

Organometallic Nickel Complexes Supported by Pyridinophane Ligands and Their Oxidative Reactivity

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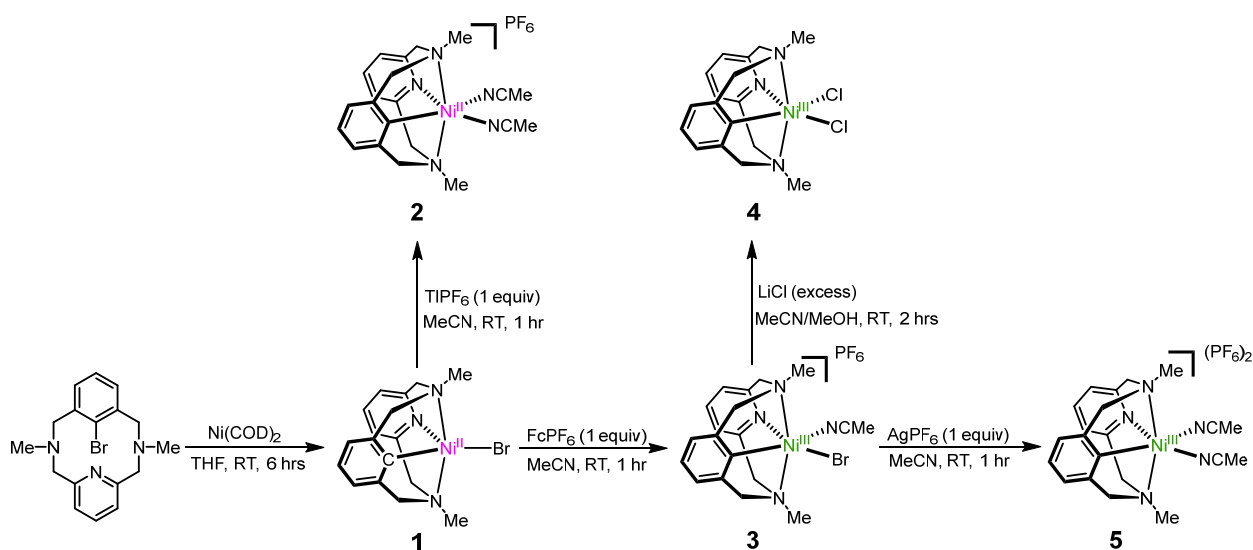
ABSTRACT

The pyridinophane ligands $^R\text{N}3\text{CX}$ ($X = \text{H}, \text{Br}$) are well-established scaffolds that facilitate and stabilize nickel oxidative addition complexes to the proximal C(aryl)–X bond. In this study, we report the synthesis, detailed characterization, and reactivity of a series of Ni(II) and Ni(III) complexes supported by the $^{\text{Me}}\text{N}3\text{CX}$ ligand. Our findings demonstrate that Ni(II) complexes can be oxidized both chemically and aerobically to yield well-defined Ni(III) species. Excitingly, the Ni-disolvento complexes exhibit catalytic trifluoroethoxylation to generate the C–O coupled product. In addition, the Ni(III)-halide complex undergoes transmetalation with a Grignard reagent and subsequent C–C reductive elimination, while the β -hydride elimination side reaction is suppressed, outperforming its Ni(II) analogue.

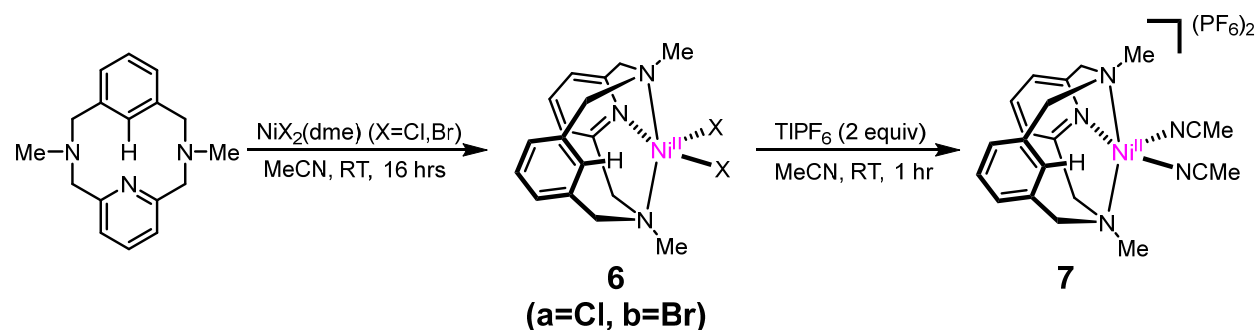
INTRODUCTION

High-valent organometallic nickel complexes have emerged as fascinating subjects of study due to their unique properties and potential applications in catalysis to forge new chemical bonds. Among the various ligands employed to stabilize Ni(III) centers, the tetradentate pyridinophane ligands, $^R\text{N}3\text{CX}$, have supported high-valent Ni species with remarkable stability.¹ Understanding the stability of these complexes is crucial for reactivity study, and in particular, the role of substituent effects has been a topic of interest.^{2–5} In this regard, replacing the tert-butyl (^tBu) N-substituents with the less bulky neopentyl (Np) groups (yielding the $^{\text{Np}}\text{N}3\text{C}$ system) has been found to reduce stability and enhance the reactivity of the high valent nickel species compared to the ^tBuN3CX system.^{4–6} Since the introduction of these systems, numerous metalloorganic systems

based on this framework and incorporating first- and second-row transition metals such as Mn, Fe, Co, and Pd have been utilized to investigate their reactivity and catalytic processes.⁷⁻¹⁶ Building upon the remarkable reactivity enhancement observed with Np N-substituents,⁴⁻⁵ our current investigation explores the potential of the less bulky N-methyl analogues in stabilizing Ni(III) intermediates.⁷ In this case, the use of these methyl-substituted ligands aimed to elucidate the influence of steric effects on the reactivity of the nickel complexes. To generate the Ni(III) centers, we employed both aerobic and anaerobic oxidants, allowing us to compare the reactivity and stability of the resulting complexes under different conditions. The investigation aimed to shed light on the factors influencing the strength and significance of the agostic interactions between the Ni center and the ipso C(aryl)-H bond. Finally, in addition to stabilizing Ni(III) intermediates and studying agostic interactions, we examined the oxidative addition of low-valent nickel systems into C-H bonds. This step allowed us to investigate the feasibility of C-H bond activation and its potential as a key step in various catalytic transformations.



Scheme 1. Synthesis of ($\text{Me}^c\text{N}_3\text{C}$)Ni complexes.



Scheme 2. Synthesis of (^{Me}N₃CH)Ni complexes.

RESULTS AND DISCUSSION

The Ni(II) complex (^{Me}N₃C)Ni^{II}Br (**1**) can be synthesized upon addition of Ni⁰(COD)₂ to ^{Me}N₃CBr following a modified literature procedure (Scheme 1).⁷ The complex **1** can be oxidized by addition of one equivalent of FcPF₆ to yield the Ni(III) complex [(^{Me}N₃C)Ni^{III}Br(MeCN)]PF₆ (**3**). The solid-state structure of **3** could not be obtained, even upon substitution with different counterions (BARF₂₄⁻, OTf⁻, or BF₄⁻), and complex identity was confirmed by ESI-MS and CHN analysis. Halide abstraction of the bromide from **3** by addition of one equivalent of TIPF₆, or abstraction and oxidation of **1** by addition of two equivalents of AgPF₆ yields the bis-solvento complex [(^{Me}N₃C)Ni^{III}(MeCN)₂](PF₆)₂ (**5**). Both **3** and **5** are stable in air at RT. Evan's method analysis of **3** and **5** returns magnetic moments of 1.89 and 1.80 μ_B, indicating that both are paramagnetic with one unpaired electron, as expected for a d⁷ Ni(III) center. Structural characterization for **5** reveals an octahedral geometry as expected for d⁷ ions, with the amine donors in the axial positions (Figure 1). Their EPR characterizations (77 K, PrCN glass) show the presence of axial signals (Figure 2) with *g*_{ave} values of 2.125 for **3**, along with superhyperfine coupling to the bromide (*I* = 3/2) is observed in the *g*_y direction, and a value of 2.109 for **5**, along with superhyperfine coupling to the two axial N donors (*I* = 1) in the *g*_z direction. Since the solid-state structure for complex **3** is elusive, excess LiCl was added for halide exchange to give rise to **4**.⁷ The solid-state structure of **4** is also consistent with the previous observations for this system. However, a rhombic EPR signal was observed with *g*_{ave} 2.132, with no superhyperfine coupling to any halide. Taken together, the observed structural and EPR parameters for complexes **3**, **4**, and **5** strongly suggest the presence of a distorted octahedral Ni^{III} d⁷ center with a d_z² ground state.

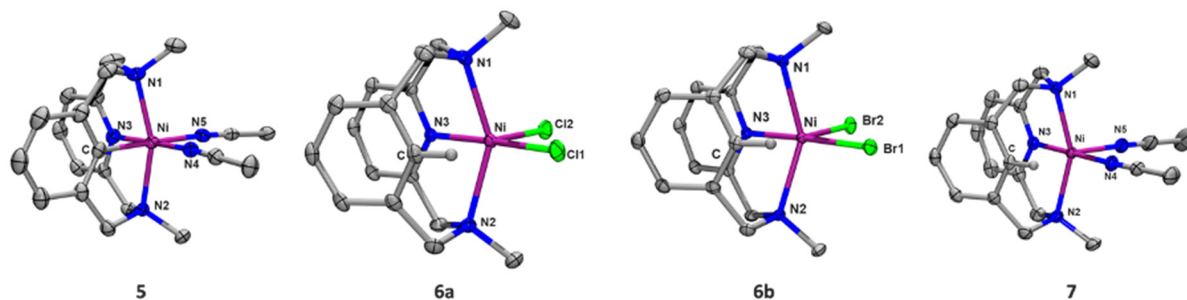


Figure 1. ORTEPs (50% probability thermal ellipsoids) of complexes **5**, **6a**, **6b**, and **7**. For clarity, hydrogen atoms and counterions have been omitted. Selected bond distances (Å), **5**: Ni–N1, 2.152(2); Ni–N2, 2.142(2); Ni–N3, 1.935(2); Ni–N4, 1.983(2); Ni–N5, 1.990(2); Ni–C, 1.921(2). **6a**: Ni–N1, 2.233(2); Ni–N2, 2.242(2); Ni–N3, 1.993(2); Ni–C, 2.539(2); Ni–Cl1, 2.312(5); Ni–Cl2, 2.329(5). **6b**: Ni–N1, 2.223(1); Ni–N2, 2.255(1); Ni–N3, 1.985(1); Ni–C, 2.482(2); Ni–Br1, 2.455(2); Ni–Br2, 2.478(2). **7**: Ni–N1, 2.179(2); Ni–N2, 2.169(2); Ni–N3, 1.971(2); Ni–N4, 2.023(2); Ni–N5, 2.037(2); Ni–C, 2.416(2).

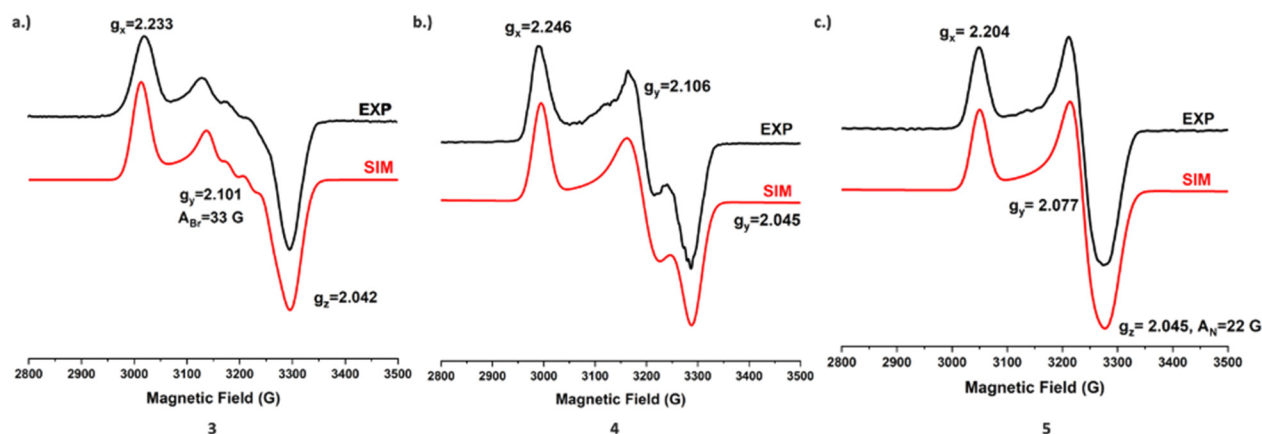


Figure 2. The experimental and simulated EPR spectra of **3**, **4**, and **5** in PrCN at 77 K. Simulations were obtained using the following parameters: **1**: $g_x = 2.233$; $g_y = 2.101$ ($A_{Br} = 33$ G); $g_z = 2.042$; **2**: $g_x = 2.246$; $g_y = 2.106$; $g_z = 2.045$; **3**: $g_x = 2.204$; $g_y = 2.077$; $g_z = 2.025$ ($A_{2N} = 22$ G).

Complexes ($^{Me}N_3CH$)Ni^{II}Cl₂ (**6a**) and ($^{Me}N_3CH$)Ni^{II}Br₂ (**6b**) were synthesized by stirring the corresponding NiX₂(dme) precursor with $^{Me}N_3CH$ overnight, in 67% and 67% yields, respectively (Scheme 2). Halide abstraction via addition of 2 equivalents of TIPF₆ yielded the bis-solvento complex ($^{Me}N_3CH$)Ni^{II}(MeCN)₂(PF₆)₂ (**7**) in 85% yield, and no C_{sp2}–H activation was

observed during the synthesis of **7**. Spin state analysis by Evan's method of **6a**, **6b**, and **7** yielded magnetic moments of 3.16 μ_B , 3.12 μ_B , and 2.96 μ_B , respectively, indicating that the three complexes are paramagnetic with two unpaired electrons. The solid-state structures show distorted square pyramidal geometries, with τ_5 values of 0.40, 0.42, and 0.41 respectively. Relatively short Ni–C lengths (2.539 Å, 2.482 Å, and 2.416 Å) and narrow Ni–H–C bond angles (94.0°, 96.3°, and 83.5°) are indicative of an agostic interaction in all three (^{Me}N₃CH)Ni^{II} complexes (Figure 1). Notably, the Ni–H–C angle and Ni–C bond length increases as the σ -donor ancillary ligands are replaced by π -donor ligands. Even though these complexes are all prone to C–H activation, we postulate that the complex bearing σ -donor ancillary ligands are more prone to activation due to potentially stronger agostic interactions. It is known that C–H agostic interaction is an electronic effect which involves the donation of electron density associated with the C–H bond to a metal center.¹⁷ Therefore, computational studies were employed in order to investigate the electronic structures. Density functional theory (DFT) calculations of the gas-phase structure of **6b** confirmed the non-bonding interactions of the C–H bond with the metal frontier orbitals (Figure 3). Natural bond orbital (NBO) analysis of the HOMO further confirmed a weaker overlap between the σ (C–H)-bond and the d_z^2 -orbital of nickel with a weak second order perturbation energy ($E_2 = 2.3 \text{ kcal}\cdot\text{mol}^{-1}$). In a reverse fashion, the LUMO analysis shows the interaction of the metal orbital with the antibonding $\sigma^*(\text{C–H})$ bond, with an energy of 3.1 $\text{kcal}\cdot\text{mol}^{-1}$. We posit that these non-bonding interactions weakens the C–H bond and confirms the agostic interactions observed in these systems. Our laboratory is currently conducting further research on the activation of these C–H bonds.

Ligand effect on properties of (^RN₃CX)Ni complexes

Characterization of the bis-solvento complexes **5** and **7** has allowed us to incorporate the (^{Me}N₃CX)Ni series into the metrical trends observed in the (^RN₃C)Ni systems bearing Np, ^tBu, and H N-substituents.^{4-5, 18} It was previously noted that moving from the ^tBuN₃C[−] system to the less sterically hindered ^{Np}N₃C[−] system led to an increase in the axial amine donation, as evidenced by a decrease in the average axial Ni–N bond length (2.302 Å vs 2.239 Å), and an increase in the superhyperfine coupling observed in the g_z direction (10 G vs. 14 G). Gratifyingly, the use of the ^{Me}N₃C[−] ligand corroborates this finding, bearing a significantly shorter average Ni–N bond distance of 2.147 Å, and a stronger superhyperfine coupling of 22 G with the axial N donors in the g_z direction. This trend also extends to the (^RN₃CH)Ni complexes, though it is less pronounced.

The (^RN3CH)NiBr₂ complexes for the N-tBu, N-Np, and N-Me derivative exhibit average axial Ni–N bond distances of 2.608 Å, 2.309 Å, and 2.239 Å, respectively, demonstrating a clear decrease (Figure 1). Similarly, when comparing the Ni-bis(solvento) complexes, the axial N donor atoms move much closer for H (2.158 Å) compared to Me (2.239 Å). However, for comparing Ni–C distances, the trend becomes much less clear. The Np and Me dibromide analogues have approximately the same Ni–C at 2.479 Å and 2.482 Å, within the reported error. For the bis(solvento) complexes, on the other hand, the Me complex bears a shorter Ni–C distance than the H complex, at 2.416 Å vs. 2.453 Å. This suggests that the ^{Me}N3CH complexes may exhibit stronger agostic interactions than the other ^RN3CH analogues, and also may bear a stronger stabilization in the organometallic Ni complex, via donation from the axial N donors. Together, these observations indicate that ^{Me}N3CH may be best poised to undergo C–H activation reactivity.

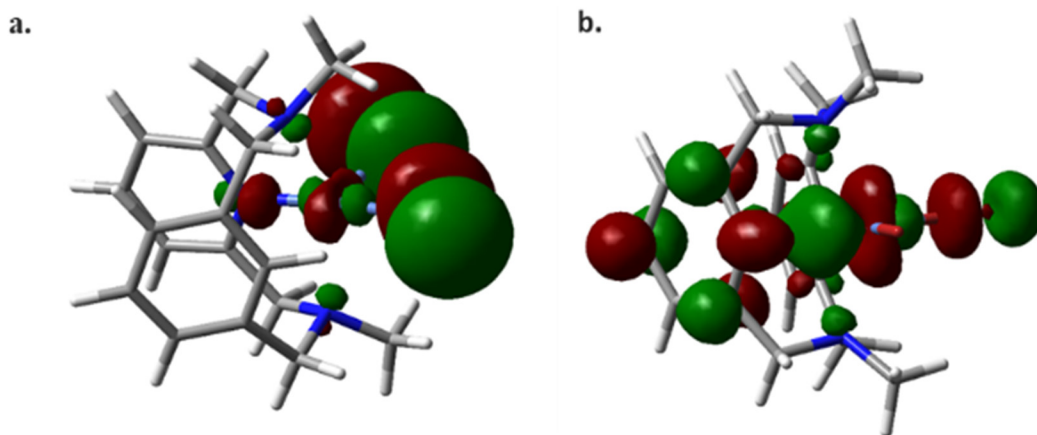


Figure 3. Frontier molecular orbitals (a. HOMO; b. LUMO, shown as 0.05 isodensity surfaces) obtained by DFT (B3LYP/def2-TZVPP), with the LUMO showing the interaction of C–H bonds with the metal d-orbital.

Catalytic trifluoroethoxylation reactivity

To investigate the catalytic properties of the complexes, we initially examined complexes **2**, **5**, and **6b** in the trifluoroalkoxylation reaction with trifluoroethanol. The Ribas group has recently reported the trifluoro- and difluoroethoxylation of Ni(II) complexes supported by the ^HN3CH ligand.¹⁸ Building on their findings, we sought to assess the steric effects of Ni complexes with the ^{Me}N3CH ligand by evaluating the reaction time and yield. The ethoxylation reaction was carried out under air in the presence of trifluoroethanol, FcPF₆ as the oxidant, and Cs₂CO₃ as the base

(Table 1). Among the complexes tested, **7** demonstrated catalytic conversion to the ethoxylated product in 49% yield which showed similar a turnover number (~ 2) as that reported by the Ribas group. Only stoichiometric amounts of the product were obtained when **2** or **6b** was used as the catalyst, suggesting that regeneration of the active catalyst during the catalytic cycle was impeded. Superstoichiometric generation of the product obtained with **5**, on the other hand, indicates that ethoxylation from a formally Ni(III) center may be more operative compared to a Ni(II) center as in **2**.

Table 1. Catalytic trifluoroethoxylation of $^{\text{Me}}\text{N}_3\text{CH}$ by Ni complexes.

Entry	Complex	mol %	yield (%)	conversion (%)
1	2	30	24	38
2	5	30	33	45
3	6b	100	12	46
4	6b	30	25	31
5	7	30	49	>99
6	7	20	27	29

Stoichiometric C-C bond formation reactivity

We then investigated C–C bond formation using the Ni complexes in various oxidation states. The $^{\text{Me}}\text{N}_3\text{CH}$ system supports Ni complexes in different oxidation states, and higher oxidation states of Ni are considered more effective in suppressing β -hydride elimination (β HE) – a common side reaction in C–C cross-coupling employing alkyl substrates. We performed stoichiometric C–C bond formation reactions using the complexes **1** and **3** with an octyl Grignard reagent. The reaction likely leads to the in-situ formation of Ni-alkyl complexes, creating a favorable environment for C–C coupling or β HE at the Ni center. Acidic workup of the reaction mixture results in the release of C–C coupled products, the β -hydride eliminated product octene, the protodemetalated product octane, and thus allowing us to measure the ratio of C–C coupling to β HE. The Ni(II) complex **1** yields 9% of the C–C coupled product, while octene was produced in 34% yield. In contrast, the Ni(III) analog, complex **3**, more effectively suppresses β HE and produces a greater yield of the C–C coupled product than **1**. Specifically, the yield of $\text{N}_3\text{C–C}_8$ was 29%, while octene was produced in 29% yield. Additionally, the homocoupled product of octyl Grignard, hexadecane, was observed at a 4% yield with **3**, none was observed for the Ni(II) complex **1**. The outcome is likely a result of a monocationic Ni(II) center being unable to bear an additional two anionic ligands, which is

supported by a reaction using dicationic **7** (see Table S2), which yields only 1% of the ^{Me}N3C–octyl coupled product but 9% hexadecane. The ratio of βHE to C–C coupling for the Ni(II) complex was 3.6 (1/0.28), whereas the Ni(III) complex showed a much lower ratio of 0.9 (1/1.13). Overall, the Ni(III) complex more efficiently suppressed βHE and yielded the C–C coupled product at a higher efficiency compared to its Ni(II) analogue. Interestingly, complex **3** reports a higher yield of both the ^{Me}N3C–octyl and hexadecane coupled products compared to the dichloride derivative **4**, even though one might expect more facile transmetallation with two chloride ligands. In this case, the loosely bound acetonitrile ligand may play a role in facilitating transmetallation and reductive elimination, as has been observed recently.¹⁹

Table 2. C-C bond coupling reaction of Ni(II) (**1**) and Ni(III) (**3**) complexes with octyl Grignard reagent.

Complex	1. octyl-MgCl (1.2 equiv) THF, RT, 1-2 h		2. H ⁺				
	Octenes	Octane	Hexadecane	N3C-octyl	N3CH	N3C-octyl/octene	C-C/octene
1	34	28	0	9	50	0.28	0.28
3	29	24	4	29	64	0.99	1.13

In summary, herein we report the synthesis and characterization of a series of Ni(II) and Ni(III) complexes bearing the ^{Me}N3C[−] or ^{Me}N3CH pyridinophane ligand framework. Analysis of the solid-state structures demonstrates the effect of the axial substituent on metrical parameters, particularly the proximity of the Ni center to the C_{sp2}–H bond to support an agnostic interaction. Catalytic ethoxylation of the ligand is observed using 2,2,2-trifluoroethanol, demonstrating its competence for C–O bond formation compared to the previously reported ^HN3CH variant. Reductive elimination studies from activated Ni(II) and Ni(III) centers shows the relevance for Ni(III) in suppressing βHE from alkyl coupling partners in favor of productive cross-coupling routes. Overall, this study sheds light on organometallic Ni catalyst design from the ligand and metal perspective for future development of Ni C–H bond catalysts.

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Notes

The authors declare no competing financial interest.

DATA AVAILABILITY

Crystallographic data for the structures reported in this article have been deposited at the Cambridge Crystallographic Data Centre, under CCDC deposition numbers 2390120-2390125.

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