



Eco-friendly and economic method for Knoevenagel condensation by employing natural catalyst of guava leaves (*Psidium guajava*)

K Kalaiselvi¹, S Shamala²

¹ Research Scholar, Department of Chemistry, PRIST University, Puducherry, India

² Assistant Professor, Department of Chemistry, PRIST University, Puducherry, India

Abstract

The antimicrobial potential of guava (*Psidium guajava*) leaf extracts against two gram-negative bacteria (*Escherichia coli* and *Salmonella enteritidis*) and two gram-positives bacteria (*Staphylococcus aureus* and *Bacillus Cereus*) which are some of food borne and spoilage bacteria. The guava leaves were extracted in four different solvents of increasing polarities (hexane, methanol, ethanol and water).

The guava (*Psidium guajava*) is a Phytotherapeutic plant used in folk medicine that is believed to have active components that help to treat and manage various diseases. The challenges in organic synthesis are develop convenient process. Reaction media conditions and utility of material based on the design of green chemistry is one of the important issues in the chemical society in resent strong synthesis effort.

Keywords: cement emissions, environmental pollution, atmospheric changes, human health

Introduction

Chemistry is undeniably a very prominent part of our daily lives. However chemical development also bring new environmental problems and harmful unexpected site effect considering a growing need for more environmentally acceptable process in the chemical industry. The challenges in organic synthesis are develop convenient process. Reaction media conditions and utility of material based on the design of green chemistry is one of the important issues in the chemical society in resent strong synthesis effort are done to use of approaches that are beneficial to industry as well as the environment [1].

Green chemistry

Green chemistry is basically environmentally begin chemical synthesis and is useful to reduce environment pollution. Green chemistry efficiently utilised raw materials, eliminated waste, and avoid the use of toxic or hazardous reagent and solvent in the manufacture and application of chemical products.

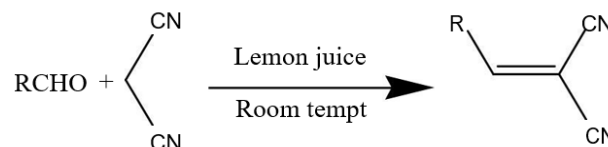
The twelve basic principles of green chemistry have been formulated.

1. It is better to prevent waste than to treat or clean up waste after it is formed.
2. Synthetic materials should be designed to maximize the incorporation of all materials used in the process into the final product.

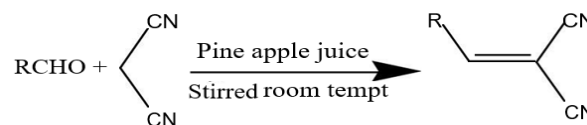
Natural Catalyst

Some of the Knoevenagel condensation employing natural catalyst such as Henna leaves, Clay, natural phosphates, Animal bone, Calcined eggshell (CES), and various fruit juices are reported due to acidic natural aqueous fruit juice like Lemon [37], Pineapple [38], Tamarind Indica [39], Star fruit juice [2], and also biocatalyst etc.

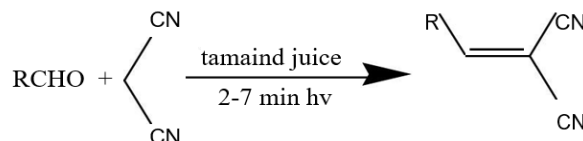
M.B. Deshmukh *et al.* [37] Have reported the Knoevenagel condensation of active methylene compound with aromatic aldehyde in the presence of lemon juice.



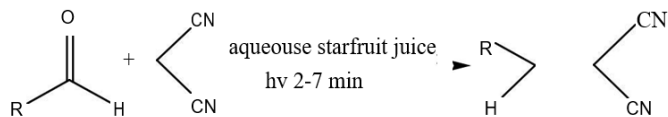
Suresh and Patil *et al.* [38] have introduced simple eco-friendly and economic method for Knoevenagel condensation of substituted aromatic as well as hetero aromatic aldehyde with malononitrile catalyzed by pineapple Juice at room temperature in the absence of any organic solvent is described.



Rammohan pal *et al.* [39] have introduced Visible light induced a highly efficient and environmentally friendly Knoevenagel condensation of various aliphatic and aromatic aldehyde with malononitrile has been achieved in excellent yield in presence of aqueous tamarind Juice.

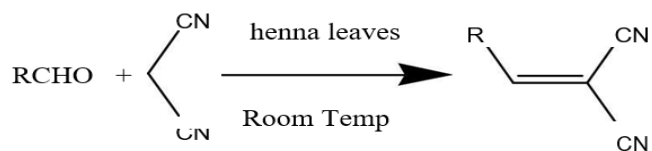


Rammohan pal, *et al* [2] have introduced a aqueous star fruit Juice catalyzed a simple and efficient knoevenagel condensation of aromatic aldehyde with malononitrile has been developed under visible light. The synthesis of intermediate such a coumarin derivative which are useful in perfumes.



Pravin chavan *et al.* [58] have introduced a simple green method for condensation of substituted aromatic aldehyde with malononitrile catalyses by extract of henna leaves at room temperature in absence of any chemical reagent. The

knoevenagel condensation by Henna juice Catalyst with good yield.



Preparation of extract of guava leaf

Guava leaf were collected and washed with water [fig.2]. The were dried and powdered by using mortar. 5g of the Powder and 50ml of methanol were taken in 250ml iodine flask and it was stirred by mechanical stirrer for one hour. The extract was filtered using filter paper and then it was used as catalyst for all reactions.

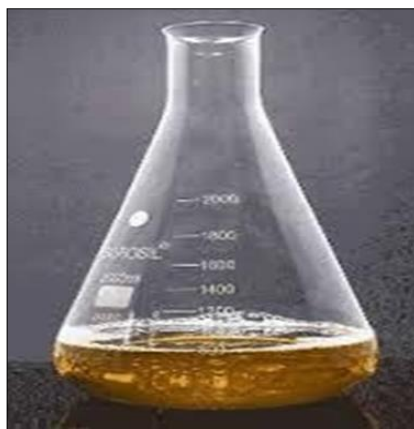


Fig 1: guava leaves extract

Results and Discussion

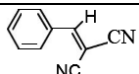
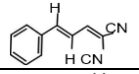
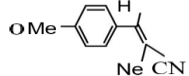
The fresh leaves are used for treating gastrointestinal problems, then also used to control blood pressure. Guava leaf extract has analgesic, anti-inflammatory, antimicrobial, hepatoprotective and antioxidant activities. The dried guava leaves are considered as waste material, it cannot be feeded for animals. But we are using the waste materials as Natural catalyst for our chemical reaction. This natural catalyst gives a good yield in the knoevenagel condensation reaction.

Kim *et al.* reported that the guava leaves contain ascorbic acid, citric acid, acetic acid, epicatechin, xanthine, protocatechuic acid, glutamic acid, asparagine, malonic acid, trans-aconitic acid, maleic acid and cis-aconitic acid. As guava leaves extract is acidic in nature it will be worked as a acid catalyst for knoevenagel condensation Hence we using for Knoevenagel condensation of active methylene compound with aromatic aldehydes in the presence of guava leaf extract (*Psidium guajava*) as a natural catalyst.



Fig 2: guava leaves

Table 1: Knoevenagel condensation of aldehydes with malononitrile catalyzed by Sweet lime peel extract.

Compound	Product	Color & nature of product	Yield (%)	Time (min)	Melting Point(c)	
					Observed	Reported
1.		White crystal	90	2	81	80-82
2.		Yellow Crystal	80	5	127	126-128
3.		Green Crystal	88	5	114	112-114

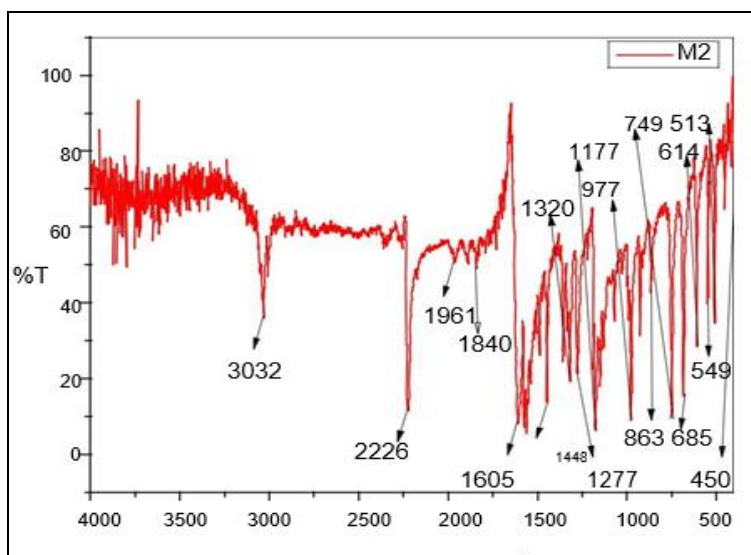
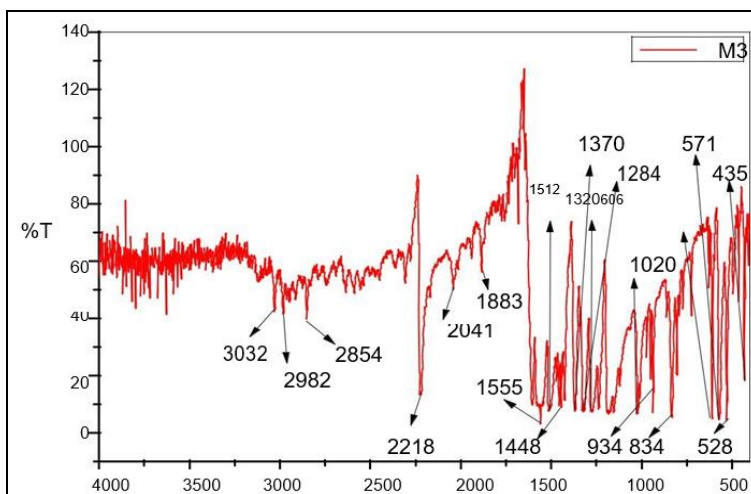
At room temperature the high yield were obtained with in few minutes by using magnetic stirrer. Their actions were monitor

ed by thin layer chromatography techniques. The melting of all the compounds are good agree with literature.

IR-Spectral Analysis

Table 2: IR data of compound (1-3)

Compounds	CN (cm-1)	Other Frequencies (cm-1)
1	2226	1591,3032,614,1220,1448
2	2226	3032,1605,1448,1227,863,685
3	2218	3032,1883,1555,1448,934,834

**Fig 3:** Wave Number (cm) IR spectrum of compound-1**Fig 4:** Wave Number (cm) IR spectrum of compound-3

¹H-Nmrspectralanalysis

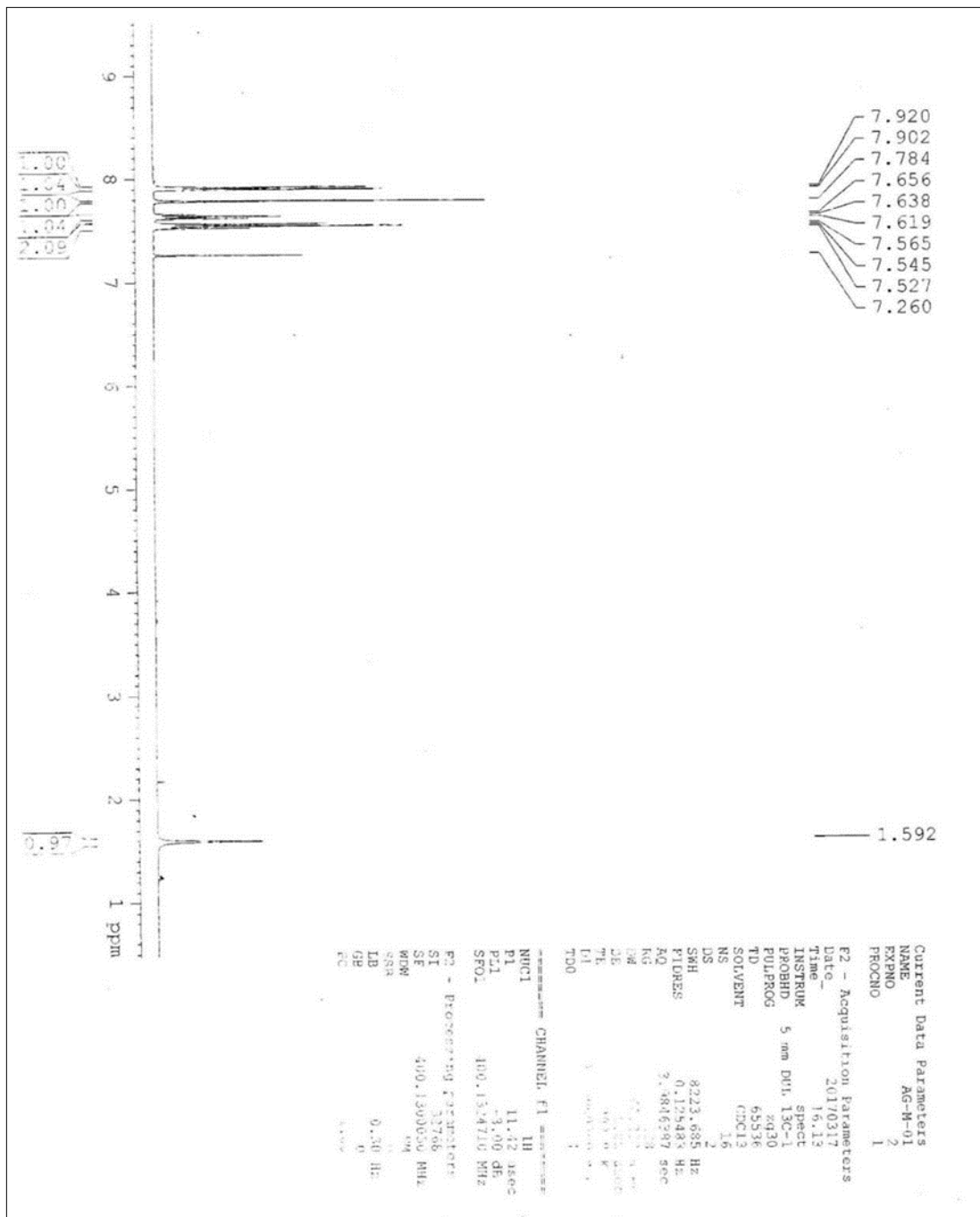
¹H-NMR spectroscopy is an important tool in the hand so fan organic chemist for getting structural information from the spectrum fanun known compound. It also helps in study in g the stereochemical details with in the molecule. Al though important, it cannot replace other tech niquessuc has ultraviolet, in frared, massetc. Leaving aside the functional groups, alarge part of anorganic mole culecons is ts of carbon-hydrogen skele ton and this tool is most use fulin the invest negation of this structural feature of the molecule-rather than the complete structure.

¹H-spectrum of compound 2(phenyl methylene) malononitrile

¹H NMR spectra recorded in CDCl₃ for the 2 (phenyl methylene) malononitrile. The data is good agree with literature.

Table 3: The data are listed in the.

Compound	CH proton	Aromatic protons		
		Ortho	Meta	Para
2(phenyl methylene)malononitrile	7.8	7.91(ppm)	7.56(ppm)	7.6(ppm)

**Fig 5**

C13nmrspectraanalysis

¹³C NMR is an important tool in chemical structure elucidation in organic Chemistry. ¹³CNMR detects only The ¹³ cisotope of carbon, who's natural Abundance is only1.1%, because the main carbon isotope, ¹²c, is not detec table by NMR since it has zero net spin. ¹³C chemical hifts follow the same principles as those of¹H, although the typical range of chemical shifts is much larger thanfor¹H (by a factor of about 20). The chemical shift references tandard for¹³C is the carbons in tetramethy lsilane (TMS), ^[1] whose chemical

shift is considered to be0.0 ppm¹³C NMR spectrarecorded inCDCl₃forthe 2 (phenylmethylene) malononitrile. The data are listed in the table -4.

Compound	CH proton	Aromatic protons		
		Ortho	Meta	Para
2(phenyl methylene)malo nonitrile	7.8	7.91(ppm)	7.56(ppm)	7.6(ppm)

¹H-spectrum of compound 2 (pheny methylene) malononitrile

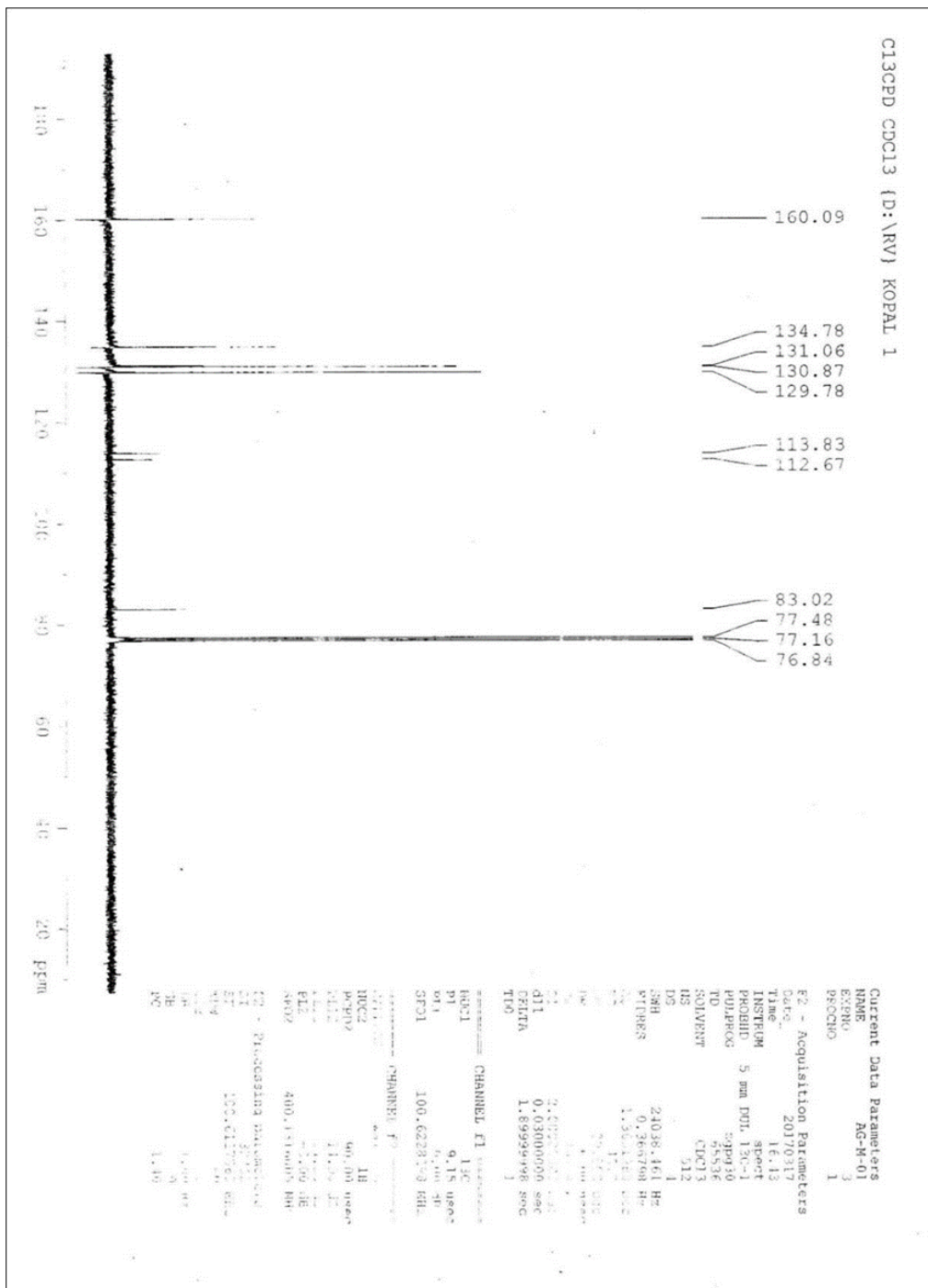


Fig 6

Conclusion

We have established and potentially efficient absolute clean and high yielding eco-friendly methodology for the Knoevenagel condensation of various aromatic aldehyde with malononitrile using sweet lime peel devoid of any toxic catalyst/solvent. Solid support and surfactant and may be considered as an excellent improvement are the existing method. The most attractive features of this protocol are its good conversions easy work up and short reaction time making it useful practical method for a synthesis of adducts. This solvent free approach is based on green chemistry principle and do not cause any harm to environment. In addition it involves mild reaction condition and simple work up.

References

1. Yu N, Aramini JM, German MW, Huang Z. Reactions of salicylaldehyde with alkyl cyanoacetates on the surface of solid catalysts: synthesis of 4H-chromene derivatives, *Tetrahedron Letters*. 2000; 41(36): 6993- 6996.
2. Gallos J, Discordia RP, Crispino GA, Li J, Grosso JA, Polniaszek V, True VC. A mild and efficient synthesis of 4-aryl-quinolin-2(1H)-ones via a tandem amidation/Knoevenagel condensation of 2-amino-benzophenones with esters or lactones, *Tetrahedron Letters*. 2003; 44(22):4271-4273.
3. Xing C, Zhu S. unexpected formation of tetrasubstituted 2, 3- dihydrofurans from the reactions of β -keto polyfluoroalkanesulfones with aldehydes, *Journal of Organic Chemistry*. 2004; 69(19):6486-6488.
4. Tietze LF, Rackelmann N. Domino reactions in the synthesis of heterocyclic natural products and analogs, *Pure and Applied Chemistry*. 2004; 76(11):1967-1983.
5. Rao PS, Vendataratnam RV. Zinc chloride as a new catalyst for Knoevenagel condensation, *Tetrahedron Letters*. 1991; 32(41):5821-5822.
6. Prajapati D, Sandhu JS. Cadmium iodide as a new catalyst for Knoevenagel condensation, *Journal of the Chemical Society, Perkin Transactions*. 1993; 1(1):739-740.
7. Lehnert W. Verbesserte variante der Knoevenagel condensation mit $\text{TiCl}_4/\text{THF}/\text{pyridine}$ (I). Alkyliden-undarylidenmaloeser bei 0-25°C, *Tetrahedron Letters*. 1970; 11(54):4723-4724.
8. Dai G, Shi D, Zhou L, Huaxue Y. Knoevenagel condensation, 1995.
9. Catalyzed by potassium fluoride/alumina, *Chinese Journal of Applied Chemistry*. 1995; 12(2): 104-108.
10. Gill C, Pandhare G, Raut R, Gore V, Gholap S. Knoevenagel condensation: A simple and efficient protocol of electrophilic alkenes catalyzed by anhydrous ferric sulphate with remarkable reusability, *Bulletin of the Catalysis society of India*. 2008; 7:153-157.
11. Mogilaiah K, Reddy CS. an efficient Friedlander condensation using sodium fluoride as catalyst in the solid state, *Synthetic Communications*. 2003; 33(18):3131-3134.
12. Mallouk S, Bougrin K, Laghzizil A, Benhida R. Microwave assisted and efficient solvent-free Knoevenagel condensation. A sustainable protocol using calcium hydroxyapatite as catalyst, *Molecules*. 2010; 15(2):813-823.
13. Bhuiyan MMH, Hossain MI, Asharaful M & Mahmud MM. Microwave assisted Knoevenagel condensation: Synthesis and antimicrobial activities of some arylidene-malononitriles, *Chemistry Journal*. 2012; 2(1):30-36.
14. McNulty J, Steere JA, Wolf S. The ultrasound promoted Knoevenagel condensation of aromatic aldehydes, *Tetrahedron Letters*. 1998; 39(44):8013-8016.
15. Palmisano G, Tibiletti F, Penoni A, Colombo F, Tollari S, Garella D, Tagliapietra S, Cravotto G. Ultrasound-enhanced one-pot synthesis of 3-(Het)arylmethyl-4 hydroxycoumarins in water, *Ultrasonics Sonochemistry*. 2011; 18(2):652-660.
16. Pratap UR, Jawale DV, Waghmare RA, Lingampalle DL, Mane RA. Synthesis of 5-arylidene-, 2,4-thiazolidinediones by Knoevenagel condensation catalyzed by baker's yeast, *New Journal of Chemistry*. 2011; 35(1):49-51.
17. Wang C, Guan Z, He Y. Biocatalytic domino reaction synthesis of 2H-1-benzopyran-2-one derivatives using alkaline protease from *Bacillus licheniformis*, *Green Chemistry*. 2011; 13(8):2048-2054.
18. Xia Y, Yang Z-Y, Brossi A, Lee K-H. Asymmetric solid phase synthesis of (3R, 4R)-di-O-cis-acyl 3-carboxyl khellactones, *Organic Letters*. 1999; 1(13):2113-2115.
19. Guo G, Arvanitis EA, Pottorf RS, Player MP. Solid phase synthesis of a tyrophostin ether library, *Journal of Combinatorial Chemistry*. 2003; 5(4):408- 413.
20. Ying AG, Liu L, Wu GF, Chen XZ, Ye WD, Chen JH, Zhang KY. Knoevenagel condensation catalyzed by DBU Bronsted ionic liquid without solvent, *Chemical Research in Chinese Universities*. 2009; 25(6):876-881.
21. Khan FA, Dash FJ, Satapathy R, Upadhyay SK. Hydrocalcite catalysis in ionic medium: a recyclable reaction system for heterogeneous Knoevenagel and nitroaldol condensation, *Tetrahedron Letters*. 2004; 45(15):3055- 3058.
22. Verdia P, Santamarta F, Tojo E. Knoevenagel reaction in $[\text{MMIm}][\text{MSO}_4]$: synthesis of coumarins, *Molecules*. 2011; 16(6):4379-4388.
23. Bigi F, Conforti ML, Maggi R, Piccinno Sartori G. clean synthesis in water: Uncatalysed preparation of arylidene malonitriles, *Green Chemistry*. 2000; 2(3):101-103.
24. Wang S, Ren Z, Cao W, Tong W. The Knoevenagel condensation of aromatic aldehydes with malononitriles or ethyl cyanoacetate in the presence of CTMAB in water, *Synthetic Communications*. 2001; 31(5): 673-677.
25. Oskooie HA, Heravi MM, Derikvand F, Khorasani M, Bamoharram FF. On water: An efficient Knoevenagel condensation using 12- Tungstophoric acid as a reusable green catalyst, *Synthetic Communications*. 2006; 36(19):2819-2823.
26. Wang QL, Ma YD, Zuo BJ. *Synth Communication* 27, 1997, 4107. Ren YM, Cai C. *Synth Commun*. 2007; 372:816.
27. Balalaie S, Bararjanian M, Hekmat S, Salehi P. *Synth Commun*. 2006; 549.

28. Abaee MS, Mojtahedi MM, Zahedi MM, Khanalizadeh G, Arkivoc, 2006; 10:48.
29. Hangarge RV, Jarikote DV, Shingare MS. Green Chem. 2002; 4:266.
30. Wang QL, Ma, YD, Zuo BJ. Synth Communication. 1997; 27:4107.
31. Gora M, Kozik B, Jamroz K, Luczyński MK, Bruzulan P, Wozny M, *et al.* 2009; 11:863.
32. Kwon PS, Kim YM, Kang CJ, Kwon TW, Chung SK, Chank YT. Synth Commun. 1997; 27:4091.
33. Ren YM, Cai C. Synth Commun, 372007, 2209.
34. Wang S, Ren Z, Cao W, Tong W. Synth Commun. 2006; 31(5):673.
35. Gill C, Pandhare G, Raut R, Gore V, Gholap S. Bull Cal Soc India, 7, 2008, 153.
36. Oskooie HA, Heravi MM, Derikvand F, Khorasani M & Bamoharram FF, Synth Commun. 2009; 36:2819.
37. Balalaie S, Bararjanian M, Hekmat S & Salehi P, Synth Commun. 2006; 36:2549.
38. Reddy BM, Patil MK, Rao KN, Reddy GK. J Mol Cat A: Chemical, 2006; 258:302.
39. Rao PS, Vengataratnam RV. Tetrahedron Letters. 1991; 32:5821.
40. Sabhitha G, Reddy BV, Babu RS, Yadav JS. Chemistry Letters. 1998; 27(8):773.
41. Lehnert W. Tetrahedron Letters. 1970; 54:4723.