

A Simple and Efficient Procedure for Knoevenagel Reaction Promoted by Imidazolium-based Ionic Liquids

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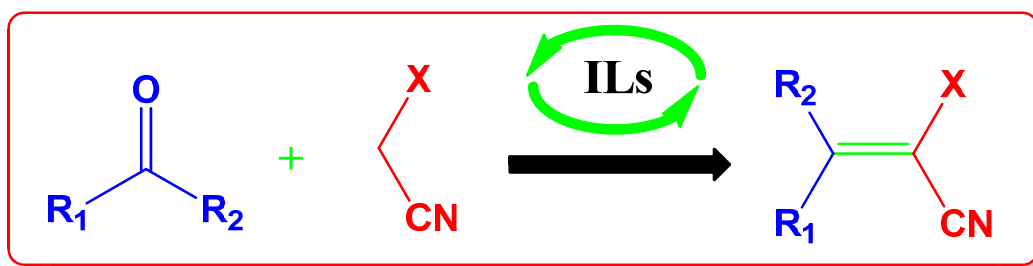
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Abstract

Various room temperature ionic liquids (RTILs), notably, 1-methoxyethyl-3-methylimidazolium trifluoroacetate $[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$, have been used to promote the Knoevenagel condensation to afford substituted olefins. All reactions proceeded effectively in the absence of any other catalysts or co-solvents with good to excellent yields. This method is simple and applicable to reactions involving a wide range of aldehydes and ketones with methylene compounds. The ionic liquid can be recycled without noticeable reduction of its catalytic activity. A plausible reaction mechanism is proposed.

Graphical Abstract



Imidazolium-based Ionic Liquids have been used to promote the Knoevenagel condensation to afford substituted olefins in good to excellent yields

Keywords: carbonyl compounds; catalysis; ionic liquids; Knoevenagel condensation; olefins.

1. Introduction

The Knoevenagel condensation reaction is a widely employed method for carbon-carbon bond formation in organic reactions [1]. Traditionally the reaction is performed in organic solvents under homogenous conditions and in the presence of weak bases, such as ethylenediamine, pyridine, piperidine or an amino acid such as glycine, β -alanine and L-proline [1]. Recently a wide range of heterogeneous catalysts has also been used for the reaction such as natural phosphate [2], metal oxides [3], modified silica [4], calcined hydrotalcites [5], alkali metals-exchanged zeolites [6], aluminophosphates oxynitrides [7], alkaline earth carbonates [8] and sulphated ZrO₂ [9]. In most cases, a combination of a catalyst and an organic solvent is used.

Recently, room temperature ionic liquids (RTILs) have gained wide popularity for their increasing applications in the areas of synthetic and biological chemistry. This is because they possess a number of interesting properties, especially their lack of vapor pressure, low flammability, a widely accessible temperature range and ease of reuse [10-14]. They are therefore considered to be environmentally friendly reaction media, which has generated an increasing interest in the application of RTILs for the Knoevenagel condensation reactions. For example, Salunkhe et al. have reported the Knoevenagel reactions in Lewis acidic ionic liquids 1-butyl-3-methylimidazolium chloroaluminate and 1-butylpyridinium chloroaluminate [15]. Base-catalysed Knoevenagel condensation in various ionic liquids has been documented, such as with proline [16, 17] and glycine [18-20]. In addition, Khan et al. have described the Knoevenagel reaction in ionic liquids catalysed by hydrotalcite [21]. Shen and co-workers have prepared a functionalized imidazolium cation-based ionic liquid immobilized on to silica gel, resulting in a solvent free system for the Knoevenagel reaction [22]. A guanidium ionic liquid [23] and 2-pyrrolidinecarboxylic acid ionic liquid [24] have been synthesized and used as catalysts for the Knoevenagel reaction. The Knoevenagel reaction has also been performed in water catalysed by ionic liquids based on 1,4-diazobicyclo[2.2.2]octane [25, 26]. Furthermore, use of dual functional ionic liquids for the condensation has been communicated by various research groups [27, 28]. We have also previously reported the Knoevenagel reaction in various ionic liquids [29-31].

However it is worthwhile to note that the aforementioned approaches have various drawbacks, including the necessity to carry out procedures in an inert atmosphere [32], harsh reaction conditions [22], long reaction times and low yields of products [17, 19]. Some approaches

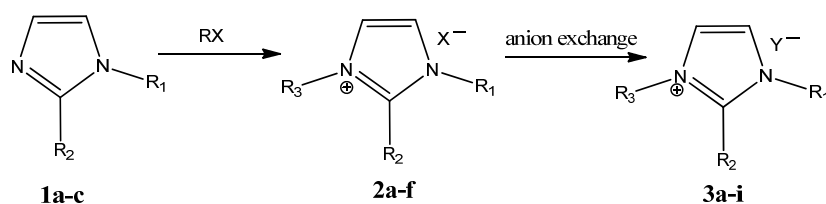
need to use catalysts in addition to ionic liquids [17-20, 31]. Moreover, in the most reported approaches, no ketones are involved since sterically hindered ketones are less reactive reagents than aldehydes in the Knoevenagel reaction [17-24, 26-30, 32-37].

In our continued efforts to use ionic liquids as environmentally friendly reaction media [29-31, 38], we have accomplished the use of various room temperature ionic liquids (RTILs), notably, 1-methoxyethyl-3-methylimidazolium trifluoroacetate $[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$, as efficient recyclable catalysts and reaction mediums for the Knoevenagel condensation. The reactions not only involve aldehydes but also ketones to afford substituted olefins in good to excellent yields under mild reaction conditions. The ionic liquid $[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$ can be recycled for up to five times with a minimal reduction in its catalytic activity. A plausible reaction mechanism is also proposed.

2. Results and Discussion

Ionic liquids **3a-h** (Table 1) were synthesised from imidazole derivatives **1a-1c** through commonly used quaternization reaction [39, 40] followed by anion metathesis [41]. The imidazole derivatives **1a-c** were reacted with appropriate halides to afford halide salts **2a-e** under anhydrous conditions. Treatment of these halide salts with an appropriate metal salt or an acid in various solvents afforded the required ionic liquids **3a-h** in good to excellent yields. The specific methods for the synthesis of these ionic liquids are described in the Experimental Section.

Table 1. Synthesis of ionic liquids



Compound No	R ₁	R ₂	R ₃	X ⁻	Y ⁻
1a	CH ₃	H			
1b	CH ₃	CH ₃			
1c	n-Bu	H			
2a	CH ₃	H	n-Bu	Cl	
2b	CH ₃	CH ₃	n-Bu	Cl	
2c	CH ₃	H	MeOMeOEt	Br	
2d	CH ₃	H	MeOEt	Cl	
2e	n-Bu	H	MeOEt	Cl	
3a	CH ₃	H	n-Bu		BF ₄
3b	CH ₃	H	n-Bu		PF ₆
3c	CH ₃	H	n-Bu		CF ₃ COO
3d	CH ₃	CH ₃	n-Bu		CF ₃ COO
3e	CH ₃	H	MeOMeOEt		CF ₃ COO
3f	CH ₃	H	MeOEt		CF ₃ COO
3g	CH ₃	H	MeOEt		CF ₃ SO ₃
3h	n-Bu	H	MeOEt		CF ₃ SO ₃

3a: 1-Butyl-3-methylimidazolium tetrafluoroborate {[BMIM]⁺[BF₄]⁻}
 3b: 1-Butyl-3-methylimidazolium hexafluorophosphate {[BMIM]⁺[PF₆]⁻}
 3c: 1-Butyl-3-methylimidazolium trifluoroacetate {[BMIM]⁺[CF₃COO]⁻}
 3d: 1-Butyl-2,3-dimethylimidazolium trifluoroacetate {[BMMIM]⁺[CF₃COO]⁻}
 3e: 1-Methoxymethoxyethyl-3-methylimidazolium trifluoroacetate {[MeOMeOEtMIM]⁺[CF₃COO]⁻}
 3f: 1-Methoxyethyl-3-methylimidazolium trifluoroacetate {[MeOEtMIM]⁺[CF₃COO]⁻}
 3g: 1-Methoxyethyl-3-methylimidazolium trifluoromethanesulfonate {[MeOEtMIM]⁺[CF₃SO₃]⁻}
 3h: 1-Methoxyethyl-3-butylimidazolium trifluoromethanesulfonate {[MeOEtBuIM]⁺[CF₃SO₃]⁻}

A number of solvents have been reported for the quaternization reactions [42, 43]. In general, it will help to drive the reaction to completion and help the product isolation if a resulting ionic liquid is immiscible with the solvent. In order to identify suitable solvents for the quaternization reaction various solvents were investigated for the reaction between 1-methylimidazole and 1-chlorobutane. The results are shown in Figure 1.

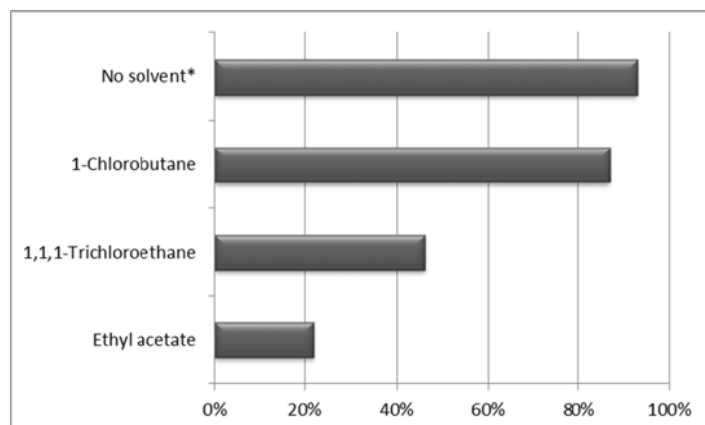


Figure 1. Comparison of various solvents for the preparation of $[\text{BMIM}]^+\text{Cl}^-$ (**2a**) (*equal molar of the starting materials without addition of other solvents).

The above data show that when chlorobutane was used as a solvent the yield was higher than that in other solvents. Additionally, the quaternization reaction with no solvent also proceeded well with an excellent yield. Consequently all quaternization reactions were performed in an appropriate halide reagent without the addition of other solvents. These halides salts (**2a-e**) were characterized with ^1H NMR, ^{13}C NMR, IR and MS before being used for the preparation of ionic liquids (**3a-h**) through anion exchange reactions (Table 1).

It is highly desirable to remove any trace of 1-alkylimidazoles from the quaternization products as they could play an unfavorable role in some applications of ionic liquids. For example, electrophilic catalysts can coordinate to 1-alkylimidazoles in an irreversible manner, resulting in deactivation of the catalysts [44]. One commonly used method to remove alkylimidazoles is by heating *in vacuo* in a rotary evaporator. But this method is not always efficient as revealed by NMR analysis of $[\text{BMIM}]^+\text{Cl}^-$ obtained after drying at $80\text{ }^\circ\text{C}$ for 8 h *in vacuo* (Fig. 2a), showing the presence of residue 1-methylimidazole. Therefore, an alternative approach was used. This was to dissolve the halide salt in dry acetonitrile at $50\text{ }^\circ\text{C}$ and add the solution to a mixture of dry acetonitrile/diethyl ether cooled in an ice bath under stirring. ^1H NMR spectrum (Fig. 2b) of $[\text{BMIM}]^+\text{Cl}^-$ obtained using this method confirmed that there was no residual 1-methylimidazole. Subsequently, all solid halide salts were purified by recrystallization before anion exchange reactions.

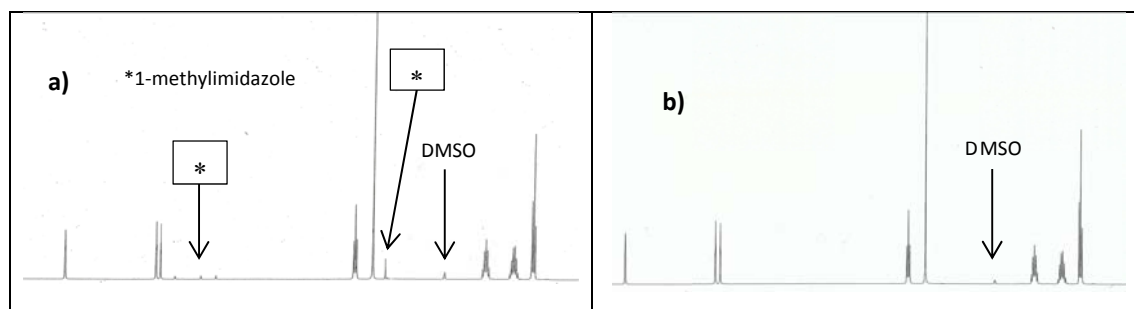


Figure 2. ^1H NMR spectrum of 1-butyl-3-methylimidazolium chloride, a) purified by heating under high vacuum; b) by recrystallization.

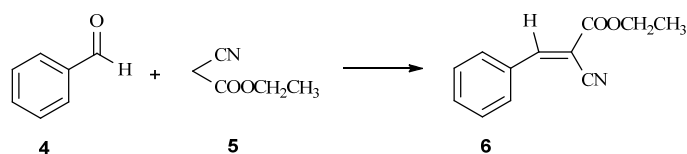
$[\text{BMIM}]^+[\text{BF}_4]^-$ (**3a**) was prepared as described [45] with some modifications. $[\text{BMIM}]^+\text{Cl}^-$ (**2a**) was dissolved in water and tetrafluoroboric acid (1 equivalent) was added slowly under stirring. The mixture was then stirred for 24 hours at room temperature. The resulting ionic liquid was extracted with chilled dichloromethane. The extract was washed with chilled water until the aqueous phase was pH neutral and free of chloride ion (AgNO_3). It was not efficient to extract water miscible ionic liquids $[\text{BMIM}]^+[\text{BF}_4]^-$ from the aqueous solution so large volume of extraction solvent was used initially. However, it was noticed later the extraction efficiency improved significantly when carried out at a low temperature. Consequently, the reaction mixture and CH_2Cl_2 were chilled in an ice bath before the extraction. $[\text{BMIM}]^+[\text{PF}_6]^-$ (**3b**) is a water-immiscible ionic liquid. Its preparation was carried out in aqueous solution by the reaction between KPF_6 with $[\text{BMIM}]^+\text{Cl}^-$ (**2a**) at room temperature [46]. A biphasic layer was formed when the reaction was complete. The aqueous layer was separated off and the organic layer was washed with water to give a pure product after being dried at $60\text{ }^\circ\text{C}$ *in vacuo*.

Ionic liquids **3c-i** were prepared through anion metathesis reactions of an appropriate halide salt with either CF_3COONa or $\text{CF}_3\text{SO}_3\text{Na}$ in acetonitrile or acetone [47]. Both CF_3COONa and $\text{CF}_3\text{SO}_3\text{Na}$ can be dissolved in acetonitrile or acetone. The reactions were completed with the resulting sodium halides precipitating out spontaneously. An initial attempt was made to synthesise these ionic liquids in an aqueous medium but with unsatisfactory yields, mainly due to poor extraction efficiency of these water soluble ionic liquids with organic solvents in aqueous solution. These ionic liquids were characterized with ^1H NMR, ^{13}C NMR and MS.

It has been widely reported that the C-2 hydrogen of imidazolium based ionic liquids plays a significant role in various organic reactions promoted by ionic liquids [48-51]. In order to

investigate its involvement in promoting the Knoevenagel reaction, ionic liquid **3d** was prepared, in which the C-2 hydrogen was substituted with a methyl group.

The ionic liquids **3a-h** prepared above were examined and compared to evaluate their efficiency for promoting the Knoevenagel reaction as outlined in Scheme 1. Benzaldehyde **4** and ethyl cyanoacetate **5** were mixed and one of the ionic liquids added and the mixture stirred for an appropriate time at 25 °C (Table 2). The product **6** was extracted with diethyl ether. After removal of the solvent the crude product was purified by recrystallization. The results summarised in Table 2 demonstrate that Knoevenagel condensation of benzaldehyde with ethyl cyanoacetate proceeded in all of the RTILs tested but with significantly variable efficiency. By comparing the results, it is evident that the structures of both the imidazolium moiety and its counter anion in the ionic liquids have a significant effect on their activity for promoting the Knoevenagel reaction. Particularly, a substantial reduction in the catalytic efficiency was observed when the C-2 hydrogen was replaced by a methyl group (entries 4).



Scheme 1. Reaction of benzaldehyde with ethyl cyanoacetate in various ILs.

Table 2. Knoevenagel condensation of benzaldehyde with ethyl cyanoacetate in various ionic liquids^a

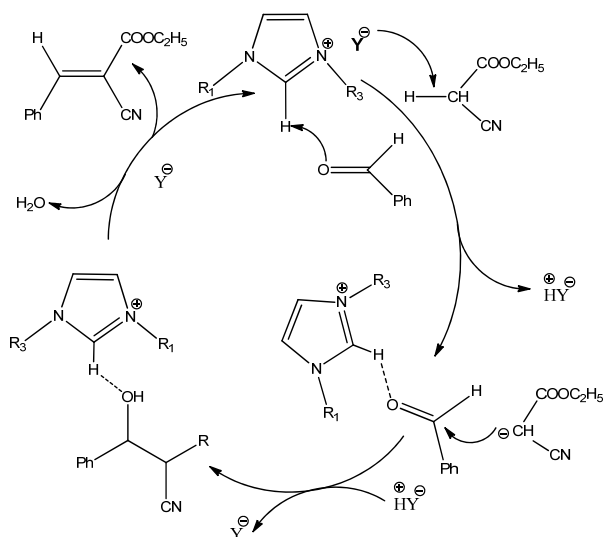
Entry	Solvent	Reaction time	Yield (%) ^b
1	[BMIM] ⁺ [BF ₄] ⁻ (3a)	24 h	85
2	[BMIM] ⁺ [PF ₆] ⁻ (3b)	22 h	63
3	[BMIM] ⁺ [CF ₃ COO] ⁻ (3c)	1 h	98
4	[BMMIM] ⁺ [CF ₃ COO] ⁻ (3d)	4 h	62
5	[MeOMeOEtMIM] ⁺ [CF ₃ COO] ⁻ (3e)	40 min	95
6	[[MeOEtMIM] ⁺ [CF ₃ COO] ⁻ (3f)	40 min	98
7	[MeOEtMIM] ⁺ [CF ₃ SO ₃] ⁻ (3g)	1 h	81
8	[MeOEtBuIM] ⁺ [CF ₃ SO ₃] ⁻ (3h)	2 h	54

^a Reaction conditions: benzaldehyde (5 mmol), ethyl cyanoacetate (5 mmol), IL (5 mmol) at 25 °C
^b Isolated yield of products.

It has been reported widely that the catalytic efficiency of ILs for various organic reactions is influenced by the structures of the cation and the counter anion. Chakraborti and co-workers reported that catalytic activity of ILs derived from 1-methyl-3-alkylimidazolium cations is affected by both cations and anions for various organic reactions [48, 50, 51]. The

involvement of the hydrogen bond donor of C-2 hydrogen in the imidazolium moiety and the hydrogen bond acceptor of the counter anion was supported by the results of various spectroscopic studies. The importance of the C-2 hydrogen of the imidazolium species was further demonstrated through a large reduction of catalytic efficiency of ILs derived from imidazolium cations having an alkyl group substituted at C-2 [48, 50, 51]. Zhu et al.[27] observed the importance of both the cations and the counter anions in N, N-dimethyl ethanolammonium-based ionic liquids. Their results demonstrated that the increase in the hydrogen bond donor ability of the cation and the hydrogen bond acceptor ability of the anion in an ionic liquid can lead to an enhancement in its catalytic activity for the Knoevenagel condensation. A possible catalytic mechanism was proposed, which involved the activation of C=O group through the formation of a hydrogen bond with the cations of the ILs and an attack by the anions of the ILs to the active methylene compounds to form a carboanion [27].

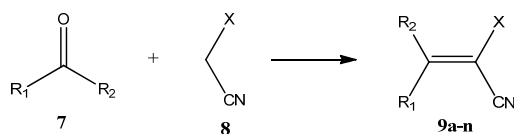
Based on the different catalytic activities observed for the ILs investigated here, particularly the importance of the involvement of C-2 hydrogen in the imidazolium moiety (entry 4) and the observations and reaction mechanism reported by others [27, 48, 50, 51] a possible mechanism is proposed and illustrated in Scheme 2. It involves an attack to the active methylene compound by the counter anion of an IL to form a carboanion. This anion then attacks the C=O group of benzaldehyde, which has been activated by the formation of a hydrogen bond with the C-2 hydrogen of the imidazolium cations, to produce an alcohol intermediate. An attack to this intermediate by the anion of the ILs produce the final dehydration product. In this reaction mechanism, the counter anions of ionic liquids play an activating role for the nucleophiles of methylene compounds to produce carboanions and the imidazolium cations act as activators for the electrophiles of carbonyl compounds by increasing the polarisation of C=O bond through the formation of a hydrogen bond with the C-2 hydrogen.



Scheme 2. Proposed mechanism for the Knoevenagel reaction.

Although all the ionic liquids tested demonstrated activity for promoting Knoevenagel reactions (Table 2), an ideal ionic liquid for promoting the Knoevenagel condensation reaction should have a strong catalytic activity, a low viscosity and be economically viable for a large scale production. According to the yields and times of the reactions (Table 2), [BMIM]⁺[CF₃COO]⁻ (**3c**), [MeOMeOEtMIM]⁺[CF₃COO]⁻ (**3e**) and [MeOEtMIM]⁺[CF₃COO]⁻ (**3f**) seemed to be better choices compared with the others. However, [MeOMeOEtMIM]⁺[CF₃COO]⁻ (**3e**) is relatively more viscous and the starting materials for its preparation are more expensive than for that of the other two ionic liquids. A further experiment was thus carried out in order to differentiate the other two ionic liquids. When the reaction of 4-methylbenzaldehyde with ethyl cyanoacetate was carried out in [BMIM]⁺[CF₃COO]⁻ (**3c**), the reaction time was 30 minutes with a yield of 74%, whereas the same reaction in [MeOEtMIM]⁺[CF₃COO]⁻ (**3f**) was 20 minutes with a yield of 89%. Considering these results, **3f** was chosen as a dual functional ionic liquid for the Knoevenagel reaction. In order to further assess its catalytic activity a wide range of reactions between aldehydes or ketones (**7**) with methylene compounds (**8**) were investigated in [MeOEtMIM]⁺[CF₃COO]⁻. The results are summarised in Table 3.

Table 3. Knoevenagel condensation of aldehydes and ketones with methylene compounds in [MeOEtMIM]⁺[CF₃COO]⁻^a



Entry	R1	R2	X	Time	Temperature	Yield (%) ^b	Product
1	Ph	H	COOEt	40 min	25 °C	98	9a
2	4-Cl-C ₆ H ₄	H	COOEt	5 min	25 °C	97	9b
3	4-Me-C ₆ H ₄	H	COOEt	20 min	25 °C	89	9c
4	4-NO ₂ -C ₆ H ₄	H	COOEt	10 min	25 °C	81	9d
5	4-HOOC-C ₆ H ₄	H	COOEt	50 min	25 °C	77	9e
6	2-Naphthyl	H	COOEt	15 min	25 °C	98	9f
7	3-Cl-C ₆ H ₄	H	COOMe	2 min	25 °C	99	9g
8	4-Me-C ₆ H ₄	H	COOMe	10 min	25 °C	91	9h
9	4-MeO-C ₆ H ₄	H	COOMe	35 min	25 °C	78	9i
10	4-HO-C ₆ H ₄	H	COOMe	40 min	25 °C	83	9j
11	-(CH ₂) ₄ -		COOMe	48 h	25 °C	85	9k
12	-(CH ₂) ₅ -		COOMe	16 h	25 °C	56	9l
13	Ph	CH ₃	COOMe	48 h	50 °C	55	9m
14	Ph	CH ₃	COOEt	48 h	50 °C	47	9n
15	n-C ₃ H ₇	H	COOEt	2 h	25 °C	82	9o
16	2-Furyl	H	COOEt	80 min	25 °C	85	9p

^a Reaction conditions: aldehyde or ketone (5 mmol), active methylene compound (5 mmol). Ionic liquid (5 mmol).
^b Isolated yield of products.

As illustrated in Table 3, various aldehydes, including aromatic aldehydes (entries 1-10) with electron-donating groups (methyl, methoxyl and hydroxyl) or electron-withdrawing groups (chloro, nitro, carboxyl), aliphatic aldehyde (entry 15) and heterocyclic aldehyde (entry 16), reacted speedily with the active methylene compounds at 25 °C with satisfactory yields. In the case of ketones, longer reaction times were needed (entries 11-12). As ketones becomes more steric-hindrance both longer reaction times and higher temperature were applied to overcome the increased steric hindrance of the substrates (entries 13-14). All products were characterized by ¹H NMR and ¹³C NMR. These reactions were usually stereoselective with the predominate products being the E-configuration, as reported previously in the literature [31, 52]. However, two isomers (E & Z) were obtained when acetophenone was reacted with methylene compounds but with E-geometry as the main product as revealed by ¹H NMR and ¹³C NMR.

One of the main advantages of ionic liquids is their ease of reuse. In order to determine the

performance of the recycled ionic liquid, the ionic liquid was recovered from the reaction mixture and reaction of benzaldehyde with ethyl cyanoacetate was carried out in the recycled $[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$ (**3f**). The results summarized in Table 4 demonstrate **3f** was still effective after 5 cycles with a minimal reduction in its activity.

Table 4. Knoevenagel condensation of benzaldehyde with ethyl cyanoacetate

Ionic liquid	Yield (%)
$[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$ (Fresh)	98
$[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$ (cycle1)	94
$[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$ (cycle2)	91
$[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$ (cycle3)	88
$[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$ (cycle4)	85
$[\text{MeOEtMIM}]^+[\text{CF}_3\text{COO}]^-$ (cycle5)	86

3. Conclusion

In summary, we have developed a simple and efficient procedure for Knoevenagel condensation reactions in various ionic liquids. The results obtained with 1-methoxyethyl-3-methylimidazolium trifluoroacetate (**3f**) show that the approach is applicable to a large number of substrates, including not only aldehydes but also ketones, to afford substituted olefins in good to excellent yields in short reaction times. The approach also has the benefit of mild reaction temperatures and easy preparation procedures. In contrast to other reported procedures in which both ionic liquids and additional catalysts are needed [17-20, 31], the ionic liquids reported here act as both a catalyst and a solvent for the reactions and can be recycled and reused for 5 times without obvious loss of its catalytic activity. Furthermore, it has been demonstrated that the catalytic activity of the ionic liquids for the Knoevenagel reactions is likely linked to their ability to form hydrogen bonds. We believe this simple and green method will be a useful alternative to the existing Knoevenagel reaction procedures.

4. Experimental section

4.1. General

Chemicals and solvents were used as received without purification. Melting points were determined on a Reichert-Jung Galen micro melting point apparatus and were uncorrected. ^1H (250 MHz) and ^{13}C (62.9 MHz) NMR spectra were recorded on a Bruker AC-250 spectrometer with TMS as an internal standard. Infrared

spectra were recorded on a Mattson 3000 FT-IR spectrophotometer using KBr discs for solids and thin films for liquids. Mass spectra were recorded on a Waters LCT Premier mass spectrometer by electron spray ionisation or electron impact ionisation. TLC was carried out on pre-coated Merck 60 F254 plates and visualized using UV (254 nm and 360 nm).

4.2. Synthesis of ionic liquids 3a-h

4.2.1. 1-Butyl-3-methylimidazolium tetrafluoroborate [BMIM]⁺[BF₄]⁻ (**3a**) [45]

1-butyl-3-methylimidazolium chloride [BMIM]⁺Cl⁻ (**2a**) [46]: 1-Methylimidazole (**1a**, 80 mL, 1 mol) was added dropwise to 1-chlorobutane (200 mL, 1.9 mol). The mixture was stirred vigorously and refluxed at 80 °C for 24 h. When the reaction was complete, the excess 1-chlorobutane was decanted and the crude ionic liquid was washed with chlorobutane (2 × 20 mL). The trace of remaining 1-chlorobutane was removed with rotary evaporation at 60 °C for 30 min, followed by at 80 °C for 4 h under reduced pressure. The crude product was further purified by recrystallization (acetonitrile/ether) and dried under vacuum to afford **2a** as a white solid (151.9 g, 87%). ¹H NMR (250 MHz, DMSO-d₆) δ: 9.57 (s, 1H, NCHN), 7.91 (t, *J* = 1.90 Hz, 1H, NCHCHN), 7.83 (t, *J* = 1.90 Hz, 1H, NHCCHN), 4.21 (t, *J* = 6.95 Hz, 2H, NCH₂), 3.88 (s, 3H, N-CH₃), 1.76 (quint, *J* = 7.58 Hz, 2H, NCH₂CH₂), 1.23 (sext, *J* = 7.58 Hz, 2H, NCH₂CH₂CH₂), 0.88 (t, *J* = 7.58 Hz, 3H, N-CH₂CH₂CH₂-CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 136.66 (NCHN), 123.47 (NCHCHN), 122.19 (NCHCHN), 48.27 (NCH₂CH₂CH₂CH₃), 35.60 (NCH₃), 31.31 (NCH₂CH₂CH₂CH₃), 18.67 (NCH₂CH₂CH₂CH₃), 13.21 (NCH₂CH₂CH₂CH₃). MS (ESI⁺) (*m/z*) = 139 {[BMIM]⁺}. IR (KBr) cm⁻¹: 3404, 3096, 2954, 1570, 1463, 1170, 756, 623.

[BMIM]⁺Cl⁻ (76 g, 0.45 mol) was dissolved in water (30 mL). Tetrafluoroboric acid (63 mL, 0.5 mol) was added dropwise to the above solution under stirring in an ice bath. The mixture was stirred at room temperature for 24 h. The resulting [BMIM]⁺[BF₄]⁻ was chilled in an ice bath and extracted with chilled dichloromethane (6 × 10 mL). The dichloromethane fraction was washed with chilled water until the aqueous fraction pH became neutral and free of chloride (AgNO₃). The dichloromethane was removed by rotary evaporation and the crude product was mixed with charcoal and stirred for 24 h at room temperature. The charcoal was filtered off and the product was dried by heating under high vacuum. [BMIM]⁺[BF₄]⁻ (**3a**) was obtained as a colorless viscous liquid (68g, 69%). ¹H NMR (250 MHz, DMSO-d₆) δ: 9.07 (s, 1H, NCHN), 7.75 (t, *J* = 1.90 Hz, 1H, NCHCHN), 7.68 (t, *J* = 1.90 Hz, 1H, NCHCHN), 4.17 (t, *J* = 6.96 Hz, 2H, NCH₂), 3.86 (s, 3H, NCH₃), 1.78 (quint, *J* = 7.74 Hz, 2H, NCH₂CH₂), 1.27 (sext, *J* = 7.60 Hz, 2H, NCH₂CH₂CH₂), 0.91 (t, *J* = 7.74 Hz, 3H, NCH₂CH₂CH₂CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 136.34 (NCHN), 123.52 (NCHCHN), 122.18 (NCHCHN), 48.47 (NCH₂CH₂CH₂CH₃), 35.62 (NCH₃), 31.29 (NCH₂CH₂CH₂CH₃), 18.79 (NCH₂CH₂CH₂CH₃), 13.16 (NCH₂CH₂CH₂CH₃). MS (ESI⁺) (*m/z*) = 139 {[BMIM]⁺}. IR (neat) cm⁻¹: 3636, 3159, 2962, 1573, 1465, 1079, 1043, 849, 751, 625.

4.2.2. 1-Butyl-3-methylimidazolium hexafluorophosphate {[BMIM]⁺[PF₆]⁻} (**3b**) [46]

This was prepared as described [46]. Yield: 21.45 g (75%). ¹H NMR (250 MHz, DMSO-d₆) δ: 9.07 (s, 1H, NCHN), 7.73 (t, *J* = 1.90 Hz, 1H, NCHCHN), 7.66 (t, *J* = 1.90 Hz, 1H, NCHCHN), 4.17 (t, *J* = 7.15 Hz, 2H,

NCH₂), 3.86 (s, 3H, NCH₃), 1.79 (quint, $J = 7.71$ Hz, 2H, NCH₂CH₂), 1.28 (sext, $J = 7.40$ Hz, 2H, NCH₂CH₂CH₂), 0.91 (t, $J = 7.22$ Hz, 3H, NCH₂CH₂CH₂CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ : 136.40 (NCHN), 123.48 (NCHCHN), 122.12 (NCHCHN), 48.49 (NCH₂CH₂CH₂CH₃), 35.57 (NCH₃), 31.26 (NCH₂CH₂CH₂CH₃), 18.69 (NCH₂CH₂CH₂CH₃), 13.09 (NCH₂CH₂CH₂CH₃). MS (ESI⁺) (m/z) = 139{[BMIM]⁺}. IR (neat) cm⁻¹: 3673, 3173, 2964, 1573, 1464, 1168, 882, 832, 745, 627.

4.2.3. 1-Butyl-3-methylimidazolium trifluoroacetate {[BMIM]⁺[CF₃COO]⁻} (3c) [42]

A solution of **2a** (20.3 g, 0.12 mol) in acetonitrile was mixed with a solution of sodium trifluoroacetate (15.82 g, 0.12 mol) in acetonitrile/acetone (1:1). The mixture was stirred for 4 h. The precipitate (NaCl) was filtered off through celite and the solvent was removed. Dichloromethane (20 mL) was added into the product and the solution was filtered off again. Dichloromethane was removed by rotary evaporation and the product was kept in high vacuum at 60 °C for 3 h. Yield: 23.18 g (79%). ¹H NMR (250 MHz, DMSO-d₆) δ : 9.35 (s, 1H, NCHN), 7.85 (t, $J = 1.90$ Hz, 1H, NCHCHN), 7.77 (t, 1H, $J = 1.90$ Hz, NCHCHN), 4.18 (t, $J = 7.58$ Hz, 2H, NCH₂), 3.87 (s, 3H, NCH₃), 1.76 (quint, $J = 7.58$ Hz, 2H, NCH₂CH₂), 1.24 (sext, $J = 7.58$ Hz, 2H, NCH₂CH₂CH₂), 0.88 (t, $J = 7.58$ Hz, 3H, CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ : 136.80 (NCHN), 123.59 (NCHCHN), 122.29 (NCHCHN), 48.42 (NCH₂CH₂CH₂CH₃), 35.58 (NCH₃), 31.38 (NCH₂CH₂CH₂CH₃), 18.72 (NCH₂CH₂CH₂CH₃), 13.13 (NCH₂CH₂CH₂CH₃). MS (ESI⁺) (m/z) = 139{[BMIM]⁺}. IR (cm⁻¹) = 3434, 3092, 2961, 1698, 1571, 1466, 1409, 1202, 1163, 1119, 825, 795, 716, 624.

4.2.4. 1-Butyl-2,3-dimethylimidazolium trifluoroacetate {[BMMIM]⁺[CF₃COO]⁻} (3d)

1-Butyl-2,3-dimethylimidazolium chloride [BMMIM]⁺Cl⁻ (**2b**): 1,2-Dimethylimidazole (**1b**, 9.6 g, 0.1 mol) was mixed with 2-chlorobutane (12 mL, 0.11 mol). The mixture was stirred at 80 °C for 24 h. The crude product was purified by recrystallisation in acetonitrile/ether and dried under reduced pressure. Yield: 12.65 g (65%). ¹H NMR (250 MHz, DMSO-d₆) δ : 7.71 (d, $J = 1.90$ Hz, 1H, NCHCHN), 7.69 (d, $J = 1.90$ Hz, 1H, NCHCHN), 4.12 (t, $J = 7.58$ Hz, 2H, NCH₂), 3.76 (s, 3H, NCH₃), 2.59 (s, 3H, CH₃), 1.69 (quint, $J = 7.58$ Hz, 2H, NCH₂CH₂), 1.28 (sext, $J = 7.58$ Hz, 2H, NCH₂CH₂CH₂), 0.91 (t, $J = 7.58$ Hz, 3H, NCH₂CH₂CH₂CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ : 144.15 (NCN), 122.25 (NCHCHN), 120.83 (NCHCHN), 47.18 (NCH₂CH₂CH₂CH₃), 34.61 (NCH₃), 31.16 (NCH₂CH₂CH₂CH₃), 18.81 (NCH₂CH₂CH₂CH₃), 13.36 (CCH₃), 9.13 (NCH₂CH₂CH₂CH₃). MS (ESI⁺) (m/z) = 153{[BMMIM]⁺}. IR (KBr) cm⁻¹: 3402, 3068, 2967, 1591, 1543, 1472, 1253, 1138, 752.

1-Butyl-2,3-dimethylimidazolium trifluoroacetate (**3d**) was prepared by following the same procedure as described for the synthesis of **3c**. Yield: 7.12 g (94%). ¹H NMR (250 MHz, DMSO-d₆) δ : 7.68 (d, $J = 1.9$ Hz, 1H, NCHCHN), 7.66 (d, $J = 1.9$ Hz, 1H, NCHCH-N), 4.11 (t, $J = 6.95$ Hz, 2H, NCH₂), 3.75 (s, 3H, NCH₃), 2.58 (s, 3H, CH₃), 1.68 (quint, $J = 7.58$ Hz, 2H, NCH₂CH₂), 1.28 (sext, $J = 7.58$ Hz, 2H, NCH₂CH₂CH₂), 0.90 (t, $J = 6.95$ Hz, 3H, CH₂CH₂CH₂CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ : 144.30 (NCN), 122.29 (NCHCHN), 120.83 (NCHCHN), 47.21 (NCH₂CH₂CH₂CH₃), 34.58 (NCH₃), 31.16 (NCH₂CH₂CH₂CH₃), 18.83 (NCH₂CH₂CH₂CH₃), 13.35 (CCH₃), 9.05 (NCH₂CH₂CH₂CH₃). HRMS calcd for C₉H₁₇N₂ {[BMMIM]⁺} 153.1392, found 153.1391. IR (KBr) cm⁻¹: 3443, 2967, 1697, 1540, 1464, 1423, 1203, 1172, 1123, 826, 799, 719.

4.2.5. 1-Methoxymethoxyethyl-3-methylimidazolium trifluoroacetate {[MeOMeOEtMIM]⁺[CF₃COO]⁻} (3e)

1-Methoxymethoxyethyl-3-methylimidazolium bromide {[MeOMeOEtMIM]⁺Br⁻} (**2c**): 1-Methylimidazole (**1a**, 2.4 g, 0.03 mol) was treated with 1-methoxymethoxyethyl bromide (5 g, 0.03 mol). The mixture was stirred at 50 °C for 20 h. The resulting product was washed with ethyl acetate (2 × 5 mL) and dried in a high vacuum for 2 h at 80 °C to afford **2c**. Yield: 7.2 g (97%). ¹H NMR (250 MHz, DMSO-d₆) δ: 9.31 (s, 1H, NCHN), 7.88 (t, *J* = 1.90 Hz, 1H, NCHCHN), 7.85 (t, *J* = 1.90 Hz, 1H, NCHCHN), 4.58 (s, 2H, OCH₂OCH₃), 4.42 (t, *J* = 5.05 Hz, 2H, NCH₂CH₂O), 3.89 (s, 3H, NCH₃), 3.82 (t, *J* = 5.05 Hz, 2H, NCH₂CH₂O), 3.16 (s, 3H, OCH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 136.93 (NCHN), 123.49 (NCHCHN), 122.66 (NCHCHN), 95.25 (OCH₂O), 69.61 (NCH₂CH₂O), 57.95 (OCH₃), 48.59 (NCH₂CH₂O), 35.62 (NCH₃). MS (ESI⁺) (m/z) = 171 {[MeOMeOEtMIM]⁺}. IR (neat) cm⁻¹: 3433, 2961, 1562, 1466, 1159, 1115, 1040, 909, 760. For the synthesis of **3e**, a similar method was followed as described for the synthesis of **3c**. [MeOMeOEtMIM]⁺Br⁻ (7 g, 0.03 mol) was reacted with CF₃COONa (4.80 g, 0.03 mol) to afford **3e** (7.23 g, 81%). ¹H NMR (250 MHz, DMSO-d₆) δ: 9.25 (s, 1H, NCHN), 7.82 (t, *J* = 1.90 Hz, 1H, NCHCHN), 7.76 (t, *J* = 1.90 Hz, 1H, NCHCHN), 4.55 (s, 2H, OCH₂OCH₃), 4.40 (t, *J* = 5.05 Hz, 2H, NCH₂CH₂O), 3.88 (s, 3H, NCH₃), 3.81 (t, *J* = 5.05 Hz, 2H, NCH₂CH₂O), 3.16 (s, 3H, OCH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 137.09 (NCHN), 123.47 (NCHCHN), 122.55 (NCHCHN), 95.50 (OCH₂O), 64.93 (NCH₂CH₂O), 54.72 (OCH₃), 48.87 (NCH₂CH₂O), 35.68 (NCH₃). HRMS calcd for C₈H₁₅N₂O₂ {[MeOMeOEtMIM]⁺} 171.1134, found 171.1138. IR (neat) cm⁻¹: 3429, 3087, 2953, 1694, 1429, 1173, 1047, 831.

4.2.6. 1-Methoxyethyl-3-methylimidazolium trifluoroacetate {[MeOEtMIM]⁺[CF₃COO]⁻} (3f) [47]

1-Methoxyethyl-3-methylimidazolium chloride {[MeOEtMIM]⁺Cl⁻} (**2d**) [41]: 1-methylimidazole (**1a**, 8 mL, 0.1 mol) was added dropwise to 2-chloroethyl methyl ether (25 mL, 0.27 mol). The mixture was stirred and refluxed at 80 °C for 50 h. The upper layer, 2-chloroethyl methyl ether, was decanted. The resulting ionic liquid was washed with 2-chloroethyl methyl ether (2 × 5 mL) and evaporated. The crude product was dissolved in acetonitrile and recrystallized in acetonitrile/ether and dried under vacuum overnight. Yield: 17.59 g (99%). ¹H NMR (250 MHz, DMSO-d₆) δ: 9.57 (s, 1H, NCHN), 7.91 (t, *J* = 1.90 Hz, 1H, NCHCHN), 7.85 (t, *J* = 1.90 Hz, 1H, NCHCHN), 4.42 (t, *J* = 5.05 Hz, 2H, NCH₂CH₂O), 3.90 (s, 3H, NCH₃), 3.71 (t, *J* = 5.05 Hz, 2H, NCH₂CH₂O), 3.24 (s, 3H, OCH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 136.93 (NCHN), 123.33 (NCHCHN), 122.51 (NCHCHN), 69.52 (NCH₂CH₂O), 57.93 (OCH₃), 48.36 (NCH₂CH₂O), 35.61 (NCH₃). MS (ESI⁺) (m/z) = 141 {[MeOEtMIM]⁺}. IR (KBr) cm⁻¹: 3391, 3072, 1570, 1449, 1177, 1117, 1013, 832.

For the synthesis of **3f**, a similar method was followed as described for the synthesis of **3c**. A solution of **2d** (36.06 g, 0.2 mol) in acetonitrile was added to a solution of sodium trifluoroacetate (27.76 g, 0.2 mol) in acetone to afford **3f** (43.07 g, 83%). ¹H NMR (250 MHz, DMSO-d₆) δ: 9.22 (s, 1H, NCHN), 7.78 (t, *J* = 1.90 Hz, 1H, NCHCHN), 7.74 (t, *J* = 1.90 Hz, 1H, NCHCHN), 4.37 (t, *J* = 5.05 Hz, 2H, NCH₂CH₂O), 3.88 (s, 3H, NCH₃), 3.69 (t, *J* = 5.05 Hz, 2H, NCH₂CH₂O), 3.26 (s, 3H, OCH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 136.87 (NCHN), 123.42 (NCHCHN), 122.58 (NCHCHN), 69.54 (NCH₂CH₂O), 57.96 (OCH₃), 48.54 (NCH₂CH₂O), 35.65 (NCH₃). MS (ESI⁺) (m/z) = 141 {[MeOEtMIM]⁺}. IR (neat) cm⁻¹: 3436, 3086, 1695, 1682, 1573, 1454, 1191, 1123, 823, 800.

4.2.7. 1-Methoxyethyl-3-methylimidazolium trifluoromethanesulfonate [MeOEtMIM]⁺[CF₃SO₃]⁻ (**3g**) [42]

A similar method was followed as described for the synthesis of **3c**. [MeOEtMIM]⁺Cl⁻ (**2d**, 6.93 g, 0.04 mol) was reacted with CF₃SO₃Na (6.75 g, 0.04 mol) to give the product **3h**. Yield: 8.92 g (79%). ¹H NMR (250 MHz, DMSO-d₆) δ: 9.10 (s, 1H, NCHN), 7.75 (t, *J* = 1.90 Hz, 1H, NCHCHN), 7.70 (t, *J* = 1.90 Hz, 1H, NCHCHN), 4.35 (t, *J* = 5.21 Hz, 2H, NCH₂), 3.86 (s, 3H, NCH₃), 3.67 (t, *J* = 5.05 Hz, 2H, CH₂O), 3.26 (s, 3H, OCH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 136.76 (NCHN), 123.41 (NCHCHN), 122.56 (NCHCHN), 69.52 (NCH₂CH₂O), 57.98 (OCH₃), 48.59 (NCH₂CH₂O), 35.68 (NCH₃). MS (ESI⁺) (*m/z*) = 141 {[MeOEtMIM]⁺}. IR (neat) cm⁻¹: 3565, 3112, 2943, 2899, 1572, 1452, 1266, 1163, 1030, 835.

4.2.8. 1-Methoxyethyl-3-butylimidazolium trifluoromethanesulfonate {[MeOEtBuIM]⁺[CF₃SO₃]⁻} (**3h**)

1-methoxyethyl-3-butylimidazolium chloride (**2e**, 6.7 g, 0.03 mol), prepared as described [42], was dissolved in acetone. CF₃SO₃Na (5.3 g, 0.03 mol) dissolved in acetone was added to the above solution. Following that a similar procedure was followed as for the synthesis of **3c**. Yield: 6.75 g (66%). ¹H NMR (250 MHz, DMSO-d₆) δ: 9.14 (t, 1H, NCHN), 7.78 (t, *J* = 1.74 Hz, 1H, NCHCHN), 7.74 (t, *J* = 1.74 Hz, 1H, NCHCHN), 4.37 (t, *J* = 4.74 Hz, 2H, NCH₂CH₂O), 4.20 (t, *J* = 7.27 Hz, 2H, NCH₂CH₂CH₂), 3.70 (t, *J* = 4.74 Hz, 2H, CH₂O), 3.26 (s, 3H, OCH₃), 1.78 (quint, *J* = 7.27 Hz, 2H, NCH₂CH₂CH₂CH₃), 1.25 (sext, *J* = 7.58 Hz, 2H, NCH₂CH₂CH₂CH₃), 0.89 (t, *J* = 7.58 Hz, 3H, NCH₂CH₂CH₂CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 136.18 (NCHN), 122.76 (NCHCHN), 122.22 (NCHCHN), 69.50 (NCH₂CH₂O), 57.94 (OCH₃), 48.74 (NCH₂CH₂O), 48.65 (NCH₂CH₂CH₂CH₃), 31.38 (NCH₂CH₂CH₂CH₃), 18.74 (NCH₂CH₂CH₂CH₃), 13.06 (NCH₂CH₂CH₂CH₃). MS (ESI⁺) (*m/z*) = 183 {[MeOEtBuIM]⁺}. HRMS calcd for C₁₀H₁₉N₂O {[MeOEtBuIM]⁺} 183.1498, found 183.1493. IR (neat) cm⁻¹: 3559, 3114, 2971, 1564, 1461, 1263, 1159, 1029, 836.

4.3. Knoevenagel Condensation Reactions

An aldehyde or a ketone (5 mmol) and an alkylcyanoacetate (5 mmol) were mixed in one of the ionic liquids (5 mmol) listed in Table 2. The reaction mixture was stirred for an appropriate time at 25 °C or 50 °C (Tables 2 and 3). When TLC analysis (diethyl ether/petroleum ether = 1/2, 1/4 or 1/6) indicated the reaction was complete, the resulting product was extracted with ether (3 × 10 mL), and the extraction was washed with water (3 × 10 mL). Ether was removed by rotary evaporation. The product was further purified by recrystallization. To recycle the ionic liquid, the used ionic liquid was washed with ethyl acetate (3 × 10 mL) and then was kept under high vacuum (4 mm Hg) at 80 °C for 3 h.

4.3.1. (E) Ethyl-2-cyano-3-phenyl-2-propenoate (**9a**) [31]

Mp: 47-48 °C. ¹H NMR (250 MHz, DMSO-d₆) δ: 8.39 (s, 1H, HC=C), 8.14-8.02 (m, 2H, Ph-H), 7.74-7.54 (m, 3H, Ph-H), 4.31 (q, *J* = 7.11 Hz, 2H, CH₂CH₃), 1.30 (t, *J* = 7.11 Hz, 3H, CH₂CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 161.72 (COO), 155.02 (HC=CCN), 133.35 (Ph-C₄), 131.27 (Ph-C₂₊₆), 130.74 (Ph-C₁), 129.25 (Ph-C₃₊₅), 115.52 (CN), 102.52 (HC=CCN), 62.33 (OCH₂CH₃), 13.90 (OCH₂CH₃). MS (ESI⁺) (*m/z*): 200 (M-H). IR (KBr) cm⁻¹: 3426, 2980, 2220, 1724, 1601, 1438, 1256, 1196, 1078, 1005, 769, 678.

4.3.2. (E) Ethyl-2-cyano-3-(4-chlorophenyl)-2-propenoate (**9b**) [31]

Mp: 91-92 °C. ¹H NMR (250 MHz, DMSO-d₆) δ: 8.39 (s, 1H, H-C=C), 8.05 (d, *J* = 8.53 Hz, 2H, Ph-H), 7.66 (d,

$J = 8.53$ Hz, 2H, Ph-H), 4.31 (q, $J = 7.11$ Hz, 2H, CH_2CH_3), 1.30 (t, $J = 7.11$ Hz, 3H, CH_2CH_3). ^{13}C NMR (62.9 MHz, DMSO-d_6) δ : 161.56 (COO), 153.60 (HC=CCN), 137.99 (Ph-C₄), 132.39 (Ph-C₂₊₆), 130.14 (Ph-C₁), 129.41 (Ph-C₃₊₅), 115.35 (CN), 103.11 (HC=CCN), 62.42 (OCH_2CH_3), 13.91 (OCH_2CH_3). MS (ESI⁺) (m/z): 234 (M-H). IR (KBr) cm^{-1} : 3434, 2989, 2225, 1717, 1609, 1586, 1488, 1263, 1200, 1074, 1007, 827, 755.

4.3.3. (E) Ethyl-2-cyano-3-(4-methylphenyl)-2-propenoate (9c) [53]

Mp: 92-93 °C. ^1H NMR (250 MHz, DMSO-d_6) δ : 8.32 (s, 1H, H-C=C), 7.95 (d, $J = 8.21$ Hz, 2H, Ph-H), 7.39 (d, $J = 8.21$ Hz, 2H, Ph-H), 4.30 (q, $J = 7.11$ Hz, 2H, CH_2CH_3), 2.39 (s, 3H, CH_3), 1.30 (t, $J = 7.11$ Hz, 3H, CH_2CH_3). ^{13}C NMR (62.9 MHz, DMSO-d_6) δ : 161.95 (COO), 154.88 (HC=CCN), 144.37 (Ph-C₄), 130.94 (Ph-C₂₊₆), 129.90 (Ph-C₃₊₅), 128.62 (Ph-C₁), 115.75 (CN), 101.01 (HC=CCN), 62.22 (OCH_2CH_3), 21.30 (PhCH₃), 13.93 (OCH_2CH_3). MS (ESI⁺) (m/z): 214 (M-H). IR (KBr) cm^{-1} : 3440, 2997, 2218, 1727, 1591, 1273, 1206, 1179, 1094, 816, 758, 494.

4.3.4. (E) Ethyl-2-cyano-3-(4-nitrophenyl)-2-propenoate (9d) [31]

Mp: 170-171 °C. ^1H NMR (250 MHz, DMSO-d_6) δ : 8.55 (s, 1H, H-C=C), 8.40 (d, $J = 9.00$ Hz, 2H, Ph-H), 8.24 (d, $J = 9.00$ Hz, 2H, Ph-H), 4.34 (q, $J = 7.11$ Hz, 2H, CH_2CH_3), 1.32 (t, $J = 7.11$ Hz, 3H, CH_2CH_3). ^{13}C NMR (62.9 MHz, CDCl_3) δ : 161.10 (COO), 152.66 (HC=CCN), 149.19 (Ph-C₄), 137.23 (Ph-C₁), 131.64 (Ph-C₂₊₆), 124.13 (Ph-C₃₊₅), 114.92 (CN), 106.62 (HC=CCN), 62.71 (OCH_2CH_3), 13.91 (OCH_2CH_3). MS (ESI⁺) (m/z): 245 (M-H). IR (KBr, cm^{-1}): 3451, 2991, 2226, 1718, 1621, 1590, 1510, 1342, 1262, 1205, 1196, 865, 763, 688.

4.3.5. (E) Ethyl-2-cyano-3-(4-carboxyphenyl)-2-propenoate (9e)

Mp: 232-234 °C. ^1H NMR (250 MHz, DMSO-d_6) δ : 13.42 (s, 1H, COOH), 8.46 (s, 1H, H-C=C), 8.13 (d, $J = 8.67$ Hz, 2H, Ph-H), 8.08 (d, $J = 8.53$ Hz, 2H, Ph-H), 4.33 (q, $J = 7.11$ Hz, 2H, CH_2CH_3), 1.32 (t, $J = 7.11$ Hz, 3H, CH_2CH_3). ^{13}C NMR (62.9 MHz, DMSO-d_6) δ : 166.36 (COOH), 161.40 (COOC_2H_5), 153.80 (HC=CCN), 135.02 (Ph-C₁), 134.22 (Ph-C₄), 130.69 (Ph-C₂₊₆), 129.82 (Ph-C₃₊₅), 115.20 (CN), 104.73 (NC=CCN), 62.52 (OCH_2CH_3), 13.90 (OCH_2CH_3). HRMS calcd for $\text{C}_{13}\text{H}_{12}\text{NO}_4$ [M+H] 246.0766, found 246.0769. IR (KBr) cm^{-1} : 3424, 2992, 2205, 1733, 1702, 1612, 1421, 1283, 1196, 1006, 850, 768, 686, 543.

4.3.6. (E) Ethyl-2-cyano-3-(2-naphthyl)-2-propenoate (9f) [54]

Mp: 111-112 °C. ^1H NMR (250 MHz, DMSO-d_6) δ : 8.60 (s, 1H, H-C=C), 8.53 (s, 1H, Ar-H), 8.26-7.99 (m, 4H, Ar-H), 7.78-7.61 (m, 2H, Ar-H), 4.34 (q, $J = 6.95$ Hz, 2H, CH_2CH_3), 1.33 (t, $J = 6.95$ Hz, 3H, CH_2CH_3). ^{13}C NMR (62.9 MHz, DMSO-d_6) δ : 161.90 (COO), 154.92 (HC=CCN), 134.76 (Ar-C), 134.43 (Ar-C), 132.29 (Ar-C), 129.27 (Ar-C), 129.24 (Ar-C), 128.95 (Ar-C), 128.91 (Ar-C), 127.81 (Ar-C), 127.41 (Ar-C), 124.48 (Ar-C), 115.77 (CN), 102.28 (HC=CCN), 62.36 (OCH_2CH_3), 13.97 (OCH_2CH_3). MS (ESI⁺) (m/z): 250 (M-H). IR (KBr) cm^{-1} : 3439, 2981, 2222, 1724, 1598, 1248, 1181, 817, 750, 476.

4.3.7. (E) Methyl-2-cyano-3-(3-chlorophenyl)-2-propenoate (9g)

Mp: 111-112 °C. ^1H NMR (250 MHz, DMSO-d_6) δ : 8.42 (s, 1H, H-C=C), 8.09 (t, $J = 1.90$ Hz, 1H, Ph-H), 8.05-7.99 (m, 1H, Ph-H), 7.73-7.58 (m, 2H, Ph-H), 3.87 (s, 3H, CH_3). ^{13}C NMR (62.9 MHz, DMSO-d_6) δ :

161.89 (COO), 153.44 (HC=CCN), 133.80 (Ph-C₃), 133.24 (Ph-C₁), 132.74 (Ph-C₅), 131.07 (Ph-C₂), 130.16 (Ph-C₄), 128.94 (Ph-C₆), 115.18 (CN), 103.95 (HC=CCN), 53.42 (OCH₃). HRMS calcd for C₁₁H₈NO₂Cl [M⁺] 221.0244, found 221.0247. IR (KBr) cm⁻¹: 3431, 3059, 2218, 1722, 1612, 1562, 1433, 1272, 1208, 1093, 781, 680.

4.3.8. (E) Methyl-2-cyano-3-(4-methylphenyl)-2-propenoate (9h)

Mp: 108-110 °C. ¹H NMR (250 MHz, DMSO-d₆) δ: 8.37 (s, 1H, H-C=C), 7.99 (d, *J* = 8.21 Hz, 2H, Ph-H), 7.41 (d, *J* = 8.06 Hz, 2H, Ph-H), 3.86 (s, 3H, OCH₃), 2.40 (s, 3H, CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 162.51 (COO), 155.07 (HC=CCN), 144.46 (Ph-C₄), 130.08 (Ph-C₂₊₆), 129.95 (Ph-C₃₊₅), 128.64 (Ph-C₁), 115.80 (CN), 100.77 (HC=CCN), 53.25 (OCH₃), 21.33 (PhCH₃). HRMS calcd for C₁₂H₁₁NO₂ [M⁺] 201.0790, found 201.0792. IR (KBr) cm⁻¹: 3434, 2952, 2218, 1722, 1590, 1435, 1271, 1209, 1183, 1094, 811, 489.

4.3.9. (E) Methyl-2-cyano-3-(4-methoxyphenyl)-2-propenoate (9i) [55]

Mp: 102-103 °C. ¹H NMR (250 MHz, DMSO-d₆) δ: 8.31 (s, 1H, H-C=C), 8.09 (d, *J* = 9.00 Hz, 2H, Ph-H), 7.15 (d, *J* = 9.00 Hz, 2H, Ph-H), 3.87 (s, 3H, OCH₃), 3.84 (s, 3H, COOCH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 163.53 (COO), 162.84 (Ph-C₄), 154.51 (HC=CCN), 133.51 (Ph-C₂₊₆), 123.87 (Ph-C₁), 116.19 (CN), 114.91 (Ph-C₃₊₅), 98.13 (HC=CCN), 55.71 (OCH₃), 53.07 (COOCH₃). MS (ESI⁻) (m/z): 216 (M-H). IR (KBr) cm⁻¹: 3434, 2949, 2213, 1720, 1585, 1554, 1509, 1423, 1257, 1208, 1168, 1024, 840, 548.

4.3.10. (E) Methyl-2-cyano-3-(4-hydroxyphenyl)-2-propenoate (9j)

Mp: 213-214 °C. ¹H NMR (250 MHz, DMSO-d₆) δ: 10.89 (s, 1H, OH), 8.26 (s, 1H, H-C=C), 8.00 (d, *J* = 8.85 Hz, 2H, Ph), 6.95 (d, *J* = 8.64 Hz, 2H, Ph), 3.83 (s, 3H, CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 163.10 (COO), 162.08 (Ph-C₁), 154.79 (HC=CCN), 133.00 (Ph-C₂₊₆), 122.42 (Ph-C₄), 116.45 (CN), 111.36 (Ph-C₃₊₅), 96.59 (HC=CCN), 53.00 (OCH₃). HRMS calcd for C₁₁H₁₀NO₃ [M+H] 204.0661, found 204.0663. IR (KBr) cm⁻¹: 3338, 2222, 1724, 1590, 1434, 1270, 1209, 1170, 1088, 815, 513.

4.3.11. Methyl-2-cyano-3-cyclopentanyl-2-propenoate (9k)

Mp: 23-25 °C. ¹H NMR (250 MHz, DMSO-d₆) δ: 3.75 (s, 3H, CH₃), 2.91 (t, *J* = 6.95 Hz, 2H, CH₂C=C), 2.75 (t, *J* = 7.11 Hz, 2H, C=CCH₂), 1.85-1.69 (m, 4H, CH₂CH₂). ¹³C NMR (62.9 MHz, DMSO-d₆): 188.79 (C=CCN), 161.77 (COO), 115.45 (CN), 99.15 (C=CCN), 52.28 (OCH₃), 37.39 (CH₂), 35.25 (CH₂), 25.97 (CH₂), 24.56 (CH₂). HRMS calcd for C₉H₁₁NO₂ [M⁺] 165.0790, found 165.0793. IR (KBr) cm⁻¹: 3440, 2957, 2224, 1727, 1612, 1439, 1279, 1203, 1088, 1026, 773.

4.3.12. Methyl-2-cyano-3-cyclohexanyl-2-propenoate (9l, liquid)

¹H NMR (250 MHz, DMSO-d₆) δ: 3.76 (s, 3H, -CH₃), 2.95 (t, *J* = 5.61 Hz, 2H, CH₂C=C), 2.62 (t, *J* = 5.85 Hz, 2H, C=CCH₂), 1.78-1.57 (m, 6H, CH₂CH₂CH₂). ¹³C NMR (62.9 MHz, DMSO-d₆) δ: 180.95 (C=CCN), 161.85 (COO), 115.41 (CN), 100.56 (C=CCN), 52.53 (OCH₃), 36.19 (CH₂), 30.92 (CH₂), 28.12 (CH₂), 27.77 (CH₂), 24.83 (CH₂). HRMS calcd for C₁₀H₁₃NO₂ [M⁺] 179.0946, found 179.0944. IR (neat) cm⁻¹: 3446, 2940, 2222, 1734, 1601, 1437, 1269, 1216, 1096, 1012, 777.

4.3.13. (E) Methyl-2-cyano-3-phenyl-2-butyrate (9m, liquid)

¹H NMR (250 MHz, DMSO-d₆, major isomer) δ: 7.63-7.28 (m, 5H, Ph-H), 3.82 (s, 3H, OCH₃), 2.66 (s, 3H, CH₃C=C). ¹³C NMR (62.9 MHz, DMSO-d₆, major isomer) δ: 173.02 (COO), 162.19 (C=CCN), 138.49 (Ph-C₁), 130.31 (Ph-C₄), 128.55 (Ph-C₃₊₅), 127.19 (Ph-C₂₊₆), 116.15 (CN), 103.99 (C=CCN), 52.77 (OCH₃), 26.65 (CH₃C=C). HRMS calcd for C₁₂H₁₁NO₂ [M⁺] 201.0790, found 201.0793. IR (neat) cm⁻¹: 3424, 2976, 2216, 1719, 1599, 1437, 1261, 1194, 1095, 762, 678.

4.3.14. (E) Ethyl-2-cyano-3-phenyl-2-butyrate (9n, liquid) [31]

¹H NMR (250 MHz, DMSO-d₆, major isomer) δ: 7.70-7.27 (m, 5H, Ph-H), 4.29 (q, *J* = 6.95 Hz, 2H, CH₂CH₃), 2.65 (s, 3H, CH₃), 1.29 (t, *J* = 6.95 Hz, 3H, CH₂CH₃). ¹³C NMR (62.9 MHz, DMSO-d₆, major isomer) δ: 172.67 (COO), 161.71 (C=CCN), 140.13 (Ph-C₁), 133.12 (Ph-C₄), 129.38 (Ph-C₃₊₅), 126.52 (Ph-C₂₊₆), 116.10 (CN), 104.29 (C=CCN), 61.73 (OCH₂CH₃), 26.48 (CH₃C=C), 13.87 (OCH₂CH₃). MS (ESI⁺) (m/z): 214 (M-H). IR (neat) cm⁻¹: 2985, 2225, 1729, 1592, 1442, 1369, 1242, 1137, 1046, 764, 700.

4.3.15. (E) Ethyl-2-cyano-2-hexenoate (9o, liquid) [56]

¹H NMR (250 MHz, CDCl₃) δ: 7.65 (t, *J* = 7.80 Hz, 1H, CH=C), 4.30 (q, *J* = 7.20 Hz, 2H, OCH₂CH₃), 2.51-2.60 (m, 2H, CH₂CH₂CH=C), 1.57-1.68 (m, 2H, CH₃CH₂CH₂CH=C), 1.38 (t, *J* = 7.20 Hz, 3H, OCH₂CH₃), 0.99 (t, *J* = 7.10 Hz, 3H, CH₃CH₂CH₂CH=C). ¹³C NMR (62.9 MHz, CDCl₃) δ: 163.46 (COO), 161.10 (HC=CCN), 113.66 (CN), 109.93 (HC=CCN), 62.38 (OCH₂CH₃), 33.75 (CH₃CH₂CH₂CH=C), 21.16 (CH₃CH₂CH₂CH=C), 14.10 (OCH₂CH₃), 13.72 (CH₃CH₂CH₂CH=C). MS (ESI⁺) (m/z): 166 (M-H). IR (neat) cm⁻¹: 3436, 2965, 2231, 1732, 1270, 760.

4.3.16. (E) Ethyl-2-cyano-3-(2-furyl)-2-propenoate (9p) [31]

Mp: 92-93 °C. ¹H NMR (250 MHz, CDCl₃) δ: 8.03 (s, 1H, CH=C), 7.76 (d, *J* = 1.62 Hz, 1H, furyl-H), 7.41 (d, *J* = 4.0 Hz, 1H, furyl-H), 6.69 (dd, *J* = 4.0, 1.62 Hz, 1H, furyl-H), 4.31 (q, *J* = 7.10 Hz, 2H, OCH₂CH₃), 1.31 (t, *J* = 7.10 Hz, 3H, OCH₂CH₃). ¹³C NMR (62.9 MHz, CDCl₃) δ: 162.13 (COO), 149.97 (furyl-C₂), 148.13 (furyl-C₃), 139.01 (HC=CCN), 124.41 (furyl-C₃), 115.22 (CN), 114.21 (furyl-C₄), 96.90 (HC=CCN), 62.07 (OCH₂CH₃), 13.91 (OCH₂CH₃). MS (ESI⁺) (m/z): 190 (M-H). IR (KBr) cm⁻¹: 3554, 3410, 2989, 2406, 1736, 1621, 1460, 1218, 779.

Conflict of Interest

The authors confirm that this article content has no conflict of interest.

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Figures:

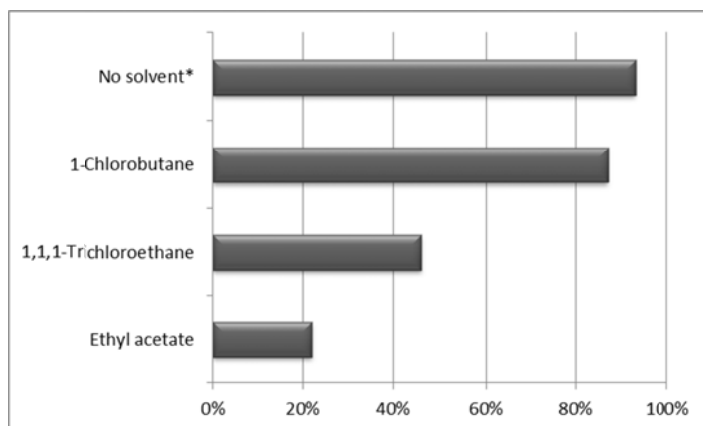


Figure 1. Comparison of various solvents for the preparation of [BMIM]⁺Cl⁻ (**2a**) (*equal molar of the starting materials without addition of other solvents).

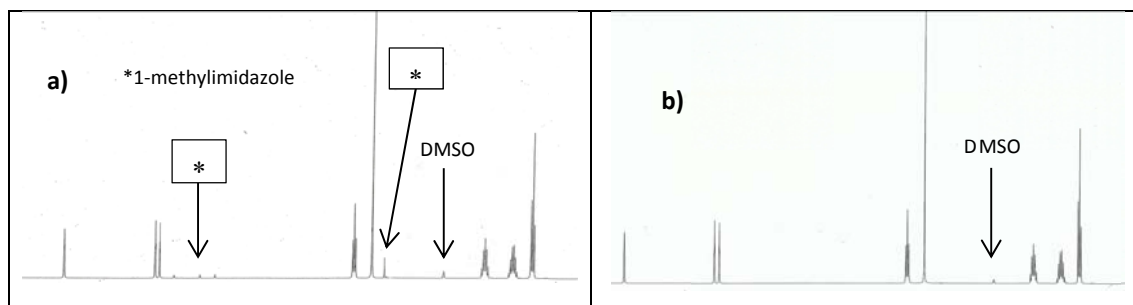
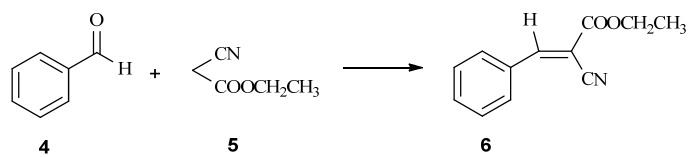
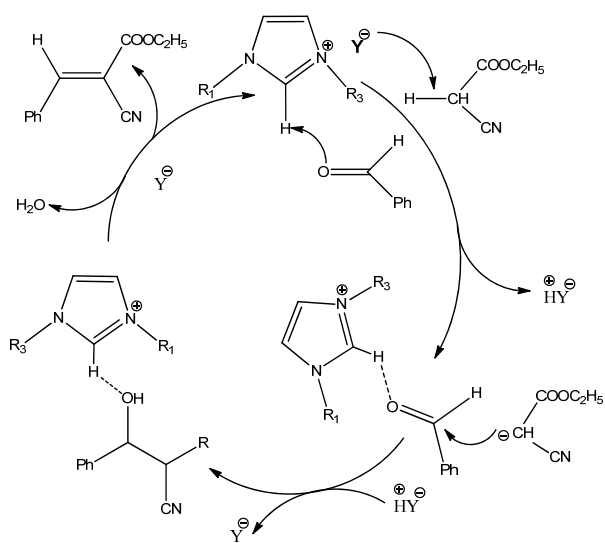


Figure 2. ¹H NMR spectrum of 1-butyl-3-methylimidazolium chloride, a) purified by heating under high vacuum; b) by recrystallization.

Schemes:



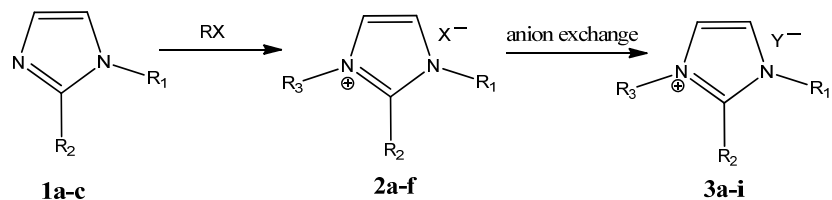
Scheme 1. Reaction of benzaldehyde with ethyl cyanoacetate in various ILs.



Scheme 2. Proposed mechanism for the Knoevenagel reaction.

Tables:

Table 1. Synthesis of ionic liquids



Compound No	R ₁	R ₂	R ₃	X ⁻	Y ⁻
1a	CH ₃	H			
1b	CH ₃	CH ₃			
1c	n-Bu	H			
2a	CH ₃	H	n-Bu	Cl	
2b	CH ₃	CH ₃	n-Bu	Cl	
2c	CH ₃	H	MeOMeOEt	Br	
2d	CH ₃	H	MeOEt	Cl	
2e	n-Bu	H	MeOEt	Cl	
3a	CH ₃	H	n-Bu		BF ₄
3b	CH ₃	H	n-Bu		PF ₆
3c	CH ₃	H	n-Bu		CF ₃ COO
3d	CH ₃	CH ₃	n-Bu		CF ₃ COO
3e	CH ₃	H	MeOMeOEt		CF ₃ COO
3f	CH ₃	H	MeOEt		CF ₃ COO
3g	CH ₃	H	MeOEt		CF ₃ SO ₃
3h	n-Bu	H	MeOEt		CF ₃ SO ₃

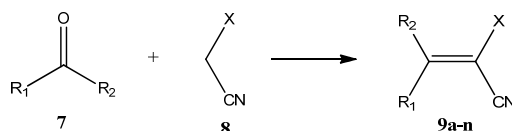
3a: 1-Butyl-3-methylimidazolium tetrafluoroborate {[BMIM]⁺[BF₄]⁻}
 3b: 1-Butyl-3-methylimidazolium hexafluorophosphate {[BMIM]⁺[PF₆]⁻}
 3c: 1-Butyl-3-methylimidazolium trifluoroacetate {[BMIM]⁺[CF₃COO]⁻}
 3d: 1-Butyl-2,3-dimethylimidazolium trifluoroacetate {[BMMIM]⁺[CF₃COO]⁻}
 3e: 1-Methoxymethoxyethyl-3-methylimidazolium trifluoroacetate {[MeOMeOEtMIM]⁺[CF₃COO]⁻}
 3f: 1-Methoxyethyl-3-methylimidazolium trifluoroacetate {[MeOEtMIM]⁺[CF₃COO]⁻}
 3g: 1-Methoxyethyl-3-methylimidazolium trifluoromethanesulfonate {[MeOEtMIM]⁺[CF₃SO₃]⁻}
 3h: 1-Methoxyethyl-3-butylimidazolium trifluoromethanesulfonate {[MeOEtBuIM]⁺[CF₃SO₃]⁻}

Table 2. Knoevenagel condensation of benzaldehyde with ethyl cyanoacetate in various ionic liquids^a

Entry	Solvent	Reaction time	Yield (%) ^b
1	[BMIM] ⁺ [BF ₄] ⁻ (3a)	24 h	85
2	[BMIM] ⁺ [PF ₆] ⁻ (3b)	22 h	63
3	[BMIM] ⁺ [CF ₃ COO] ⁻ (3c)	1 h	98
4	[BMMIM] ⁺ [CF ₃ COO] ⁻ (3d)	4 h	62
5	[MeOMeOEtMIM] ⁺ [CF ₃ COO] ⁻ (3e)	40 min	95
6	[[MeOEtMIM] ⁺ [CF ₃ COO] ⁻ (3f)	40 min	98
7	[MeOEtMIM] ⁺ [CF ₃ SO ₃] ⁻ (3g)	1 h	81
8	[MeOEtBuIM] ⁺ [CF ₃ SO ₃] ⁻ (3h)	2 h	54

^a Reaction conditions: benzaldehyde (5 mmol), ethyl cyanoacetate (5 mmol), IL (5 mmol) at 25 °C

^b Isolated yield of products.

Table 3. Knoevenagel condensation of aldehydes and ketones with methylene compounds in [MeOEtMIM]⁺[CF₃COO]⁻^a

Entry	R1	R2	X	Time	Temperature	Yield (%) ^b	Product
1	Ph	H	COOEt	40 min	25 °C	98	9a
2	4-Cl-C ₆ H ₄	H	COOEt	5 min	25 °C	97	9b
3	4-Me-C ₆ H ₄	H	COOEt	20 min	25 °C	89	9c
4	4-NO ₂ -C ₆ H ₄	H	COOEt	10 min	25 °C	81	9d
5	4-HOOC-C ₆ H ₄	H	COOEt	50 min	25 °C	77	9e
6	2-Naphthyl	H	COOEt	15 min	25 °C	98	9f
7	3-Cl-C ₆ H ₄	H	COOMe	2 min	25 °C	99	9g
8	4-Me-C ₆ H ₄	H	COOMe	10 min	25 °C	91	9h
9	4-MeO-C ₆ H ₄	H	COOMe	35 min	25 °C	78	9i
10	4-HO-C ₆ H ₄	H	COOMe	40 min	25 °C	83	9j
11	-(CH ₂) ₄ -		COOMe	48 h	25 °C	85	9k
12	-(CH ₂) ₅ -		COOMe	16 h	25 °C	56	9l
13	Ph	CH ₃	COOMe	48 h	50 °C	55	9m
14	Ph	CH ₃	COOEt	48 h	50 °C	47	9n
15	n-C ₃ H ₇	H	COOEt	2 h	25 °C	82	9o
16	2-Furyl	H	COOEt	80 min	25 °C	85	9p

^a Reaction conditions: aldehyde or ketone (5 mmol), active methylene compound (5 mmol). Ionic liquid (5 mmol).

^b Isolated yield of products.

Table 4. Knoevenagel condensation of benzaldehyde with ethyl cyanoacetate

Ionic liquid	Yield (%)
[MeOEtMIM] ⁺ [CF ₃ COO] ⁻ (Fresh)	98
[MeOEtMIM] ⁺ [CF ₃ COO] ⁻ (cycle1)	94
[MeOEtMIM] ⁺ [CF ₃ COO] ⁻ (cycle2)	91
[MeOEtMIM] ⁺ [CF ₃ COO] ⁻ (cycle3)	88
[MeOEtMIM] ⁺ [CF ₃ COO] ⁻ (cycle4)	85
[MeOEtMIM] ⁺ [CF ₃ COO] ⁻ (cycle5)	86