydrous cobalt(II) chloride, as an indicator showed the characteristic color change from blue to purple only when the Ru complex was present, thus confirming formation of water as a by-product. These experiments clearly indicate that oxygen is essential for this dehydrogenation acting as the terminal oxidant. While an inert atmosphere impedes the reaction, air leads to full conversion of the amine in 1 h to an imine/nitrile mixture, and an atmosphere of oxygen further accelerates the dehydrogenation and produces the nitrile in high yield and selectivity within the same time.

The formation of high ratios of nitrile from intermittent imine/nitrile ratios implies that N-substituted aldimines are converted to nitriles. This transformation is suggested to involve a presumably non-catalyzed equilibrium between the primary and secondary aldimine **III**, with the former prone to undergo a second dehydrogenation to generate the nitrile product **II** (Scheme 2). We note that the aldehyde **IV** is formed in much smaller quantities than the nitrile once all amine is consumed (*cf.* Fig. 2), which excludes quantitative imine transformation to the aldehyde. Formation of aldehyde is surmised to be a side reaction from competition of water with ammonia in the reaction with the N-substituted imine.

Scheme 2. Benzylamine dehydrogenation using oxygen as terminal oxidant by ruthenium(II) complex bearing pyridyl-triazolylidene ligand.
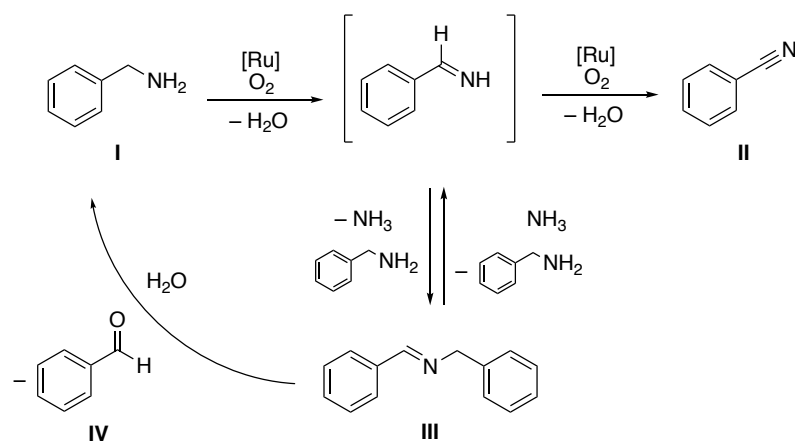
In order to evaluate the role of the pyridyl-triazolylidene ligand, the activity of complexes **4–6** with different ligand substitution patterns were compared in amine oxidation catalysis under aerobic reaction conditions. Complexes **4a–c** with different substituents on the pyridyl moiety feature an essentially identical time-conversion profile with no significant difference in activity nor selectivity (Fig. S7, entries 1–3, Table 2). This outcome suggests that the pyridyl site does not have a direct impact on the catalytic activity. However, complexes **5** and **6a** bearing a propyl

and an ethoxy group on the triazolylidene unit, respectively, were considerably more active at early stage. Conversions reached 90% already after 30 min and were quantitative within 1 h compared to about 50% at 30 min with complexes **4a**–**c** (entries 4,5). Higher activity is also quantified by the improved turnover frequency at 50% conversion, TOF_{50}, which reaches 33 h^{-1} with complex **6a** containing an OEt substituent at the triazolylidene unit (*cf.* $TOF_{50} = 20$ h^{-1} for **4a**). Moreover, only trace amounts of aldehyde were detectable when using complexes **5** and **6a** as catalyst precursors. The higher activity of complexes **5** and **6a** indicates that the electron-donating properties of the triazolylidene are relevant for imparting high catalytic activity. Moreover, complexes **5** and **6a** showed formation of nitrile and imine in an equal ratio within the first 30 min of the reaction, whereas complexes **4a**–**c** formed preferentially the imine over the nitrile at early stage (Table 2). Although the conversion of the substrate took less time and the amount of imine formed was lower when using complex **5** and **6a** compared to complexes **4a**–**c**, it is worth noting that the transformation of the initially formed imine intermediate to benzonitrile required about the same time for all complexes **4**–**6**, suggesting a rate-limiting step for this transformation that is not metal-catalyzed. We assume that this second transformation is rate-limited by the NH_3 concentration, which is required to generate the primary aldimine that is then dehydrogenated to nitrile (*cf.* Scheme 2).

Table 2. Conversions and yields at early and late stage of catalytic 4-methylbenzylamine oxidation using complexes **4a**–**c**, **5** and **6a**.[a]

entry	complex	conversion /%		product yield /% [b]		TOF_{50} (h^{-1})
		1 h	3h	**III** / **II** at 1 h	**III** / **II** / **IV** at 3 h	
1	**4a**	90	>99%	48/28	0/85/13	20
2	**4b**	94	>99%	49/29	0/86/11	17
3	**4c**	88	>99%	54/29	0/86/13	17
4	**5**	98	>99%	21/67	1/86/6	24
5	**6a**	99	>99%	23/62	3/85/10	33

[a] general conditions: 4-methylbenzylamine (0.2 mmol), [Ru] (0.01 mmol, 5 mol%), 1,2-dichlorobenzene (2 mL), 150 ºC, under aerobic reaction conditions (*cf.* entry 3, Table S1); [b] **II** (nitrile), **III** (imine), **IV** (aldehyde), see Scheme 2.

Overall, the activities for complexes **5** and **6a** vary only subtly and therefore, complex **5** was selected for further optimization since its synthesis is more facile than **6a**. Under molecular oxygen, complex **5** reaches full conversion already after 20 min. At this stage, the nitrile/imine ratio is almost 3:1 (67% nitrile, 24% imine; Fig. S8), considerably higher than the 1:2 ratio

observed for **4b** at the same stage in air. After 2 h, the reaction is essentially complete, with 96% yield of nitrile together with 4% of aldehyde as side-product.

Since conversion of the secondary imine to the nitrile is independent of the ruthenium complex, yet limits the rate of nitrile formation once all amine is converted, we reasoned that the equilibrium between the primary and secondary imine (*cf.* Scheme 2) will be favorably shifted to the former if ammonia is supplied. Indeed, a catalytic run performed under oxygen in the presence of gaseous ammonia (6 mL) accelerated nitrile formation considerably. While amine conversion is not affected (20 min, Fig. S9), the fraction of nitrile increased to 81% (*cf.* 67% in the absence of ammonia) and the concentration of imine was considerably reduced from 24% to 12%. After 40 min, the reaction was complete and yielded essentially quantitatively the nitrile product (<1% aldehyde detected). These optimized conditions provided an estimated $TOF_{50} = 75$ h^{-1}, about a fourfold increase compared to the initial experiments under air and in the absence of added ammonia. Moreover, these conditions also successfully suppress the formation of aldehyde as a side-product due to the kinetically favorable reaction of the secondary imine with NH_3 *vs.* H_2O, which allows for further dehydrogenation and nitrile formation. This experiment supports the mechanistic proposal and in particular the uncatalyzed re-activation of the secondary imine for nitrile formation. Controlling this equilibrium with the primary imine is critical to accomplish selective oxidation to the nitrile. In the absence of ammonia, competitive reaction of the imine with water, presumably formed as a by-product from aerobic H_2 fixation, produces the aldehyde and compromises the selectivity. The shorter reaction time also indicates that while the first dehydrogenation is fast (quantitative conversion of the amine), the second dehydrogenation is impeded by competitive imine formation, *i.e.* the rate of dehydrogenation to form the nitrile is slower than the rate of condensation of the primary imine with amine to form the secondary imine **III**. This rate difference depletes the concentration of primary imine if no ammonia is present, while excess ammonia is reversing the formation of the secondary imine and eliminates the otherwise rate-limiting and uncatalyzed aminolysis of the secondary imine for nitrile production. From a process point of view, ammonia addition is unproblematic as it is removed easily due to its low boiling point.

Different primary amine substrates were oxidized by complex **5** under these optimized conditions in an atmosphere of oxygen and ammonia. Both *para*-substituted benzylamines with electron-donating MeO and electron-withdrawing CF_3 groups were converted. Reaction rates are highly similar (Fig. 3), suggesting that the electronically sensitive benzylic C–H bond activation is not part of the turnover-limiting step. In contrast, aliphatic amines were oxidized significantly faster and time to nitrile formation was reduced to 20 min (*cf.* 40 min for benzylamines).
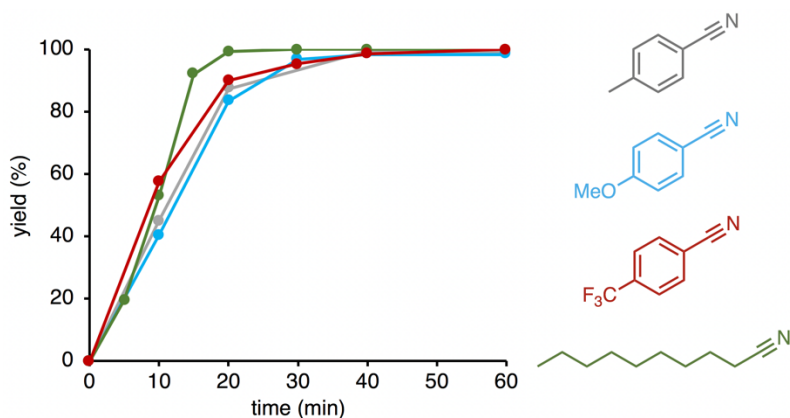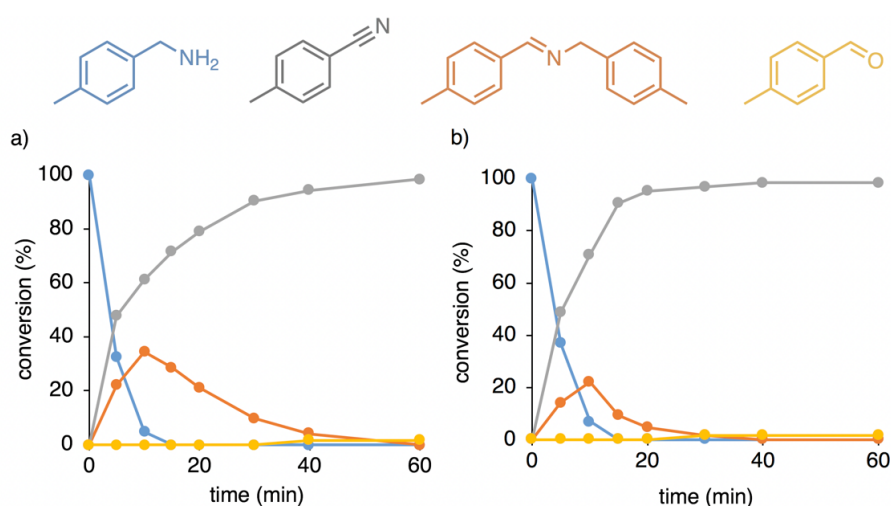
Figure 3. Time-dependent profiles for the catalytic oxidation towards nitrile with complex **5** under ammonia gas (6 mL) and molecular oxygen. General conditions: amine substrate (0.2 mmol), [Ru] (0.01 mmol, 5 mol%), 1,2-dichlorobenzene (2 mL), 150 ºC. Conversions were determined by ^{1}H NMR integration (1,3,5-trimethoxybenzene as internal standard) and are averaged over 2 runs. Isolated yields between 52% (4-Me-benzonitrile) and 86% (4-MeO-benzonitrile).

Mechanistic insights into the nature of the active species were obtained from ^{1}H NMR spectroscopic analysis of the reaction mixture, which revealed the complete disappearance of the resonances due to the ruthenium-bound cymene within 2 h when the reaction was heated at 150 ºC. At room temperature, the cymene ligand was stably bound to the ruthenium center over extended periods. These observations suggest a thermally induced activation of the ruthenium complexes **4**–**6** by cymene dissociation. In order to eliminate this potentially temperature-limiting step, the solvento complex **7** was prepared from the parent cymene analogue **5** upon reaction with AgOTf in refluxing MeCN (Scheme 3) *via* established procedures.[41,46,47,48,49,50] In contrast to complex **5**, the solvento complex **7** is moderately air-sensitive and gradually degrades over several days, as indicated by a color change from yellow to green. Successful ligand exchange was indicated by ^{1}H NMR spectroscopy, which showed the absence of the cymene resonances and the appearance of two singlets attributed to coordinated MeCN in 1:2 ratio (δ_H = 2.54 and 2.07, respectively, CD₃CN solution). The observation of only three MeCN ligands is in agreement with fast exchange of the MeCN ligand *trans* to the carbene with CD₃CN.[41,46] Unequivocal evidence for the ligand exchange and the octahedral geometry around the ruthenium center of complex **7** was obtained by X-ray diffraction studies.

Scheme 3. Synthesis of the solvento ruthenium complex **7** from complex **5** and ORTEP plot (50% probability ellipsoids).
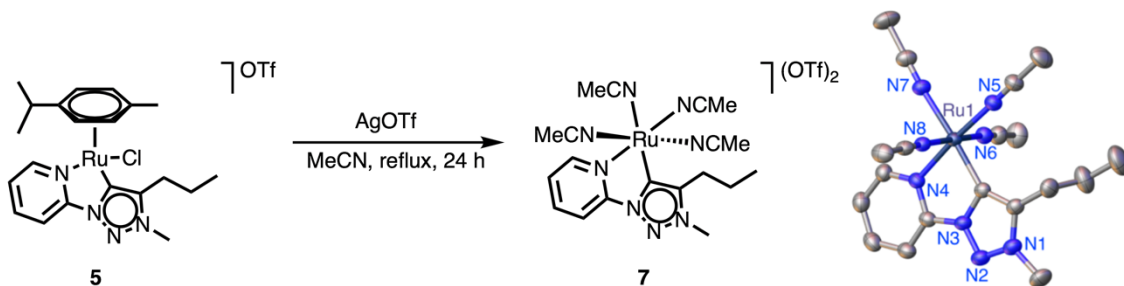
Catalytic amine oxidation using the solvento complex **7** as catalyst precursor was indeed considerably accelerated when compared to reactions with complex **5** under identical conditions, reaching full amine conversion already after 10 min (*cf.* 57% conversion with complex **5** after this time, Fig. 4a). After these initial stages, complex **5** and **7** show similar catalytic profile with formation of nitrile in 81% and 79% yield respectively, suggesting the evolution of an identical catalytic species, which supports the initial loss of cymene from complex **5**. However, the solvento complex **7** showed only very modest activity at lower temperature and reached a mere 50% conversion after 3 d at 80 ºC, indicating that unlike N-coordinated systems, [29,51] high temperatures are needed also for the catalytic dehydrogenation and not only for cymene dissociation during catalyst activation. We note that nitrile formation was more favored when running the catalysis with complex **7** under higher concentration of ammonia gas (30 instead of 6 mL), reaching 71% nitrile and 22% imine after 10 min, and almost full conversion to nitrile (95% yield) after 20 min reaction (Fig. 4b).

Figure 4. Time-dependent profile for the catalytic 4-methylbenzylamine oxidation with complex **7** under molecular oxygen and a) 6 mL ammonia gas and b) 30 mL ammonia gas conditions. General conditions: amine substrate (0.2 mmol), [Ru] (0.01 mmol, 5 mol%), 1,2-dichlorobenzene (2 mL), 150 ºC. Conversions

were determined by ^{1}H NMR integration (1,3,5-trimethoxybenzene as internal standard) and are averaged over 2 runs.

The solvento complex **7** is also competently catalyzing the dehydrogenation of aliphatic amines, such as decylamine (Fig. S10). After 10 min reaction, 93% of the substrate was converted and nitrile formation reached 99% after 15 min, displaying a TOF_{max} of about 150 h^{-1}. Due to the high activity of complex **7**, formation of the imine intermediate or aldehyde as a side-product were not detected by NMR spectroscopy, indicating an excellent selectivity with this catalyst under the applied reaction conditions.

Further studies using complex **7** were carried out to evaluate its catalytic activity at lower catalyst loading. Thus, 4-methylbenzylamine was fully converted at 0.1 mol% catalyst loading with a maximum turnover frequency $TOF_{max} = 250$ h^{-1}. Decreasing the catalyst loading to 0.01 mol% increased the time required for full conversion to 6 days, (TON = 10,000), indicating a highly robust catalytic cycle and little catalyst degradation. Under these conditions TOF_{max} reach a remarkable 400 h^{-1}. The catalytic activity is comparable to that reported for simpler systems like [Ru(cym)Cl$_2$]$_2$. [30,31] Control experiments indicate that under identical conditions (*i.e* 150 ºC in 1,2-dichlorobenzene), the activity of this simple complex is indeed comparable to that of complex **7**, despite the different ligand set of the two ruthenium species.

Conclusion

A new series of 1,2,3-triazolylidene ruthenium(II) complexes were prepared. Simple modifications on the ligand through electronic modulation by introducing electron-donating methoxy substituents on the pyridyl ring (**4a–c**), or by altering the electronic properties on the triazolylidene heterocycle through the incorporation of different substituents on the C4-position (**4a**, **5**, **6a–c**) were evaluated in the catalytic oxidation of 4-methylbenzylamine. While the modifications on the pyridyl moiety series have no significant effects and activities are very similar, the electronic modulation on the triazolylidene however, significantly modulate the catalytic performance. Complexes bearing electron-withdrawing substituent (–COOEt) showed slower conversion of the substrate than those complexes containing electron-donating groups (–OEt or –*n*Pr). These results suggest that the electron donating character on the triazolylidene heterocycle influences the ruthenium metal center in amine oxidation reactions.

Overall, all ruthenium(II) complexes were active catalyst precursors for the oxidation of amine under aerobic conditions, achieving very good yields of nitrile product without the need of any additive (86% after 3 h). In the presence of ammonia and molecular oxygen, the catalytic activity is boosted towards formation of nitrile in quantitative yield and with exclusive selectivity towards the nitrile (99% after 40 min). Further optimization of the catalyst by removing the *p*-cymene

ligand to give the acetonitrile solvento complex yielded a high activity catalyst which induces fast and selective amine oxidation to the corresponding nitrile (full conversion within 15 min, TOF_{max} ~400 h^{-1}) and tolerating catalyst loadings as low as 0.01 mol% (10,000 TON), thus constituting one of the most efficient catalyst for the direct oxidation of amines to nitriles to date.

Experimental section

General. The metalation reactions were carried out under nitrogen atmosphere using standard Schlenk techniques, and all the reagents and solvents were used as obtained from commercial suppliers. Triazolium salts **1a**,[41] **1b**,[41] **1c**,[39] **3b**,[39] and ruthenium complex **4a**[41] were prepared according to previously reported procedures. Unless specified, NMR spectra were recorded at 25 °C on Bruker spectrometers operating at 400 MHz (^{1}H NMR) and 100 MHz (^{13}C NMR), respectively. Chemical shifts (δ in ppm, coupling constants J in Hz) were referenced to residual solvent signals (^{1}H, ^{13}C). Assignments are based on homo- and heteronuclear shift correlation spectroscopy. All complexes show a quartet around 120 ppm in the ^{13}C NMR spectrum due to the OTf counterion. Purity of the complexes has been established by NMR spectroscopy, and by elemental analysis, which were performed by the University of Bern Microanalytic Laboratory using a Thermo Scientific Flash 2000 CHNS-O elemental analyzer. High-resolution mass spectrometry was carried out with a Thermo Scientific LTQ Orbitrap XL (ESI-TOF).

Compound 2

a) Synthesis of 2-(4-propyl-1H-1,2,3-triazol-1-yl)pyridine. A solution of 2-azidopyridine (200 mg, 1.66 mmol), 1-pentyne (197 μL, 1.99 mmol), $CuSO_4 \cdot 5\ H_2O$ (83 mg, 0.33 mmol) and Cu powder (74 mg, 1.16 mmol) in tBuOH:H_2O (9:9 mL) were placed into a sealed microwave vial. The reaction was irradiated at 100 ºC for 15 h. The mixture reaction was diluted with 100 mL of CH_2Cl_2 and extracted with NH_4OH 10% sol. (2 x 100 mL), H_2O (2 x 100 mL) and brine (2 x 100 mL). The organic layer was dried over $MgSO_4$, filtered and all volatiles were evaporated under reduced pressure. The residue was washed with pentane to afford compound **1** as a white solid (285 mg, 91%). ^{1}H NMR (400 MHz, $CDCl_3$): δ = 8.44–8.37 (m, 1H, $C_{py}H$), 8.26 (s, 1H, $C_{trz}H$), 8.11 (d, $^3J_{HH}$ = 8.2 Hz, 1H, $C_{py}H$), 7.87–7.78 (m, 1H, $C_{py}H$), 7.25 (ddd, $^3J_{HH}$ = 7.4 Hz, $^3J_{HH}$ = 4.9 Hz, $^4J_{HH}$ = 1.0 Hz, 1H, $C_{py}H$), 2.73 (t, $^3J_{HH}$ = 7.4 Hz, 2H, $C_{trz}CH_2$), 1.71 (h, 2H, $^3J_{HH}$ = 7.4 Hz, $C_{trz}CH_2CH_2$), 0.95 (t, $^3J_{HH}$ = 7.4 Hz, 3H, CH_2CH_3). ^{13}C{^{1}H} NMR (100 MHz, $CDCl_3$): δ = 149.4 (C_{py}–N_{trz}), 148.7 ($C_{trz}CH_2$), 148.4 ($C_{py}H$), 139.0 ($C_{py}H$), 123.2 ($C_{py}H$), 118.1 ($C_{trz}H$), 113.7 ($C_{py}H$), 27.7 ($CH_2CH_2CH_3$), 22.5 ($CH_2CH_2CH_3$), 13.8 (CH_2CH_3). HR-MS (CH_3CN): m/z calculated for $C_{10}H_{13}N_4$ $[M+H]^+$ = 189.1135; found, 189.1131.

b) Synthesis of triazolium salt 2. To a solution of the 2-(4-propyl-triazol-1-yl)pyridine (420 mg, 2.2 mmol) in CH_2Cl_2 at 0 ºC was added MeOTf (270 μL, 2.45 mmol) and the mixture was stirred at 0 ºC for 30 min. Addition of Et_2O gave a white solid (740 mg, 94%). ^{1}H NMR (400 MHz,

CDCl$_3$): δ = 8.85 (s, 1H, C$_{trz}$H), 8.61 (d, $^4J_{HH}$ = 3.8 Hz, 1H, C$_{py}$H), 8.16 (d, $^3J_{HH}$ = 8.1 Hz, 1H, C$_{py}$H), 8.11–8.05 (m, 1H, C$_{py}$H), 7.60 (dd, $^3J_{HH}$ = 7.3 Hz, $^4J_{HH}$ = 4.9 Hz, 1H, C$_{py}$H), 4.43 (s, 3H, NCH$_3$), 2.99 (t, $^3J_{HH}$ = 7.7 Hz, 2H, C$_{trz}$CH$_2$), 1.89 (m, 2H, C$_{trz}$CH$_2$CH_2), 1.13 (t, $^3J_{HH}$ = 7.3 Hz, 3H, CH$_2$CH_3). ^{13}C{^{1}H} NMR (100 MHz, CDCl$_3$): δ = 149.4 (C$_{py}$H), 146.7 (C$_{py}$–N$_{trz}$), 146.2 (C$_{trz}$CH$_2$), 140.6 (C$_{py}$H), 127.1 (C$_{py}$H), 124.2 (C$_{trz}$H), 115.4 (C$_{py}$H), 39.9 (NCH$_3$), 25.7 (CH$_2$CH$_2$CH$_3$), 20.5 (CH$_2$CH$_2$CH$_3$), 13.7 (CH$_2$CH$_3$). Anal. Calcd for C$_{12}$H$_{15}$F$_3$N$_4$O$_3$ (352.33): C, 40.91; H, 4.29; N, 15.90. Found: C, 41.08; H, 4.18; N, 15.77. HR-MS (CH$_3$CN): m/z calculated for C$_{11}$H$_{15}$N$_4$ [M–OTf]$^+$ = 203.1291; found, 203.1285

Compound 3a

a) Synthesis of 2-(4-ethoxy-1H-1,2,3-triazol-1-yl)pyridine. The reaction of 2-azidopyridine (500 mg, 4.16 mmol), ethoxyacetylene (2.9 mL, 12.48 mmol), CuI (795 mg, 4.16 mmol) and 2,6-lutidine (970 µL, 8.32 mmol) in THF (27 mL) and DMSO (470 µL) was stirred under reflux for 24 h and afforded this compound as an off-white solid (608 mg, 77%). ^{1}H NMR (300 MHz, CDCl$_3$): δ = 8.47 (ddd, $^3J_{HH}$ = 4.9 Hz, $^4J_{HH}$ = 1.9 Hz, $^5J_{HH}$ = 0.9 Hz, 1H, C$_{py}$H), 7.99 (s, 1H, C$_{trz}$H), 7.89 (ddd, $^3J_{HH}$ = 8.3 Hz, $^3J_{HH}$ = 7.4 Hz, $^4J_{HH}$ = 1.9 Hz, 1H, C$_{py}$H), 7.53–7.46 (m, 1H, C$_{py}$H), 7.32 (ddd, $^3J_{HH}$ = 7.4 Hz, $^4J_{HH}$ = 4.9 Hz, $^5J_{HH}$ = 0.9 Hz, 1H, C$_{py}$H), 4.33 (q, $^3J_{HH}$ = 7.1 Hz, 2H, OCH$_2$Me), 1.46 (t, $^3J_{HH}$ = 7.1 Hz, 3H, OCH$_2$CH_3).

b) Synthesis of triazolium salt 3a. To a solution of the 2-(4-ethoxy-triazol-1-yl)pyridine (607 mg, 3.2 mmol) in CH$_2$Cl$_2$ at 0 ºC was added MeOTf (400 µL, 3.67 mmol) and the mixture was stirred at 0 ºC for 30 min. Addition of Et$_2$O gave a white precipitate (245 mg, 69%). ^{1}H NMR (400 MHz, CDCl$_3$): δ = 8.78 (s, 1H, C$_{trz}$H), 8.63–8.60 (m, 1H, C$_{py}$H), 8.15 (d, $^3J_{HH}$ = 8.2 Hz, 1H, C$_{py}$H), 8.07 (td, $^3J_{HH}$ = 7.4 Hz, $^4J_{HH}$ = 4.8 Hz, 1H, C$_{py}$H), 4.65 (q, $^3J_{HH}$ = 7.0 Hz, 2H, OCH$_2$Me), 4.23 (s, 3H, NCH$_3$), 1.60 (t, $^3J_{HH}$ = 7.0 Hz, 3H, OCH$_2$CH_3). ^{13}C{^{1}H} NMR (100 MHz, CDCl$_3$): δ = 153.9 (C$_{trz}$–OEt), 149.4 (C$_{py}$H), 140.5 (C$_{py}$H), 127.1 (C$_{py}$H), 115.1 (C$_{py}$H), 108.6 (C$_{trz}$H), 72.9 (OCH$_2$Me), 36.1 (NCH$_3$), 14.5 (OCH$_2$CH$_3$), (C$_{py}$–N$_{trz}$) n.d. HR-MS (CH$_3$CN): m/z calculated for C$_{10}$H$_{13}$ON$_4$ [M–OTf]$^+$ = 205.1084; found, 205.1079.

Compound 3c

a) Synthesis of 2-(4-ethoxy-1H-1,2,3-triazol-1-yl)-4-methoxypyridine. The reaction of 2-azido-4-methoxypyridine (500 mg, 3.33 mmol), ethoxyacetylene (2.4 mL, 9.98 mmol), CuI (635 mg, 3.33 mmol) and 2,6-lutidine (770 µL, 6.65 mmol) in THF (27 mL) and DMSO (470 µL) was stirred under reflux for 72 h and afforded this compound as an off-white solid (270 mg, 36%). ^{1}H NMR (300 MHz, CDCl$_3$): δ = 8.23 (d, $^3J_{HH}$ = 5.8 Hz, 1H, C$_{py}$H), 7.96 (s, 1H, C$_{trz}$H), 7.66 (d, $^4J_{HH}$ = 2.3 Hz, 1H, C$_{py}$H), 6.82 (dd, $^3J_{HH}$ = 5.8 Hz, $^4J_{HH}$ = 2.3 Hz, 1H, C$_{py}$H), 4.29 (q, $^3J_{HH}$ = 7.0 Hz, 2H, OCH$_2$Me), 3.93 (s, 3H, OCH$_3$), 1.44 (t, $^3J_{HH}$ = 7.0 Hz, 3H, OCH$_2$CH_3).

b) Synthesis of triazolium salt 3c. To a solution of the 2-(4-ethoxy-triazol-1-yl)-4-methoxypyridine (190 mg, 0.86 mmol) in CH_2Cl_2 at 0 ºC was added MeOTf (110 µL, 0.99 mmol) and the mixture was stirred at 0 ºC for 15 min. Addition of Et_2O gave a white precipitate (150 mg, 45%). ^{1}H NMR (300 MHz, $CDCl_3$): δ = 8.78 (s, 1H, C$_{trz}$H), 8.36 (d, $^3J_{HH}$ = 5.8 Hz, 1H, C$_{py}$H), 7.69 (d, $^4J_{HH}$ = 2.2 Hz, 1H, C$_{py}$H), 7.05 (dd, $^3J_{HH}$ = 5.8 Hz, $^4J_{HH}$ = 2.2 Hz, 1H, C$_{py}$H), 4.63 (q, $^3J_{HH}$ = 7.0 Hz, 2H, OCH$_2$Me), 4.23 (s, 3H, NCH$_3$), 4.02 (s, 3H, OCH$_3$), 1.60 (t, $^3J_{HH}$ = 7.0 Hz, 3H, OCH$_2$C*H*$_3$).

General procedure for the synthesis of the complexes 4–6: The corresponding triazolium salt **1–3** (1 eq), Ag$_2$O (2 eq), Me$_4$NCl (1.5 eq) and [RuCl$_2$(p-cymene)]$_2$ (0.5 eq), were suspended in CH_2Cl_2 and stirred under exclusion of light at 40 ºC for 24–48 h. The reaction was filtered through Celite and all volatiles were evaporated under reduced pressure. The crude solid was purified by gradient column chromatography (SiO$_2$; CH_2Cl_2 to CH_2Cl_2/acetone 10:2). Precipitation with CH_2Cl_2/Et_2O gave the title complexes as pure orange solids.

Complex 4b

According to the general procedure, the triazolium salt **1b** (150 mg, 0.49 mmol), Ag$_2$O (225 mg, 0.98 mmol), Me$_4$NCl (80 mg, 0.74 mmol) and [RuCl$_2$(p-cymene)]$_2$ (149 mg, 0.25 mmol), were suspended in CH_2Cl_2 (12 mL) and stirred under exclusion of light at 40 ºC for 24 h. The reaction mixture was worked-up and purified as described in the general procedure and gave complex **4b** as a pure orange solid (190 mg, 57%). ^{1}H NMR (400 MHz, $CDCl_3$): δ = 8.79 (d, $^4J_{HH}$ = 2.6 Hz, 1H, C$_{py}$H), 8.07 (d, $^3J_{HH}$ = 9.0 Hz, 1H, C$_{py}$H), 7.65 (dd, $^3J_{HH}$ = 9.0 Hz, $^4J_{HH}$ = 2.6 Hz, 1H, C$_{py}$H), 6.24 (d, $^3J_{HH}$ = 5.8 Hz, 1H, C$_{cym}$H), 6.18 (d, $^3J_{HH}$ = 6.2 Hz, 1H, C$_{cym}$H), 5.91 (d, $^3J_{HH}$ = 6.2 Hz, 1H, C$_{cym}$H), 5.61 (d, $^3J_{HH}$ = 5.8 Hz, 1H, C$_{cym}$H), 4.59 (s, 3H, NCH$_3$), 4.65–4.56 (m, 2H, OCH$_2$Me), 4.08 (s, 3H, OCH$_3$), 2.70–2.53 (m, 1H, CHMe$_2$), 2.14 (s, 3H, C$_{cym}$–CH$_3$), 1.53 (t, $^3J_{HH}$ = 7.1 Hz, 3H, OCH$_2$C*H*$_3$), 1.09 (d, $^3J_{HH}$ = 6.8 Hz, 3H, CHC*H*$_3$), 0.99 (d, $^3J_{HH}$ = 6.8 Hz, 3H, CHC*H*$_3$). ^{13}C{^{1}H} NMR (100 MHz, $CDCl_3$): δ = 175.3 (C$_{trz}$–Ru), 157.9 (C=O), 157.4 (C$_{py}$–OMe), 143.2 (C$_{py}$–N$_{trz}$), 142.4 (C$_{py}$H), 138.9 (*C*$_{trz}$–COOEt), 125.8 (C$_{py}$H), 115.2 (C$_{py}$H), 108.5, 105.1 (2 × *C*$_{cym}$–C), 90.2, 89.0, 87.6, 84.3 (4 × C$_{cym}$H), 63.3 (OCH$_2$Me), 57.3 (OCH$_3$), 42.7 (NCH$_3$), 31.3 (CHMe$_2$), 22.6, 22.4 (2 × CH–CH$_3$), 19.0 (C$_{cym}$–CH$_3$), 14.4 (OCH$_2$C*H*$_3$). Anal. Calcd for C$_{23}$H$_{28}$ClF$_3$RuN$_4$O$_6$S (682.07): C, 40.50; H, 4.14; N, 8.21. Found: C, 40.35; H, 3.84; N, 7.95. HR-MS (CH$_3$CN): m/z calculated for C$_{22}$H$_{28}$O$_3$N$_4$ClRu [M–OTf]$^+$ = 533.0888; found, 533.0882.

Complex 4c

The triazolium salt **1c** (200 mg, 0.49 mmol), Ag_2O (225 mg, 0.97 mmol), Me_4NCl (80 mg, 0.73 mmol) and $[RuCl_2(p\text{-cymene})]_2$ (149 mg, 0.24 mmol), were suspended in CH_2Cl_2 (17 mL) and stirred under exclusion of light at 40 °C for 48 h. The reaction was filtered through Celite and purified by column chromatography as described in the general procedure to give complex **4c** as a pure orange solid (110 mg, 33%). ^{1}H NMR (400 MHz, $CDCl_3$): δ = 8.98 (d, $^3J_{HH}$ = 6.6 Hz, 1H, $C_{py}H$), 7.55 (d, $^4J_{HH}$ = 2.6 Hz, 1H, $C_{py}H$), 7.12 (dd, $^3J_{HH}$ = 6.6 Hz, $^4J_{HH}$ = 2.6 Hz, 1H, $C_{py}H$), 6.13 (d, $^3J_{HH}$ = 6.0 Hz, 1H, $C_{cym}H$), 6.09 (d, $^3J_{HH}$ = 6.1 Hz, 1H, $C_{cym}H$), 5.86 (d, $^3J_{HH}$ = 6.1 Hz, 1H, $C_{cym}H$), 5.51 (d, $^3J_{HH}$ = 6.0 Hz, 1H, $C_{cym}H$), 4.63–4.49 (m, 3H, NCH_3, 2H, OCH_2Me), 3.98 (s, 3H, OCH_3), 2.60–2.46 (m, 1H, $CHMe_2$), 2.07 (s, 3H, $C_{cym}–CH_3$), 1.47 (t, $^3J_{HH}$ = 7.2 Hz, 3H, OCH_2CH_3), 1.00 (d, $^3J_{HH}$ = 6.9 Hz, 3H, $CHCH_3$), 0.91 (d, $^3J_{HH}$ = 6.9 Hz, 3H, $CHCH_3$). $^{13}C\{^1H\}$ NMR (100 MHz, $CDCl_3$): δ = 177.8 ($C_{trz}–Ru$), 169.5 ($C_{py}–OMe$), 157.9 (C=O), 156.2 ($C_{py}H$), 150.9 ($C_{py}–N_{trz}$), 139.0 ($C_{trz}–COOEt$), 114.1 ($C_{py}H$), 107.7, 105.3 (2 × $C_{cym}–C$), 100.8 ($C_{py}H$), 89.9, 88.5, 87.9, 84.1 (4 × $C_{cym}H$), 63.4 (OCH_2Me), 57.3 (OCH_3), 42.9 (NCH_3), 31.3 ($CHMe_2$), 22.7, 22.4 (2 × $CH–CH_3$), 19.1 ($C_{cym}–CH_3$), 14.4 (OCH_2CH_3). Anal. Calcd for $C_{23}H_{28}ClF_3RuN_4O_6S$ (682.07): C, 40.50; H, 4.14; N, 8.21. Found: C, 40.27; H, 3.75; N, 7.98. HR-MS (CH_3CN): m/z calculated for $C_{22}H_{28}O_3N_4ClRu$ $[M–OTf]^+$ = 533.0888; found, 533.0865.

Complex 5

According to the general procedure, the triazolium salt **2** (180 mg, 0.51 mmol), Ag_2O (237 mg, 1.20 mmol), Me_4NCl (84 mg, 0.77 mmol) and $[RuCl_2(p\text{-cymene})]_2$ (157 mg, 0.26 mmol), were suspended in CH_2Cl_2 (15 mL) and stirred under exclusion of light at 40 °C for 48 h. The reaction was filtered through Celite and the crude solid was purified by gradient column chromatography as described in the general procedure affording complex **5** as a pure orange solid (120 mg, 38%). ^{1}H NMR (400 MHz, $CDCl_3$): δ = 9.27 (d, $^3J_{HH}$ = 5.5 Hz, 1H, $C_{py}H$), 8.13–8.01 (m, 2H, $C_{py}H$), 7.64–7.58 (m, 1H, $C_{py}H$), 6.09 (d, $^3J_{HH}$ = 6.0 Hz, 1H, $C_{cym}H$), 6.06 (d, $^3J_{HH}$ = 6.2 Hz, 1H, $C_{cym}H$), 5.83 (d, $^3J_{HH}$ = 6.2 Hz, 1H, $C_{cym}H$), 5.64 (d, $^3J_{HH}$ = 6.0 Hz, 1H, $C_{cym}H$), 4.31 (s, 3H, NCH_3), 3.36–3.10 (m, 2H, $C_{trz}CH_2$), 2.58 (septet, $^3J_{HH}$ = 6.9 Hz, 1H, $CHMe_2$), 2.07 (s, 3H, $C_{cym}–CH_3$), 1.92–1.68 (m, 2H, $CH_2CH_2CH_3$), 1.10 (t, $^3J_{HH}$ = 7.3 Hz, 3H, CH_2CH_3), 1.05 (d, $^3J_{HH}$ = 7.0 Hz, 3H, $CHCH_3$), 1.04 (d, $^3J_{HH}$ = 7.0 Hz, 3H, $CHCH_3$). $^{13}C\{^1H\}$ NMR (100 MHz, $CDCl_3$): δ = 167.9 ($C_{trz}–Ru$), 155.8 ($C_{py}H$), 150.7 ($C_{py}–N_{trz}$), 149.1 ($C_{trz}–nPr$), 141.2 ($C_{py}H$), 126.2 ($C_{py}H$), 114.3 ($C_{py}H$), 107.0, 100.7 (2 × $C_{cym}–C$), 90.5, 87.7, 86.6, 85.4 (4 × $C_{cym}H$), 37.8 (NCH_3), 31.3 ($CHMe_2$), 27.6 ($C_{trz}–CH_2$), 22.7, 22.4, 22.3 (2 × $CH–CH_3$, $CH_2CH_2CH_3$), 18.8 ($C_{cym}–CH_3$), 14.3 (CH_2CH_3). Anal. Calcd for $C_{22}H_{28}ClF_3RuN_4O_3S$ (622.07): C, 42.48; H, 4.54; N, 9.01. Found: C, 42.08; H, 4.25; N, 9.04. HR-MS (CH_3CN): m/z calculated for $C_{21}H_{28}N_4ClRu$ $[M–OTf]^+$ = 473.1046; found, 473.1025.

Complex 6a

The triazolium salt **3a** (160 mg, 0.45 mmol), Ag_2O (210 mg, 0.90 mmol), Me_4NCl (74 mg, 0.68 mmol) and $[RuCl_2(p\text{-cymene})]_2$ (138 mg, 0.23 mmol), were suspended in CH_2Cl_2 (12 mL) and stirred under exclusion of light at 40 ºC for 48 h. Following the general procedure, complex **6a** was obtained as a pure orange solid (70 mg, 25%). ^1H NMR (400 MHz, $CDCl_3$): δ = 9.42 (d, $^3J_{HH}$ = 5.7 Hz, 1H, $C_{py}H$), 8.15–8.00 (m, 2H, $C_{py}H$), 7.71–7.68 (m, 1H, $C_{py}H$), 7.69 (t, $^3J_{HH}$ = 6.4 Hz, 1H, $C_{py}H$), 6.18 (d, $^3J_{HH}$ = 6.1 Hz, 1H, $C_{cym}H$), 6.10 (d, $^3J_{HH}$ = 6.0 Hz, 1H, $C_{cym}H$), 5.73 (d, $^3J_{HH}$ = 6.1 Hz, 1H, $C_{cym}H$), 5.36 (d, $^3J_{HH}$ = 6.0 Hz, 1H, $C_{cym}H$), 5.41–4.69 (m, 2H, OCH_2Me), 4.19 (s, 3H, NCH_3), 2.66–2.53 (m, 1H, $CHMe_2$), 2.17 (s, 3H, $C_{cym}–CH_3$), 1.52 (t, $^3J_{HH}$ = 7.1 Hz, 3H, OCH_2CH_3), 1.11 (d, $^3J_{HH}$ = 6.9 Hz, 3H, $CHCH_3$), 1.00 (d, $^3J_{HH}$ = 6.9 Hz, 3H, $CHCH_3$). $^{13}C\{^1H\}$ NMR (100 MHz, $CDCl_3$): δ = 158.5 ($C_{trz}–OEt$), 156.9 ($C_{py}H$), 155.2 ($C_{trz}–Ru$), 150.2 ($C_{py}–N_{trz}$), 141.4 ($C_{py}H$), 127.2 ($C_{py}H$), 113.7 ($C_{py}H$), 108.3, 104.9 (2 × $C_{cym}–C$), 89.9, 87.2, 86.0, 82.8 (4 × $C_{cym}H$), 74.2 (OCH_2Me), 35.5 (NCH_3), 31.4 ($CHMe_2$), 22.4, 22.3 (2 × $CH–CH_3$), 19.1 ($C_{cym}–CH_3$), 15.7 (OCH_2CH_3). Anal. Calcd for $C_{21}H_{26}ClF_3RuN_4O_4S$ (624.04): C, 40.42; H, 4.20; N, 8.98. Found: C, 40.55; H, 4.32; N, 8.59. HR-MS (CH_3CN): m/z calculated for $C_{20}H_{26}ON_4ClRu$ [M–OTf]$^+$ = 475.0839; found, 475.0826.

Complex 6b

The triazolium salt **3b** (138 mg, 0.36 mmol), Ag_2O (166 mg, 0.72 mmol), Me_4NCl (60 mg, 0.54 mmol) and $[RuCl_2(p\text{-cymene})]_2$ (110 mg, 0.18 mmol), were suspended in CH_2Cl_2 (12 mL) and stirred under exclusion of light at 40 ºC for 48 h. The reaction mixture was treated according to the general procedure to obtain complex **6b** as a pure orange solid (50 mg, 21%). ^1H NMR (400 MHz, $CDCl_3$): δ = 8.81 (d, $^4J_{HH}$ = 2.4 Hz, 1H, $C_{py}H$), 7.96 (d, $^3J_{HH}$ = 9.1 Hz, 1H, $C_{py}H$), 7.63 (dd, $^3J_{HH}$ = 9.1 Hz, $^4J_{HH}$ = 2.4 Hz, 1H, $C_{py}H$), 6.10 (d, $^3J_{HH}$ = 6.1 Hz, 2H, $C_{cym}H$), 5.71 (d, $^3J_{HH}$ = 6.1 Hz, 1H, $C_{cym}H$), 5.39 (d, $^3J_{HH}$ = 6.1 Hz, 1H, $C_{cym}H$),4.67–4.40 (m, 2H, OCH_2Me), 4.16 (s, 3H, NCH_3), 4.10 (s, 3H, OCH_3), 2.73–2.58 (m, 1H, $CHMe_2$), 2.16 (s, 3H, $C_{cym}–CH_3$), 1.52 (t, $^3J_{HH}$ = 7.1 Hz, 3H, OCH_2CH_3), 1.17 (d, $^3J_{HH}$ = 6.9 Hz, 3H, $CHCH_3$), 1.07 (d, $^3J_{HH}$ = 6.9 Hz, 3H, $CHCH_3$). $^{13}C\{^1H\}$ NMR (100 MHz, $CDCl_3$): δ = 158.4 ($C_{trz}–OEt$), 157.4 ($C_{py}–OMe$), 153.4 ($C_{trz}–Ru$), 143.8 ($C_{py}–N_{trz}$), 142.6 ($C_{py}H$), 126.4 ($C_{py}H$), 114.4 ($C_{py}H$), 108.9, 103.9 (2 × $C_{cym}–C$), 89.1, 87.8, 85.5, 82.3 (4 × $C_{cym}H$), 74.2 (OCH_2Me), 57.3 (OCH_3), 35.2 (NCH_3), 31.5 ($CHMe_2$), 22.6, 22.2 (2 × $CH–CH_3$), 19.1 ($C_{cym}–CH_3$), 15.7 (OCH_2CH_3). Anal. Calcd for $C_{22}H_{28}ClF_3RuN_4O_5S$ + 1/2 Et_2O (691.12): C, 41.71; H, 4.81; N, 8.11. Found: C, 41.49; H, 4.55; N, 8.11. HR-MS (CH_3CN): m/z calculated for $C_{21}H_{28}O_2N_4ClRu$ [M–OTf]$^+$ = 505.0944; found, 505.0922.

Complex 6c

According to the general procedure, the triazolium salt **3c** (125 mg, 0.33 mmol), Ag_2O (151 mg, 0.66 mmol), Me_4NCl (54 mg, 0.49 mmol) and $[RuCl_2(p\text{-cymene})]_2$ (100 mg, 0.17 mmol), were suspended in CH_2Cl_2 (12 mL) and stirred under exclusion of light at 40 ºC for 24 h. The reaction was filtered through Celite and purified by gradient column chromatography as described in the general procedure to afford complex **6c** as a pure orange solid (70 mg, 33%). ^1H NMR (400 MHz, $CDCl_3$): δ = 9.14 (d, $^3J_{HH}$ = 6.6 Hz, 1H, $C_{py}H$), 7.49 (d, $^4J_{HH}$ = 2.6 Hz, 1H, $C_{py}H$), 7.22 (dd, $^3J_{HH}$ = 6.6 Hz, $^4J_{HH}$ = 2.6 Hz, 1H, $C_{py}H$), 6.13 (d, $^3J_{HH}$ = 6.1 Hz, 1H, $C_{cym}H$), 6.02 (d, $^3J_{HH}$ = 5.9 Hz, 1H, $C_{cym}H$), 5.68 (d, $^3J_{HH}$ = 6.1 Hz, 1H, $C_{cym}H$), 5.28 (d, $^3J_{HH}$ = 5.9 Hz, 1H, $C_{cym}H$), 4.67–4.41 (m, 2H, OCH_2Me), 4.18 (s, 3H, NCH_3), 4.02 (s, 3H, OCH_3), 2.62–2.50 (m, 1H, $CHMe_2$), 2.15 (s, 3H, $C_{cym}–CH_3$) 1.50 (t, $^3J_{HH}$ = 7.1 Hz, 3H, OCH_2CH_3), 1.08 (d, $^3J_{HH}$ = 6.9 Hz, 3H, $CHCH_3$), 0.97 (d, $^3J_{HH}$ = 6.9 Hz, 3H, $CHCH_3$). $^{13}C\{^1H\}$ NMR (100 MHz, $CDCl_3$): δ = 169.4 (C_{py}–OMe), 158.4 (C_{trz}–OEt), 157.2 ($C_{py}H$), 155.5 (C_{trz}–Ru), 151.2 (C_{py}–N_{trz}), 113.5 ($C_{py}H$), 107.3, 104.7 (2 × C_{cym}–C), 100.4 ($C_{py}H$), 89.7, 86.4, 85.9, 82.1 (4 × $C_{cym}H$), 74.2 (OCH_2Me), 57.3 (OCH_3), 35.4 (NCH_3), 31.4 ($CHMe_2$), 22.4, 22.3 (2 × CH–CH_3), 19.0 (C_{cym}–CH_3), 15.7 (OCH_2CH_3). Anal. Calcd for $C_{22}H_{28}ClF_3RuN_4O_5S$ (654.06): C, 40.40; H, 4.32; N, 8.57. Found: C, 40.25; H, 3.56; N, 8.10. HR-MS (CH_3CN): m/z calculated for $C_{21}H_{28}O_2N_4ClRu$ $[M–OTf]^+$ = 505.0944; found, 505.0919.

Complex 7

To a solution of complex **5** (100 mg, 0.16 mmol) in MeCN (10 mL) was added AgOTf (61 mg, 0.24 mmol), and the suspension was stirred at reflux for 18 h. The reaction mixture was filtered through Celite, and the solvent was removed under reduced pressure. Precipitation with MeCN/Et_2O afforded a spectroscopically pure yellow solid. Yield: 80 mg, 66%. ^1H NMR (400 MHz, CD_3CN): δ = 8.95 (d, $^3J_{HH}$ = 5.6 Hz, 1H, $C_{py}H$), 8.22–8.11 (m, 2H, $C_{py}H$), 7.62 (ddd, $^3J_{HH}$ = 7.3 Hz, $^3J_{HH}$ = 5.6 Hz, $^4J_{HH}$ = 1.7 Hz, 1H, $C_{py}H$), 4.26 (s, 3H, NCH_3), 3.03 (t, $^3J_{HH}$ = 7.5 Hz, 2H, $C_{trz}CH_2$), 2.54 (s, 3H, CH_3CN), 2.07 (s, 6H, CH_3CN), 1.78 (sextet, $^3J_{HH}$ = 7.5 Hz, 2H, $CH_2CH_2CH_3$), 1.02 (t, $^3J_{HH}$ = 7.5 Hz, 3H, CH_2CH_3). $^{13}C\{^1H\}$ NMR (100 MHz, CD_3CN): δ = 172.5 (C_{trz}–nPr), 154.6 ($C_{py}H$), 154.4 (C_{py}–N_{trz}), 150.3 (C_{trz}–Ru), 141.7 ($C_{py}H$), 126.9 ($C_{py}H$), 114.4 ($C_{py}H$), 38.0 (NCH_3), 26.7 (C_{trz}–CH_2), 23.3 ($CH_2CH_2CH_3$), 13.8 (CH_2CH_3). HR-MS (CH_3CN): m/z calculated for $C_{20}H_{26}F_3N_8O_3RuS$ $[M–OTf]^+$ = 617.0839; found, 617.0841.

General procedure for the 4-methylbenzylamine oxidation. In a 100 mL Schlenk flask, a mixture of complex (0.01 mmol), 4-methylbenzylamine (0.2 mmol) and 1,3,5-trimethoxybenzene (5.6 mg, 0.033 mmol; internal standard) in 1,2-dichlorobenzene (2 mL) was mixed. Then, molecular oxygen was bubbled into the solution for 5 min and 6 mL of gaseous ammonia was injected. An oxygen balloon was connected to the reaction vessel *via* a septum. The reaction mixture was heated to 150 ºC. Aliquots were taken at specific times, diluted with $CDCl_3$ and analyzed by ^1H NMR spectroscopy. Conversions and yields were determined relative to

1,3,5-trimethoxybenzene as internal standard. Products were isolated at the end of the reaction by cooling the mixture to room temperature, and subsequent purification by column chromatography using SiO2 as stationary phase and elution first with pentane (100 mL) to remove the 1,2-dichlorobenzene, followed by elution with pentane:Et$_2$O (either 80:20 or 90:10). The NMR spectra of the isolated products showed less than 5% of aldehyde (due to imine hydrolysis).

Crystal structure determinations: Crystal data for **4b**, **4c**, **6a** and **7** were collected using an Oxford Diffraction SuperNova area-detector diffractometer[52] with mirror optics monochromated Mo *K*α radiation (λ = 0.71073 Å) and Al filtered.[53] Data reduction was performed using the CrysAlisPro program. The intensities were corrected for Lorentz and polarization effects, and numerical absorption correction based on gaussian integration over a multifaceted crystal model was applied. The structure was solved by direct methods using SHELXT,[54] which revealed the positions of all not disordered non-hydrogen atoms of the title compound. The non-hydrogen atoms were refined anisotropically. All H-atoms were placed in geometrically calculated positions and refined using a riding model where each H-atom was assigned a fixed isotropic displacement parameter with a value equal to 1.2Ueq of its parent atom (1.5Ueq for the methyl groups and water). Refinement of the structure was carried out on F^2 using full-matrix least-squares procedures, which minimized the function $\Sigma w(F_o^2 - F_c^2)^2$. The weighting scheme was based on counting statistics and included a factor to downweight the intense reflections. All calculations were performed using the SHELXL-2014/7 program.[55] Further crystallographic details are compiled in Tables S2–5. Crystallographic data for all structures have been deposited with the Cambridge Crystallographic Data Centre (CCDC) as supplementary publication numbers **4b** (1958588), **4c** (1958592), **6a** (1958633) and **7** (1958593).

Electronic Supporting Information
Synthetic procedures for ligand precursors, NMR spectra of all complexes, catalytic and crystallographic details.

Conflicts of interest
There are no conflicts to declare.

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