

An Update on Half-decade Recent Advances in Functionalized Fe₃O₄ Nanoparticles as Heterogeneous Nanocatalysts for the Synthesis of Six-Membered Compounds Containing Nitrogen: A Mini-Review

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ABSTRACT

Over the past half-decade, the functionalized Fe₃O₄ nanoparticles have been applied to organic synthesis as heterogeneous nanocatalysts. Due to the magnetic properties, the functionalized Fe₃O₄ nanoparticles have been preferred over other catalytic systems. In addition, the unique physical, chemical, and magnetic field properties of Fe₃O₄ nanoparticles have led to the development of them for efficient organic synthesis as heterogeneous catalytic systems. Over the past half-decade, all the researchers tried to design and develop different functionalized Fe₃O₄ nanoparticles to perform the organic transformation. In this mini-review, different methodologies for the synthesis of six-membered compounds (heterocyclic compounds) containing nitrogen catalyzed by functionalized Fe₃O₄ nanoparticles have been reviewed. This short review aims to comprehensively investigate the use of functionalized Fe₃O₄ nanoparticles in the synthesis of six-membered compounds (heterocyclic compounds) containing nitrogen with emphasis on their reusability of the catalysts and magnetic field of them in separation from the organic reaction medium.

Keywords: Fe₃O₄ nanoparticles, heterocyclic compounds, magnetic, nanocatalyst, synthesis, reusability

1. Introduction

Since the first research paper in 1916 titled “The oxides of iron. I. Solid solution in the system Fe₂O₃-Fe₃O₄” by Sosman and Hostetter, the Fe₃O₄ topic has grown into the main research area [1]. In this short review, we aim to present a wide overview of the functionalized Fe₃O₄ nanoparticles as a recoverable and magnetically reusable nanocatalyst. In particular, the main focus of this study is on the use of functionalized Fe₃O₄ nanoparticles in the synthesis of six-membered compounds (heterocyclic compounds) containing nitrogen such as pyridobenzoazepines, 2-amino-3-

cyanopyridine, pyrido[2,1-a]isoquinolines, pyrido[1,2-a]quinolones, and so on. 36549 articles were found in a search on 31st March 2022 on the Scopus website for the term “Fe₃O₄” in articles title, abstract, and keywords. In **Fig. 1** the different applications of Fe₃O₄ in various fields of sciences are presented. In addition, there is a growing number of published original articles on Fe₃O₄ nanoparticles in organic synthesis. To the best of our knowledge, this is the first review paper about the application of functionalized Fe₃O₄ nanoparticles in the synthesis of six-membered compounds (heterocyclic compounds) containing nitrogen. Therefore, we decided to start a review paper on this matter.

Pharmaceutical and medicinal chemistry are interesting filed in organic synthetic chemistry for the preparation

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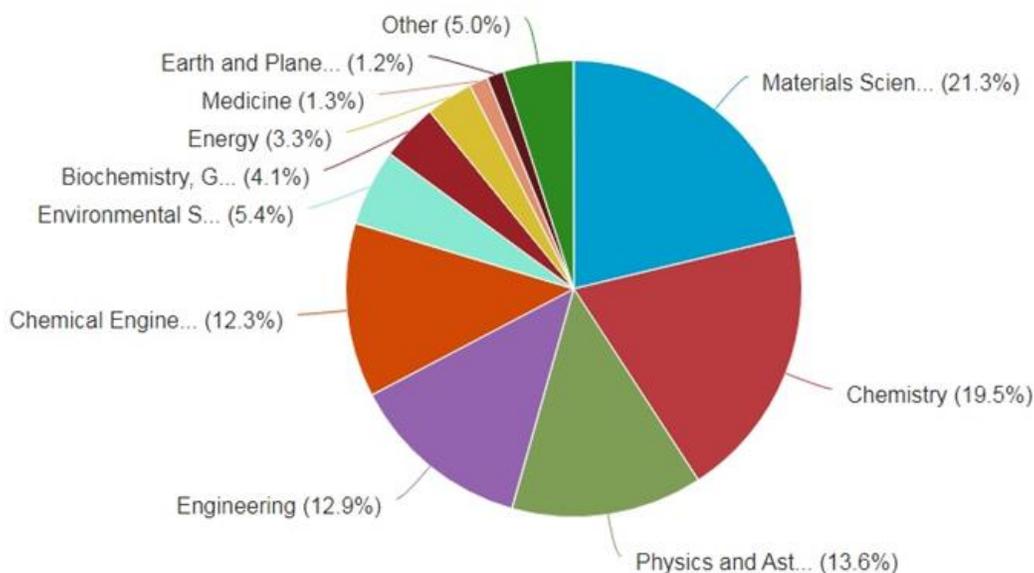


Fig. 1. Application of Fe_3O_4 nanoparticles in different fields of sciences

of different biologically active compounds dealing with drug effectiveness. In addition, nitrogen heterocyclic compounds especially six-membered rings more attention in scientific research papers [2-11]. The heterocyclic six-membered ring compounds are widely spread in the drug and natural compounds. The chemical structure of some of these compounds is shown in Fig 2.

2. Synthesis of Six-Membered Heterocyclic Compound Containing Nitrogen Catalyzed by Functionalized Fe_3O_4 Nanoparticles

In 2022, Savari and co-workers described the synthesis of new compounds of pyridobenzoazepines in the presence of $\text{Ag-Fe}_3\text{O}_4\text{-TiO}_2\text{-MWCNTs}$ at room temperature in the water as a green solvent [12] (Scheme 1). In this research, the $\text{Ag-Fe}_3\text{O}_4\text{-TiO}_2\text{-MWCNTs}$ nanocomposites were prepared via biology and green route using *S. oleracea* bacteria. At first, they synthesized $\text{Ag-Fe}_3\text{O}_4\text{-TiO}_2$ using AgNO_3 , FeCl_3 , and FeCl_3 in the presence of *S. oleracea* as a green reducing agent to reduce Ag (I) to Ag nanoparticles. Subsequently, 0.1 g of multi-walled carbon nanotubes (MWCNTs) was added to the $\text{Ag-Fe}_3\text{O}_4\text{-TiO}_2$ under ultrasound irradiation. Then, the prepared composite was heated at 500 °C for 45 min and washed with the mixture of water and ethanol to obtain pure $\text{Ag-Fe}_3\text{O}_4\text{-TiO}_2\text{-MWCNTs}$ nanocomposites. The interesting characteristic of the pointed catalyst was its attraction to an external magnet. After preparation of the catalyst, to synthesize pyridobenzoazepines, a mixture of isatoic anhydride **1** (2 mmol), *N*-methylimidazole **2** (2 mmol),

activated acetylinic **3** and **4** (2 mmol) in the presence of 0.015 g of $\text{Ag-Fe}_3\text{O}_4\text{-TiO}_2\text{-MWCNTs}$ nanocomposites for 15 min. Then, 2 mmol of α -haloketone **5** and ammonium acetate **6** were added and stirred for 20 other min. Finally, 2 mmol of 2-methyloxirane (epoxide) **7** was added and stirred in water for 1 h to obtain corresponding compounds **8** and **9** in 83-96% yield. If R in compound **4** was hydrogen the compound **8** could be formed. In the current study, the $\text{Ag-Fe}_3\text{O}_4\text{-TiO}_2\text{-MWCNTs}$ nanocomposite shows reusability for seven catalytic cycles in 92% to 87% yield. Notably, some of the synthesized pyridobenzoazepine derivatives show antioxidant activity towards DPPH radical trapping through the FRAP method. The good to excellent yield, moderate reaction time, and high atomic economic, as well as ambient temperature are some of the worthwhile advantages of the presented method.

In 2021, Motahari and Khazaei reported an efficient approach for the synthesis of 2-amino-3-cyanopyridine derivatives **13** using $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-Si}(\text{CH}_2)_3\text{NH-CC-Im-Co}$ (II) nanoparticles at 100 °C under solvent-free conditions (Scheme 2) [13]. The 2-amino-3-cyanopyridine derivatives **13** have biological activity including anti-HIV, anti-acid, and anti-tumor. They prepared the $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-Si}(\text{CH}_2)_3\text{NH-CC-Im-Co}$ (II) nanoparticles through the five sequence steps involving (i): preparation of $\text{Fe}_3\text{O}_4\text{-SiO}_2$ (ii): supporting 3-aminopropyltriethoxy silane on $\text{Fe}_3\text{O}_4\text{-SiO}_2$ (iii): addition of cyanuric chloride to obtain $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-Si}(\text{CH}_2)_3\text{NH-CC}$, (iv): addition of imidazole to $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-Si}(\text{CH}_2)_3\text{NH-CC}$ to afford $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-Si}(\text{CH}_2)_3\text{NH-CC-Im}$, and (v): using $\text{Co}(\text{OAc})_2$ as

precarious source of cobalt to form $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-Si-(CH}_2)_3\text{NH-CC-Im-Co (II)}$ nanoparticles. The chemical representation of the catalyst was shown in **Fig. 3**. After the preparation of the pointed catalyst, they used a mixture of 1 mmol of aryl aldehydes **10**, 1 mmol of acetophenones **11**, 1 mmol of malononitrile **12**, and 1.25

mmol of ammonium acetate **6** in the presence of 0.05 g of $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-Si-(CH}_2)_3\text{NH-CC-Im-Co (II)}$ at 100 °C under solvent-free conditions. The $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-Si-(CH}_2)_3\text{NH-CC-Im-Co (II)}$ show reusability for eight consequence catalytic cycles.

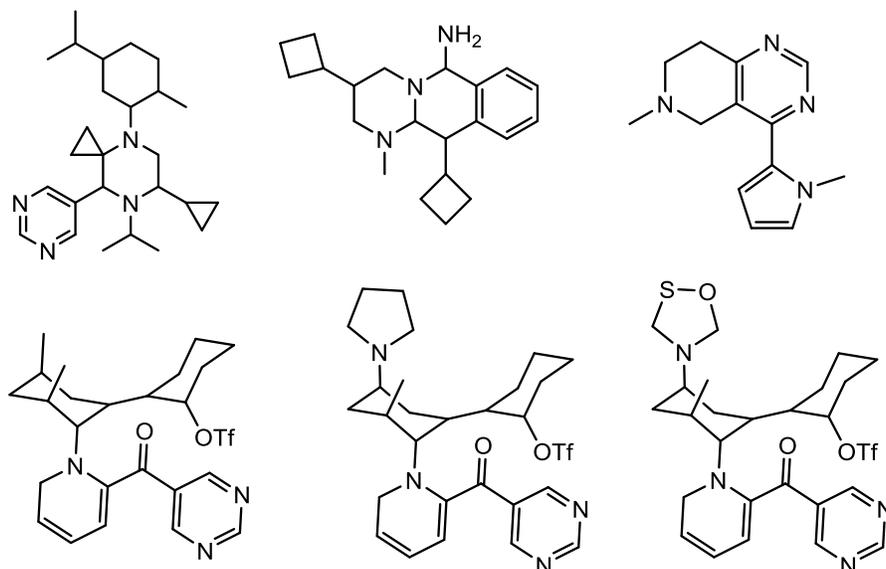
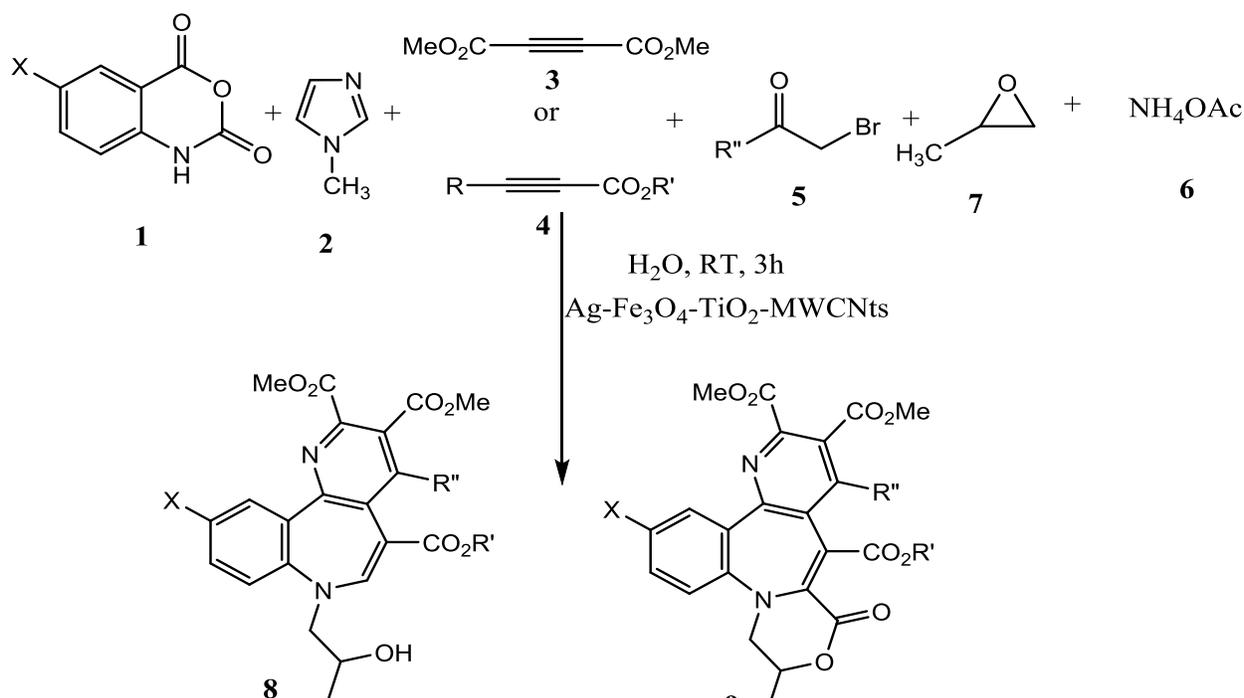


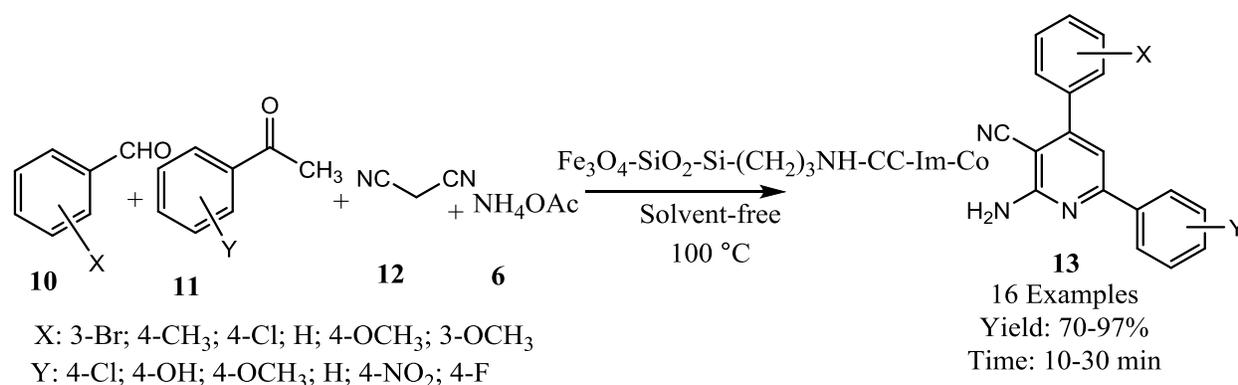
Fig. 2. Some of the biologically active of six-membered nitrogen compounds



R: H, CO_2Et , CO_2Me
 R': CH_3 , C_2H_5
 R'': 4- OCH_3 -Ph, 4- CH_3 -Ph, 4- NO_2 -Ph, CO_2Et ,
 X: H, Me, NO_2 , Cl

11 Examples
 Yield: 83-96%
 Time: 3 h

Scheme 1. Synthesis of pyridobenzazepine derivatives using $\text{Ag-Fe}_3\text{O}_4\text{-TiO}_2\text{-MWCNTs}$ nanocomposites



Scheme 2. Synthesis of 2-amino-3-cyanopyridine derivatives using Fe₃O₄-SiO₂-Si-(CH₂)₃NH-CC-Im-Co (II) nanoparticles

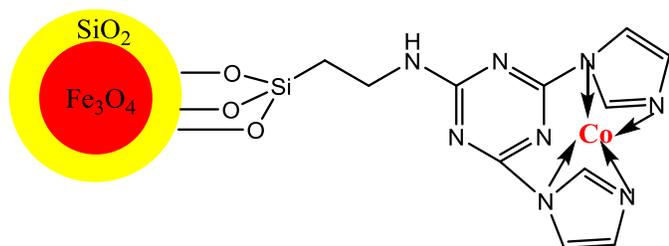
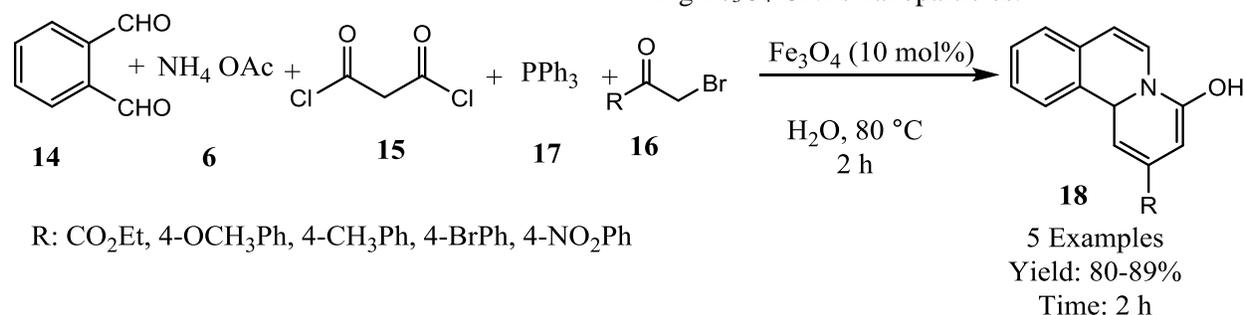


Fig. 3. The schematic of Fe₃O₄-SiO₂-Si-(CH₂)₃NH-CC-Im-Co (II) nanoparticles.

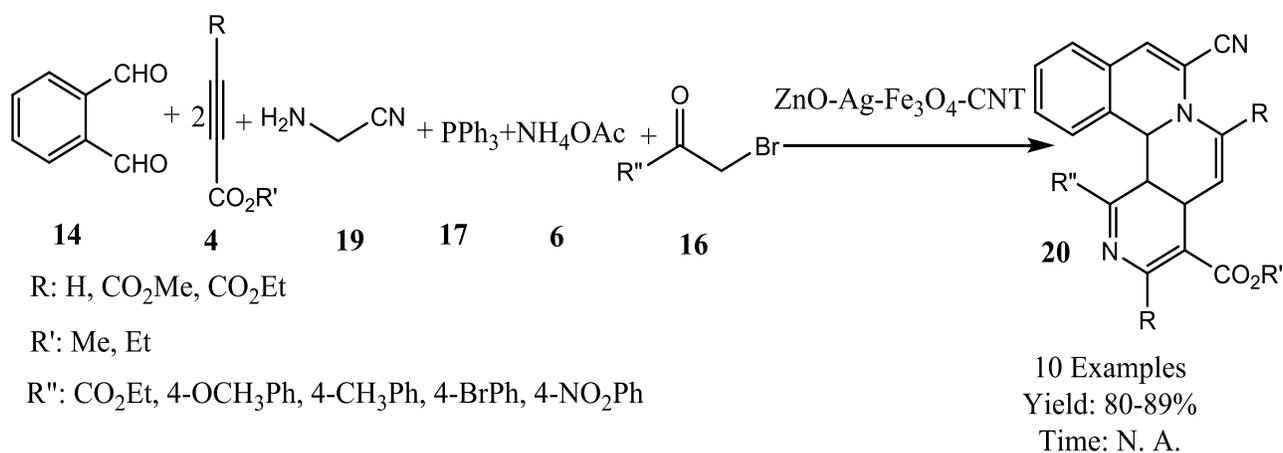
In 2021, Ezzatzadeh and co-workers described a green method for the synthesis of pyrido[2,1-a]isoquinolines and pyrido[1,2-a]quinolones **18** using bio-Fe₃O₄ nanoparticles (**Scheme 3**) [14]. In this multicomponent reaction, in the mixture of 2 mmol of phthalaldehyde **14**, 2 mmol of ammonium acetate **6**, 2 mmol of malonyl chloride **15**, α -bromoketone **16**, and triphenylphosphine **17** the final products **18** were obtained in good to excellent yields within 2 h. The optimum reaction conditions were 10 mol% of Fe₃O₄ nanoparticles, water as the solvent, and 80 °C. Notably, they synthesized Fe₃O₄ nanoparticles via the reduction of ferric chloride solution using Clover leaf water. In addition, they investigated some of the antioxidant activity of the prepared pyrido[2,1-a]isoquinolines and pyrido[1,2-a]quinolones using DPPH trapping in the presence of TBHQ and BHT as antioxidant references. Moreover, the antimicrobial activity of pyrido[2,1-a]isoquinolines and pyrido[1,2-a]quinolones were checked and the



Scheme 3. Synthesis of pyrido[2,1-a]isoquinolines and pyrido[1,2-a]quinolones using bio-Fe₃O₄ nanoparticles

obtained results showed that some of the pointed compounds were active in the diffusion test on Gram-positive and Gram-negative bacteria.

In 2021, Hossaini and co-workers developed a new strategy for the synthesis of naphthyridines **20** using the mixture of phthalaldehyde **14**, 2-aminoacetonitrile **19**, activated acetylinic compounds **4**, triphenyl phosphine **17**, ammonium acetate **6**, and α -bromoketone **16** in the presence of ZnO-Ag-Fe₃O₄-CNTs nanoparticles in aqueous media (**Scheme 4**) [15]. In this research, they investigated antioxidant properties of some of the synthesized compounds by DPPH radical trapping. In addition, the antimicrobial activity of the synthesized compounds was checked. Their results showed that the synthesized compounds have high biological activity such as antioxidant and antimicrobial activity. They prepared the ZnO-Ag-Fe₃O₄-CNTs nanoparticles using the following procedure: First, 0.5 g of AgNO₃, 1 g of Zn(OAc)₂, and 1.5 g of FeCl₂·4H₂O were added to 100 mL of *Petasites hybridus rhizome* water extract to afford the colloids of AgOH, Zn(OH)₂, and Fe(OH)₂ at ambient temperature. Then, the pointed colloids were heated to 350 °C for 2 h to obtain brown solid ZnO-Ag-Fe₃O₄. In continue, 0.1 g of ZnO-Ag-Fe₃O₄ and 0.1 g of carbon nanotubes (CNTs) were mixed in 100 mL of *P. hybridus rhizome* water extract under ultrasound irradiation at 100 °C. Then, the obtained black solid was washed several times with water and ethanol to produce ZnO-Ag-Fe₃O₄-CNTs nanoparticles.

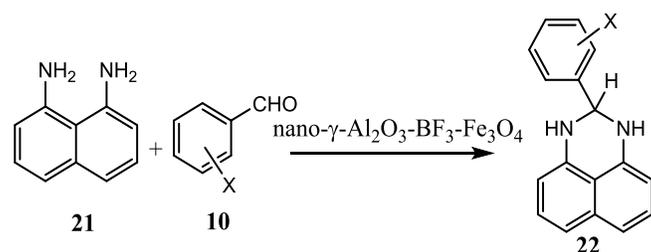


Scheme 4. Synthesis of naphthyrindines in the presence of ZnO-Ag-Fe₃O₄-CNTs nanoparticles

In 2021, Mazoochi and co-workers established different new methods for the synthesis of 2,3-dihydroperimidines **22** in the presence of nano- γ -Al₂O₃-BF₃-Fe₃O₄ nanoparticles under grinding, reflux, microwave, and ultrasound conditions (**Scheme 5**) [16]. The 2,3-dihydroperimidines **22** were synthesized using the reaction of naphthalene-1,8-diamine **21** and aryl aldehydes **10** under grinding, reflux, microwave, and ultrasound irradiations. The catalytic amount of nano- γ -Al₂O₃-BF₃-Fe₃O₄ and nano- γ -Al₂O₃-BF₃ for grinding, reflux, microwave, and ultrasound conditions were (0.06, and 0.12 g), (0.06, and 0.06 g), (0.06, and 0.03 g), and (0.03, and 0.03 g), respectively. They investigated the chemical structure of the synthesized compounds using ¹H NMR, ¹³C NMR, and melting point. Their results show that ultrasound irradiation was the best reaction condition due to the reaction time and yield of the final products for the synthesis of 2,3-dihydroperimidines **22**. In addition, the reusability of the catalyst was checked and the pointed catalyst showed five consequence catalytic cycles.

In 2020, Moussa and Rahmati designed a new catalyst (Fe₃O₄-SiO₂-TEAPOM) for the synthesis of tetrahydrobenzimidazo[2,1-b]quinazolin-1(2H)-ones **26** via a multicomponent reaction including 2-aminobenzazoles **23**, benzaldehyde **24**, and cyclic β -diketone **25** in ethanol at 70-75 °C to obtain tetrahydrobenzimidazo[2,1-b]quinazolin-1(2H)-ones **26** in moderate to excellent yields (**Scheme 6**) [17]. The chemical structure of Fe₃O₄-SiO₂-TEAPOM was shown in **Fig. 4**. The pointed catalyst was characterized by FT-IR, TGA, SEM, TEM, EDX, and VSM techniques. Due to the presence of polyoxometalate, the catalytic of Fe₃O₄-SiO₂-TEAPOM was superior in the synthesis of tetrahydrobenzimidazo[2,1-b]quinazolin-1(2H)-ones **26**. Interestingly, the pointed catalyst showed excellent reusability up to ten catalytic cycles. Easy separation of

the Fe₃O₄-SiO₂-TEAPOM using an external magnetic field, and green solvent were other worthwhile advantages of the presented work.



X: H, 2-NO₂, 3-NO₂, 4-COOH, 4-CH₃, 4-Cl, 2,4-Cl, 2,6-Cl, 2-OH, 3-OH, 2,4-OH, 2-OCH₃, 3,4-OCH₃, 2-OH-3-OCH₃, 4-OH-3-OEt

15 Examples

Grinding: Yield: 69-82% Reflux: Yield: 69-84%
 Time: 20-45 min Time: 20-40 min

US: Yield: 74-95% MW: Yield: 70-92%
 Time: 3-5 min Time: 4-6 min

Scheme 5. Synthesis of 2,3-dihydroperimidines in the presence of nano- γ -Al₂O₃-BF₃-Fe₃O₄ nanoparticles under grinding, reflux, microwave (MW), and ultrasound (US) conditions.

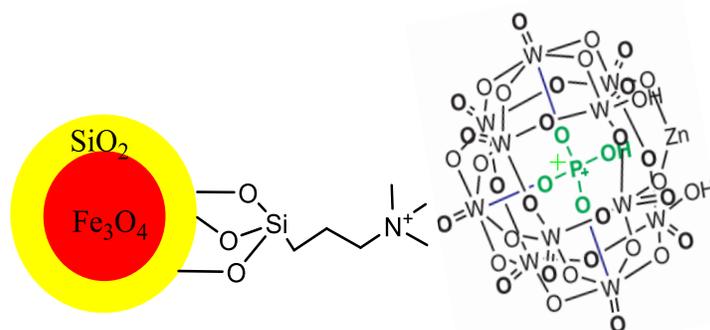
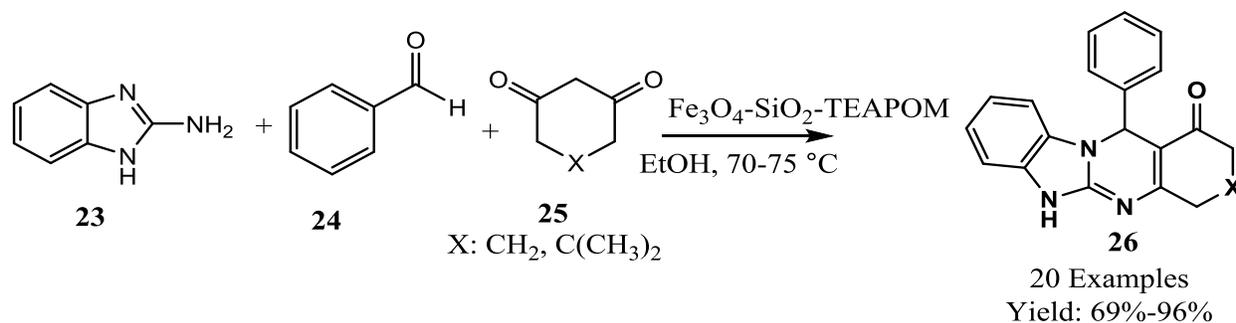


Fig. 4. The schematic representation of Fe₃O₄-SiO₂-TEAPOM



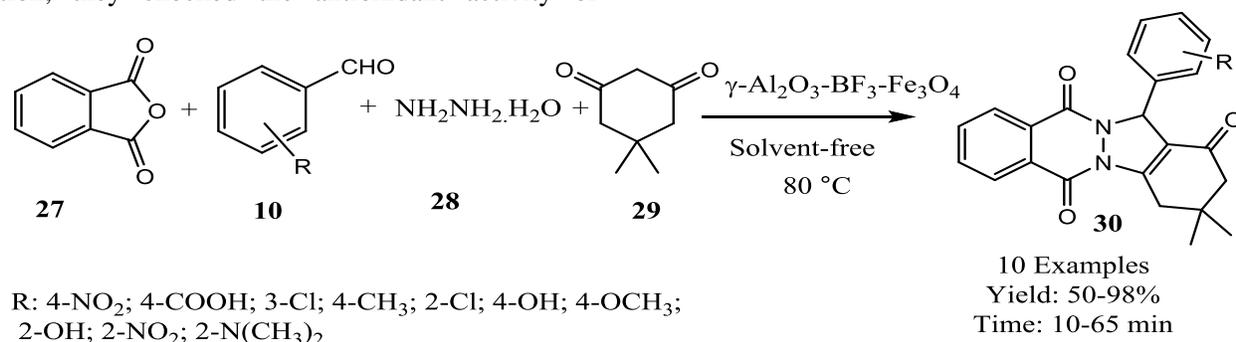
Scheme 6. Synthesis of tetrahydrobenzimidazo[2,1-b]quinazolin-1(2H)-ones using $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-TEAPOM}$

In 2020, Chegeni and co-workers described the synthesis of 2H-indazolo[2,1-b]phthalazine-triones **30** using nano- $\gamma\text{-Al}_2\text{O}_3\text{-BF}_3\text{-Fe}_3\text{O}_4$ nanoparticles at 80 °C under solvent-free conditions (**Scheme 7**) [18]. The 2H-indazolo[2,1-b]phthalazine-triones **30** was prepared through the reaction of phthalicanhydride **27**, aryl aldehyde **10**, hydrazine hydrate **28**, and dimedone **29** to produce 10 examples of 2H-indazolo[2,1-b]phthalazine-triones **30** in moderate to excellent yields within 10 to 56 min. The pointed catalyst could be reused for five consequence catalytic cycles with decreasing yield from 98% to 65% vs. increasing the reaction time from 12 min to 30 min. In addition, they calculated the HOMO-LUMO band gap ($\Delta E_{\text{HOMO-LUMO}}$) of the T1, T2 conformers of one of the synthesized compounds.

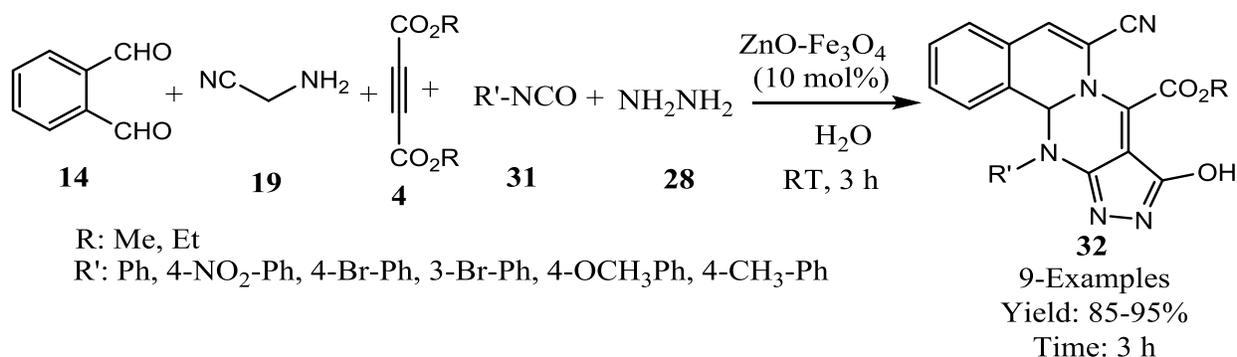
In 2020, Goochibeigi and co-workers promoted a green approach for the synthesis of pyrazolo pyrimidinone derivatives **32** in the presence of 10 mol% of $\text{ZnO-Fe}_3\text{O}_4$ nanoparticles at room temperature in water for 3 h (**Scheme 8**) [19]. In this regard, they prepared the pyrazolo pyrimidinone derivatives using a five component reactions including 2 mmol of phthaldehyde **14**, 2 mmol of cyano methylamine **19**, 2 mmol of dialkyl acetylene dicarboxylate **4**, and 2 mmol of isocyanate **31**, as well as hydrazine **28** in water using $\text{ZnO-Fe}_3\text{O}_4$ nanoparticles as a robust heterogeneous nanocatalyst. They reported 9 examples of pyrazolo pyrimidinone derivatives in good to excellent yields for 3 h. In addition, they checked the antioxidant activity of

pyrazolo pyrimidinone and their results showed that the mentioned compounds have a great potential antioxidant activity towards DPPH radical trapping.

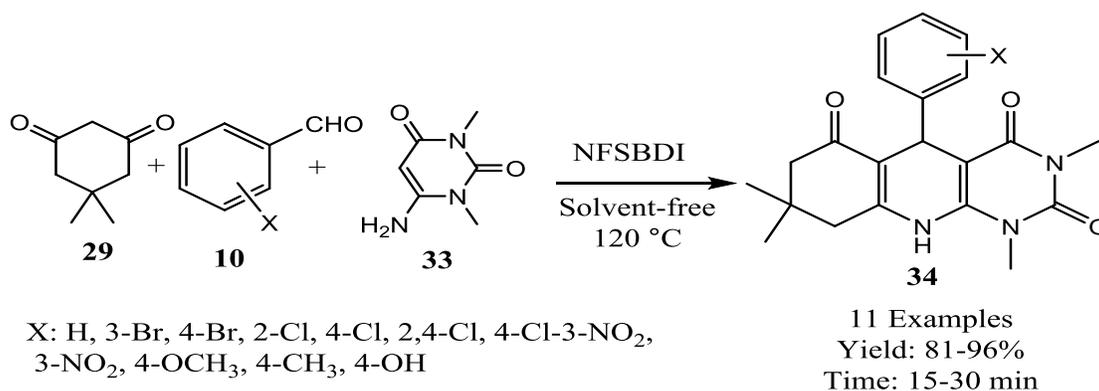
In 2020, Zare and Barzegar designed a new and fantastic heterogeneous catalyst for the synthesis of pyrimido[4,5-b]quinolones using NFSBDI as dicationic ionic liquid grafted with silica-coated nanoparticles of Fe_3O_4 (**Scheme 9**) [20]. The chemical structure of NFSBDI was shown in **Fig. 5**. They prepared NFSBDI using five consequences steps including (i) Fe_3O_4 preparation (ii), SiO_2 coated on Fe_3O_4 to afford $\text{Fe}_3\text{O}_4\text{-SiO}_2$, (iii) anchored of 3-chloropropyltrimethoxysilane on $\text{Fe}_3\text{O}_4\text{-SiO}_2$ to afford $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-SiPrCl}$, (iv) reaction of N,N,N',N' -tetramethylenediamine to obtain $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-SiPrN}(\text{CH}_3)_2\text{CH}_2\text{CH}_2\text{N}(\text{CH}_3)_2$, (iv) treatment of H_2SO_4 to corresponding dicationic ionic liquid grafted with silica-coated nanoparticles of Fe_3O_4 (NFSBDI). After preparation and characterization of NFSBDI as a heterogeneous catalyst, they synthesized some of the pyrimido[4,5-b]quinolones **34**, using the reaction of 1 mmol of dimedone **29**, 1 mmol of aldehyde **10**, and 1 mmol of 6-amino-1,3-dimethyluracil **33** in the presence of 0.048 g of NFSBDI to obtain pyrimido[4,5-b]quinolones **34** in good to excellent yields within 15-30 min. They characterized and approved the NFSBDI using FT-IR, FE-SEM, VSM, XRD, TGA, and EDS techniques. In addition, they investigated the reusability and recyclability of the catalyst and their results showed that the NFSBDI could be reused for four runs without loss of their catalytic activity.



Scheme 7. Synthesis of 2H-indazolo[2,1-b]phthalazine-triones using nano- $\gamma\text{-Al}_2\text{O}_3\text{-BF}_3\text{-Fe}_3\text{O}_4$ nanoparticles



Scheme 8. Synthesis of pyrazolo pyrimidinone derivatives using ZnO-Fe₃O₄ nanoparticles



Scheme 9. Synthesis of pyrimido[4,5-b]quinolones using NFSBDI

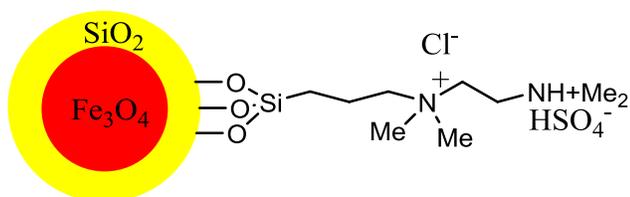
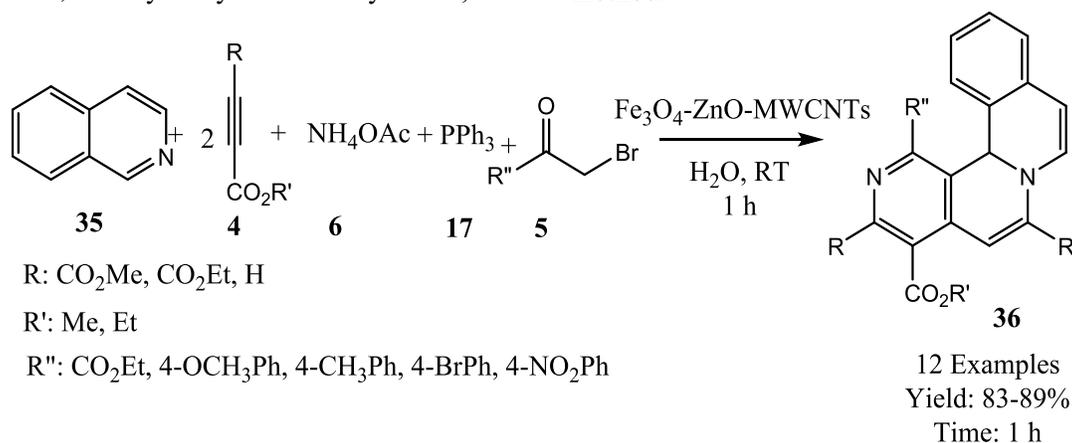


Fig. 5. The chemical structure of NFSBDI

In 2020, Hossaini and co-workers introduced the Fe₃O₄-ZnO-MWCNTs as high magnetic and efficient nanocomposites for the synthesis of naphthyridine **36** through a five-component reaction including isoquinoline **35**, dialkylacetylenedicarboxylate **4**, α -

haloketone **5**, triphenyl phosphine **17**, and ammonium acetate **6** (**Scheme 10**) [21]. They reported that the synthesized naphthyridine derivative has the potential for antimicrobial and anticancer activities. To approve the preparation of Fe₃O₄-ZnO-MWCNTs, they characterized the Fe₃O₄-ZnO-MWCNTs using SEM, TEM, EDX, XRD, and VSM techniques. The prepared catalyst can be reused for seven cycles. The easy separation of the catalyst using an external magnetic field, high yield of products, short reaction times, and biological activity of synthesized compounds were some of the worthwhile advantages of the pointed method.



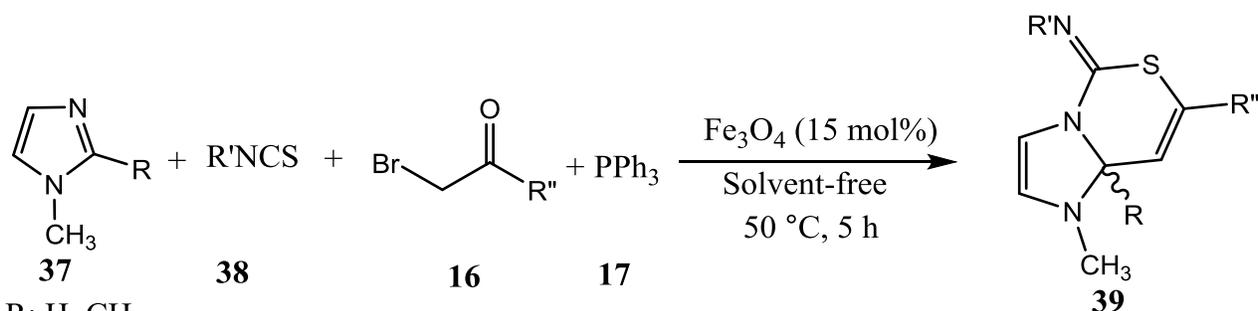
Scheme 10. Synthesis of naphthyridine using Fe₃O₄-ZnO-MWCNTs nanocatalyst

In 2019, Shafaei and Sharafian reported a green synthesis of imidazole derivatives **39** using 15 mol% of Fe₃O₄ nanoparticles at 50 °C for 5 h under solvent-free conditions (**Scheme 11**) [22]. They synthesized the imidazole derivatives **39** using a multicomponent reaction of N-methylimidazole **37**, isothiocyanate **38**, triphenyl phosphine **17**, and α -bromoketone **16**. It is notable that the Fe₃O₄ nanoparticles were synthesized through the green approach using the reduction of ferric chloride solution in the presence of *Clover Leaf* water extract.

In 2019, Hamedian and co-workers described a green method synthesis of pyrido[2,1-*a*]isoquinolines and pyrido[1,2-*a*]quinolins **42** in the presence of 10 mol% of Fe₃O₄ nanoparticles and 10 mol% of sodium hydroxide at 80 °C for 6 h with good yields of the 5 examples of pyrido[2,1-*a*]isoquinolines and pyrido[1,2-*a*]quinolins (**Scheme 12**) [23]. In this work, they used phthalaldehyde **14**, methyl amine **40**, methyl malonyl chloride **41**, triphenyl phosphine **17**, and α -bromoketone **16** to produce pyrido[2,1-*a*]isoquinolines and pyrido[1,2-*a*]quinolins **42**. In addition, they investigated the antioxidant activity of some of the synthesized compounds using DPPH radical trapping in comparison with BHT and TBHQ as references standard.

In China, Ashraf and co-workers reported a new method for the synthesis of polyhydroquinolines **44** in the presence of Fe₃O₄-Schiff base Cu as a heterogeneous catalyst in water (**Scheme 13**) [24]. The prepared catalyst was characterized by FT-IR, XRD, TEM, SEM, VSM, TGA, and ICP techniques. Then, they applied the Fe₃O₄-Schiff base Cu (**Fig. 6**) in the synthesis of polyhydroquinolines **44** using the reaction of 1 mmol of dimedone **29**, 1 mmol of aldehyde **10**, 1.3 mmol of ammonium acetate **6**, and 1 mmol of ethylacetoacetate **43** in the presence of 0.25 mol% of Cu in 3 mL of water under reflux conditions. Interestingly, they observed that the prepared catalyst could be reused for 11 catalytic cycles in the yield of 100% to 86% and turnover frequency of (TOF) (s⁻¹) 5760000-4953600.

In 2019, Kalhor and Zarnegar developed a new multifunctional and magnetic nanocatalyst (Fe₃O₄/SO₃H@zeolite-Y) for the synthesis of perimidine derivatives in ethanol at room temperature (**Scheme 14**) [25]. In this regard, they used different aromatic aldehydes **10**, and 1,8-diaminonaphthalene **21** in the presence of 0.008 g of Fe₃O₄/SO₃H@zeolite-Y in ethanol at room temperature. Their results showed that both electron-withdrawing and electron-donating groups promoted the products in good to excellent yields. Also, the catalyst can be applied for five cycles.



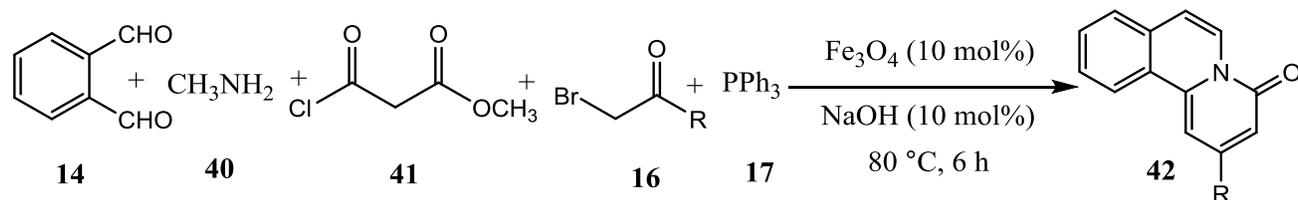
R: H, CH₃

R': Ph, 4-OCH₃-Ph, 4-CH₃-Ph, 4-NO₂-Ph, 4-Br-Ph, 4-CH₃-Ph, n-Bu

R'': CO₂Et, 4-OCH₃-Ph, 4-NO₂-Ph

10 Examples
Yield: 80-97%

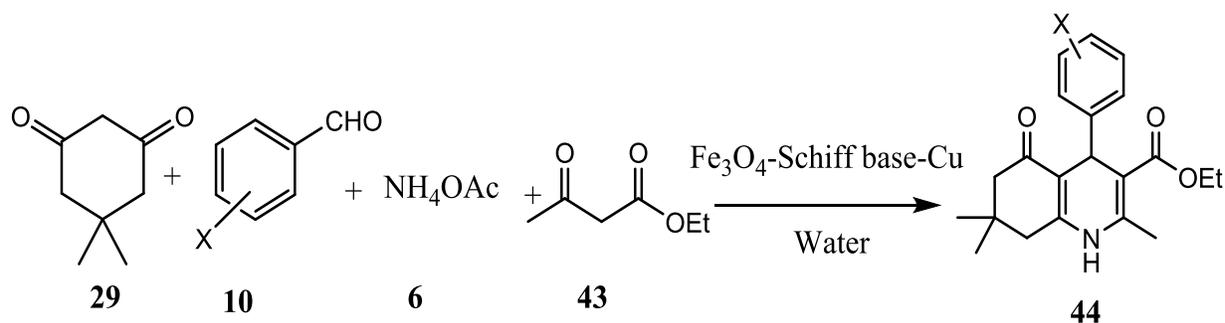
Scheme 11. Synthesis of imidazole derivatives using bio Fe₃O₄ nanoparticles



R: CO₂Et; 4-OCH₃-Ph; 4-CH₃-Ph; 4-Br-Ph; 4-NO₂-Ph

5 Examples
Yield: 80-89%
Time: 6 h

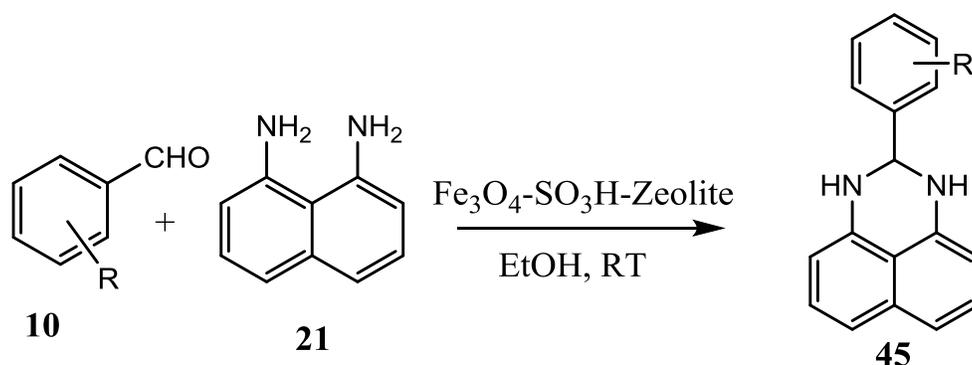
Scheme 12. Synthesis of pyrido[2,1-*a*]isoquinolines and pyrido[1,2-*a*]quinolins using Fe₃O₄ nanoparticles



X: 4-Cl, 4-CH₃, 4-OCH₃, 3,4-Di(OCH₃), 4-CF₃, 4-CN, H, 3-OH, 4-OH, 3,4-Di(OH), 4-Br, 3-Br, 4-F, 4-NO₂, 3-NO₂, 2-NO₂-5-OH, 4-N(CH₃)₂, 4-SCH₃

18 Examples
Yield: 90-100%
Time: 15-40 min

Scheme 13. Synthesis of polyhydroquinolines using Fe₃O₄-Schiff base Cu



R: H, 2-NO₂, 3-NO₂, 4-NO₂, 4-OCH₃, 3,4-OCH₃, 2-OH, 13 Examples
2-OH-5-Br, 2-OH-4-OCH₃, 3-Cl, 4-Cl, 2,3-Cl, 5-NO₂ Yield: 88-98%
Time: 3-6 min

Scheme 14. Synthesis of perimidine derivatives using Fe₃O₄/SO₃H@zeolite-Y

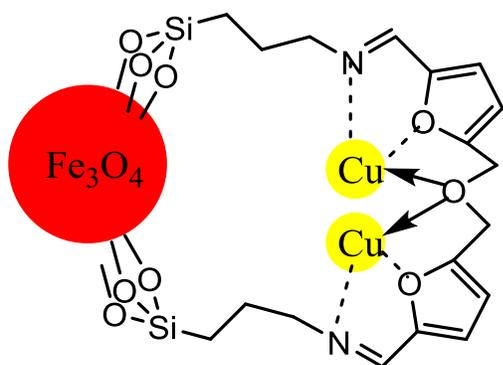
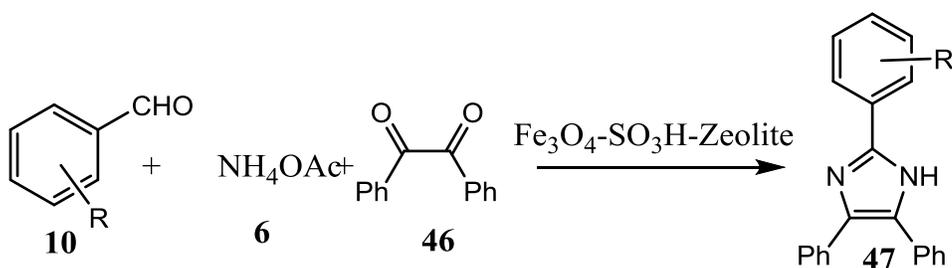


Fig. 6. The chemical structure of Fe₃O₄-Schiff base Cu

Furthermore, the same group reported the synthesis of various three substituted imidazole derivatives **47** via the reaction of 1 mmol of different aromatic aldehydes **10**, 2 mmol of ammonium acetate **6**, and 1 mmol of benzil **46** in the presence of 0.02 g of Fe₃O₄/SO₃H@zeolite-Y in ethanol at 80 °C (**Scheme**

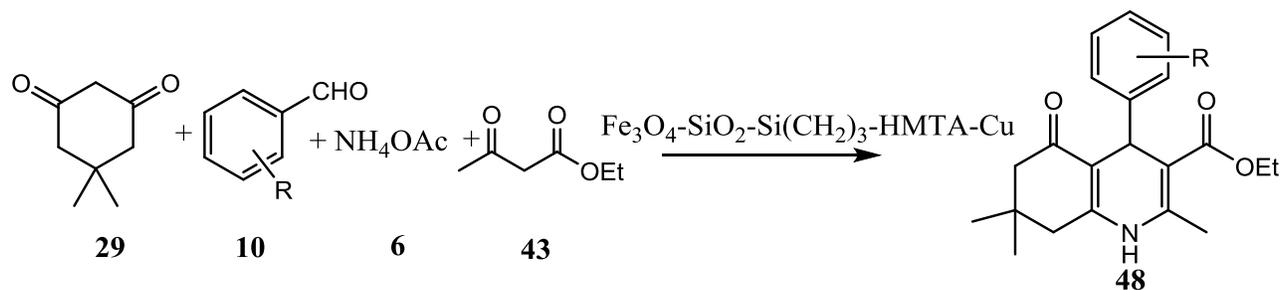
15) [25]. All of the desired three substituted imidazoles **47** were obtained in good to excellent yields within 20-65 min. In addition, the pointed catalyst can be used for five consequence catalytic cycles.

In 2019, the Fe₃O₄-SiO₂-SiPr-HMTA-Cu was prepared by Salem and co-workers for the synthesis of hexahydroquinoline derivatives **48** using the treatment of 1 mmol of dimedone **29**, 1 mmol of aromatic aldehydes **10**, 1.2 mmol of ammonium acetate **6**, and 1 mmol of beta-keto ester **43** in the presence of 0.05 g of Fe₃O₄-SiO₂-SiPr-HMTA-Cu at 60 °C (**Scheme 16**) [26]. They identified the prepared catalyst with different techniques including FT-IR, FE-SEM, TEM, VSM, and TGA. Their result approved that the metal copper was successfully chelated with hexamethylenetetramine (HMTA). The presented method has some advantages including high yield, short reaction times, easy separation of the catalyst and products, and excellent atomic economic conditions.



R: H, 2-NO₂, 3-NO₂, 3-OH-4-OCH₃, 2-OCH₃, 3,4-OCH₃, 3-OH, 4-OCH₃, 2-Cl, 2,3-Cl, 2,4-Cl, 4-Cl, 3-Br, 4-CH₃ 14 Examples
Yield: 85-98%
Time: 20-65 min

Scheme 15. Synthesis of imidazole derivatives using $\text{Fe}_3\text{O}_4/\text{SO}_3\text{H@zeolite-Y}$



R: H, 3-OCH₃, 3-OH, 4-OCH₃, 3-F, 3-NO₂, 2,3-Cl, 2,4-Cl, 4-N(CH₃)₂, 4-Cl, 2-NO₂, 4-CHO, 2-F, 4-CH₃

15 Examples
Yield: 50-95%
Time: 3-15 min

Scheme 16. Synthesis of hexahydroquinoline derivatives using $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-SiPr-HMTA-Cu}$

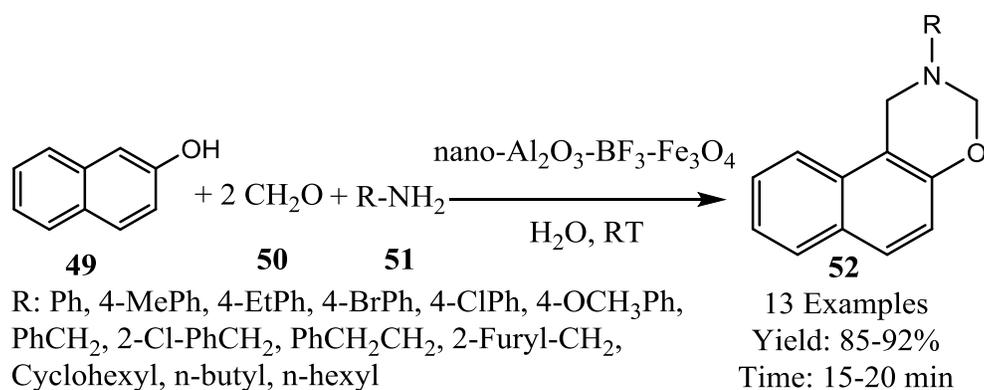
In 2019, Babaei and Mirjalili reported a one-pot synthesis of 1,3-oxazine derivatives **52** using nano- $\text{Al}_2\text{O}_3\text{-BF}_3\text{-Fe}_3\text{O}_4$ as reusable nanocatalyst in water at room temperature (**Scheme 17**) [27]. To synthesize 1,3-oxazine derivatives **52**, they mixed 1 mmol of β -naphthol **49**, 1 mmol of primary amine **50**, and 2.5 mmol of formaldehyde **51** in 1.5 mL of water in the presence of 0.04 g of nano- $\text{Al}_2\text{O}_3\text{-BF}_3\text{-Fe}_3\text{O}_4$. All of the 2-aryl/alkyl-2,3-dihydro-1H-naphtho[1,2-e][1,3]oxazine derivatives **52** were obtained in good to excellent yields within 15-20 min. In addition, they investigated the reusability of the nano- $\text{Al}_2\text{O}_3\text{-BF}_3\text{-Fe}_3\text{O}_4$ and their results showed that the catalyst could be reused for five catalytic cycles. Furthermore, the chemical structure of the synthesized compounds was checked by ¹H NMR and ¹³C NMR, FT-IR spectroscopy, and melting point. Interestingly, they compared their work with other catalysts such as ionic liquid [bmim]HSO₄ (5 mol%), and vitamin B₁ (10 mol%). Their results showed that the nano- $\text{Al}_2\text{O}_3\text{-BF}_3\text{-Fe}_3\text{O}_4$ was better than the other pointed catalysts.

In Iran, Karimiyan and Rostamzadeh reported a green and efficient approach for the synthesis of aryl-substituted imidazo[1,2-a]pyridines and pyrido[1,2-a]pyrimidines **55** using 40 mg of $\text{Fe}_3\text{O}_4\text{-GO-SO}_3\text{H}$ at 90

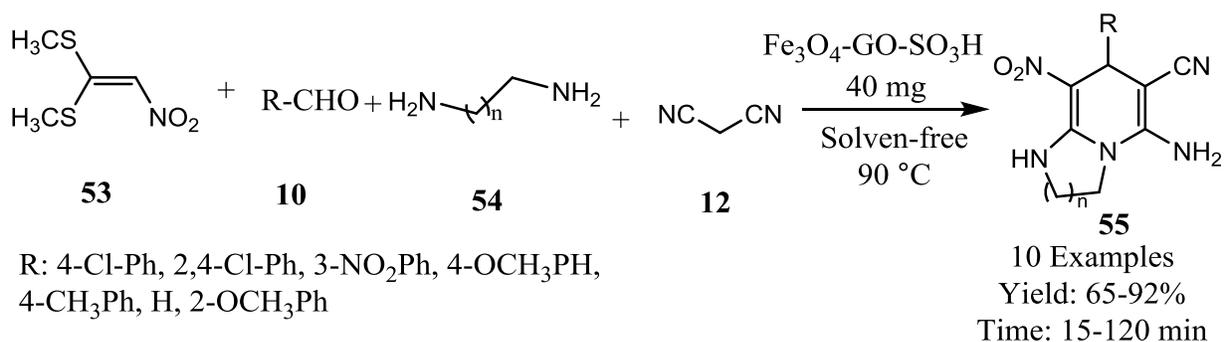
°C under solvent-free conditions (**Scheme 18**) [28]. They used the following starting materials for the production of pyrido[1,2-a]pyrimidines **55** as follows: 1,1-bis(methylthio)-2-nitroethylene **53**, aldehyde **10**, 1,n-diamine **54**, and malononitrile **12**. The final products were produced in moderate to excellent yields within 15-120 min.

In addition, the same group synthesized spiro-oxindoloimidazo[1,2-a]pyridines and -pyrido[1,2-a]pyrimidines **57** using the same previous conditions (**Schemes 19 and 20**) [28].

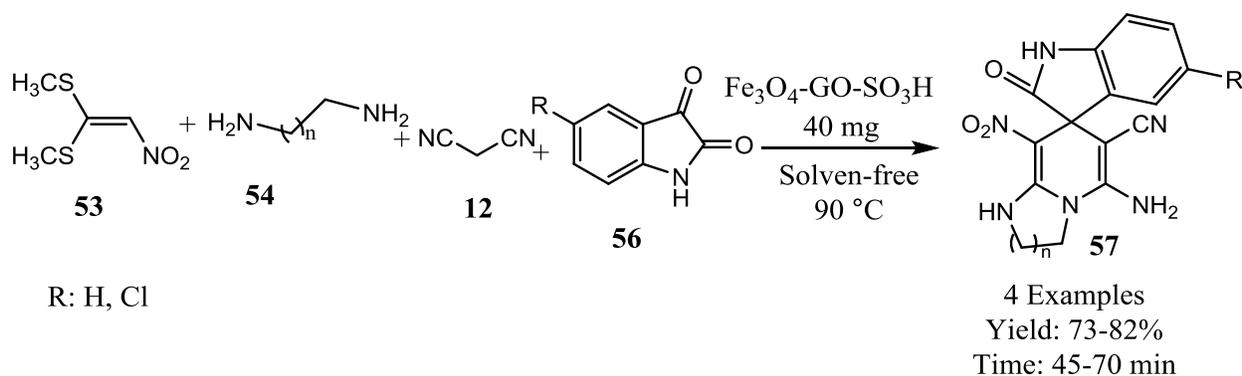
In Korea, Basavegowda and co-workers described an efficient and suitable method for the synthesis of polyfunctionalized pyridines using $\text{Fe}_3\text{O}_4\text{-MWCNTs}$ in water under reflux conditions (**Scheme 21**) [29]. After the preparation of $\text{Fe}_3\text{O}_4\text{-MWCNTs}$, they characterized the pointed catalyst using XRD, FE-SEM, TEM, XPS, TGA, EDX, and VSM. Hence, 1 mmol of ketone **60**, 1 mmol of cinnamaldehyde **61**, and 3 equivalents of ammonium acetate **6** in the presence of 5.0 mg of $\text{Fe}_3\text{O}_4\text{-MWCNTs}$ were used to synthesize polyfunctionalized pyridines **62**. The pointed catalyst showed 5 consequence catalytic cycles in 85% to 83% yields.



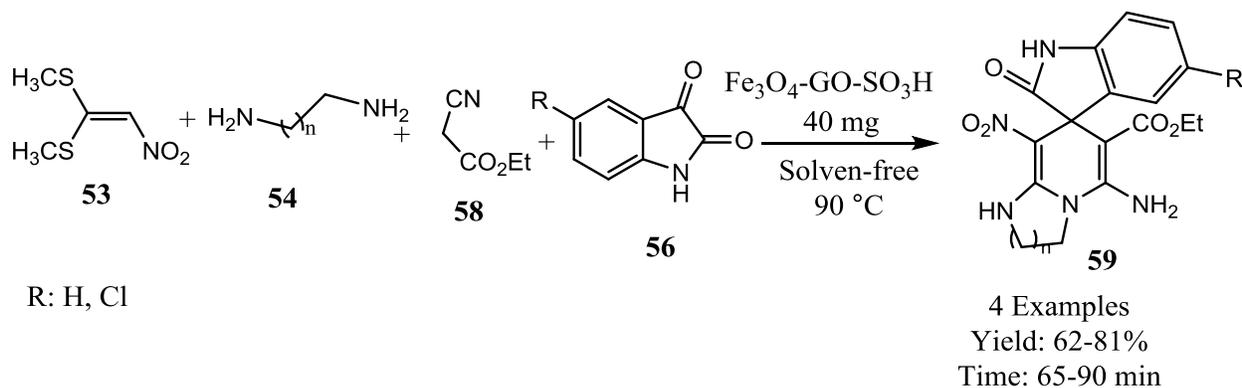
Scheme 17. Synthesis of 2-aryl/alkyl-2,3-dihydro-1H-naphtho[1,2-e][1,3]oxazine derivatives using nano-Al₂O₃-BF₃-Fe₃O₄



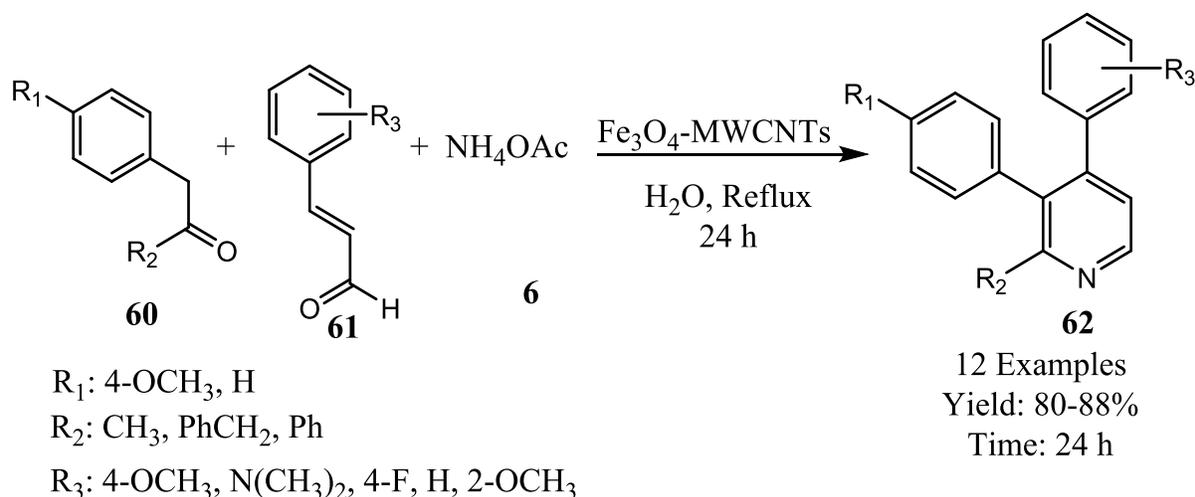
Scheme 18. Synthesis of pyrido[1,2-a]pyrimidines using Fe₃O₄-GO-SO₃H



Scheme 19. Synthesis of spiro-oxindolo-imidazo[1,2-a]pyridines and -pyrido[1,2-a]pyrimidines using Fe₃O₄-GO-SO₃H



Scheme 20. Synthesis of spiro-oxindolo-imidazo[1,2-a]pyridines and -pyrido[1,2-a]pyrimidines using Fe₃O₄-GO-SO₃H

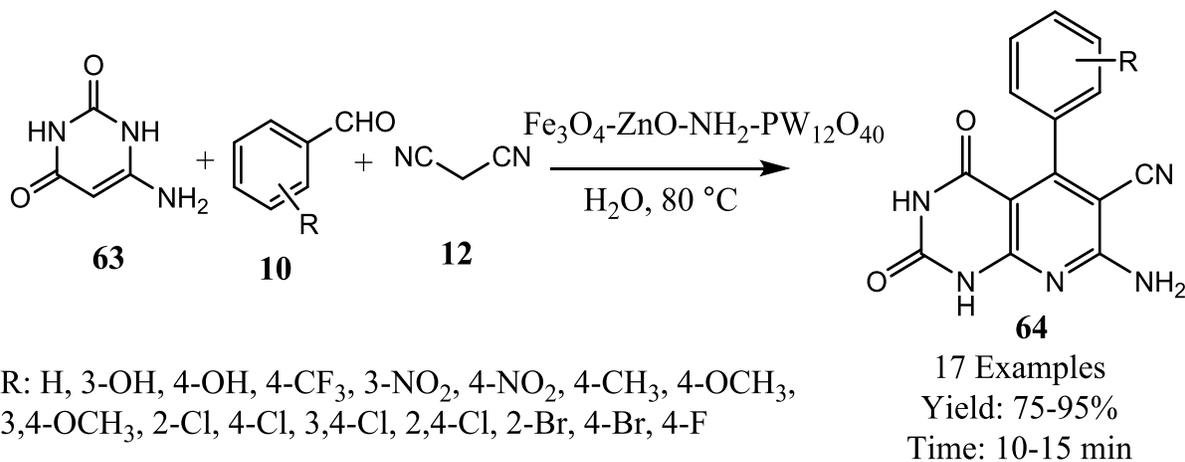


Scheme 21. Synthesis of polyfunctionalized pyridines using Fe₃O₄-MWCNTs nanocomposites

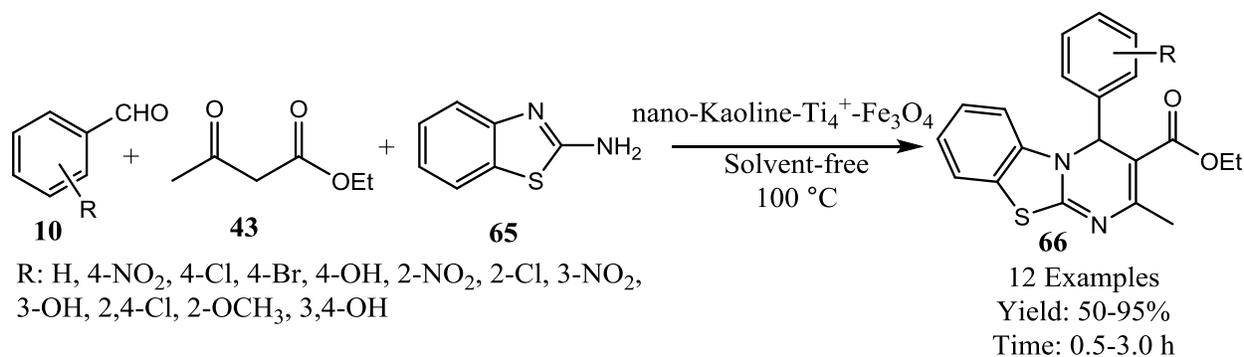
In 2019, Farokhian and co-workers reported an efficient method for the synthesis of pyrido[2,3-*d*]pyrimidine derivatives **64** using Fe₃O₄-ZnO-NH₂-PW₁₂O₄₀ at 80 °C in water (**Scheme 22**) [30]. The catalyst was characterized by FT-IR, XRD, VSM, SEM, EDX, and TEM analysis.

Mirjalili and Soltani developed a new method for the synthesis of pyrimido[2,1-*b*]benzothiazoles **66** using 1 mmol of aryl aldehydes **10**, 1 mmol of ethyl acetoacetate **43**, and 1 mmol of 2-aminobenzothiazole **65** in the presence of 0.03 g nano-Kaoline-Ti⁴⁺-Fe₃O₄ at 100 °C under solvent-free conditions (**Scheme 23**) [31]. They characterized the nano-Kaoline-Ti⁴⁺-Fe₃O₄ by use of FE-SEM, TEM, XRD, EDX, VSM, and TGA techniques. The pointed catalyst could be reused for 7 catalytic cycles without loss of their catalytic activity.

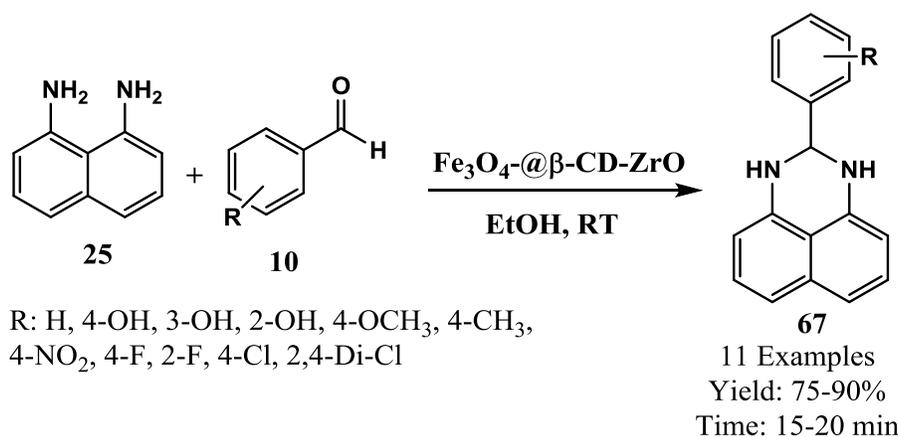
In 2018, Amorllahi and Vahidnia reported the synthesis of 2,3-dihydro-1H-perimidine derivatives **67** using a new heterogeneous nanocatalyst naming decoration of β-CD-ZrO supported on Fe₃O₄ nanoparticles (**Scheme 24**) [32]. The Fe₃O₄-@β-CD-ZrO nano-catalyst showed reusability for the continuous catalytic cycles. They used 1,8-diaminonaphthalene **25** in the treatment of different aromatic aldehydes **10** in the presence of Fe₃O₄-@β-CD-ZrO to produce 2,3-dihydro-1H-perimidine derivatives. The schematic representation of the prepared catalyst was shown in **Fig. 7**. In this study, for the characterization of Fe₃O₄-@β-CD-ZrO, the authors applied different techniques including TGA, XRD, SEM, TEM, and FT-IR techniques. The presented methodology has several worthwhile advantages including easy separation of the final product from the catalyst, reusability of the catalyst, high yields of the final products, and green solvent of the reaction.



Scheme 22. Synthesis of pyrido[2,3-*d*]pyrimidine derivatives using Fe₃O₄-ZnO-NH₂-PW₁₂O₄₀



Scheme 23. Synthesis of pyrimido[2,1-b] benzothiazoles using nano-Kaoline-Ti⁴⁺-Fe₃O₄



Scheme 24. Synthesis of 2,3-dihydro-1H-perimidine derivatives using Fe₃O₄-@β-CD-ZrO

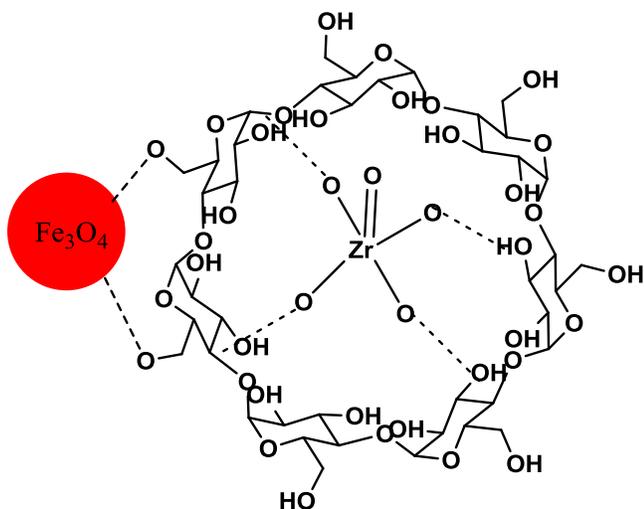


Fig. 7. The schematic representation of Fe₃O₄-@β-CD-ZrO

In a separate study, Babaei and Mirjalili introduced a one-pot method for the synthesis of tetrahydropyridines **70** using nano Al₂O₃-BF₃-Fe₃O₄ (**Scheme 25**) [33]. They characterized the prepared nanocatalyst using TEM, VSM, XRD, XRF, BET, FESEM, and FT-IR spectroscopy. They used 2 mmol of aniline derivatives **68** and 1 mmol of ethyl acetoacetate **69** in the presence of 0.03 g of Al₂O₃-BF₃-Fe₃O₄ for 3-4 h at 80 °C under solvent-free conditions. They prepared the Al₂O₃-BF₃-

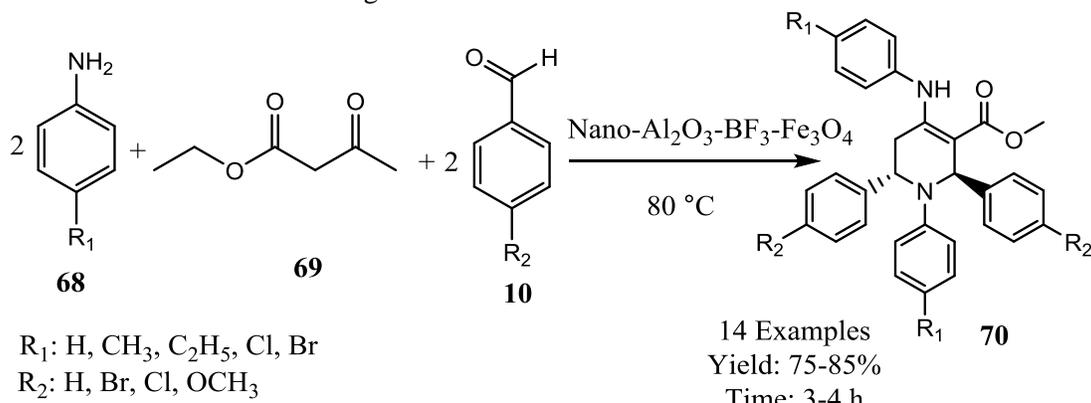
Fe₃O₄ as follows procedure. 1.0 g of γ-Al₂O₃, and 10 mL of dichloromethane were mixed and 0.5 mL of BF₃ was added drop-wise. In this step, the nano- Al₂O₃-BF₃ was obtained. Finally, for the synthesis of Al₂O₃-BF₃-Fe₃O₄, 1.0 g of nano- Al₂O₃-BF₃ was added to 1.0 g of Fe₃O₄ nanoparticles in 20 mL of dichloromethane and stirred for 1 h. They compared the prepared catalyst with other catalysts such as ZrCl₄, ZrOCl₂.8H₂O, Fe(NO₃)₃.9H₂O, FeCl₃/SiO₂ nanoparticles, acetic acid, CAN, PTSA, and L-proline/THF. Their results show that the nano Al₂O₃-BF₃-Fe₃O₄ was superior in terms of yield and time of the reaction.

Dehbalaei and co-workers reported a novel method for the synthesis of 1,8-dioxo-decahydroacridine derivatives **71** using Fe₃O₄-SiO₂-N-propylbenzoguanamine-SO₃H nanocomposite (**Schemes 26** and **27**) [34]. For the preparation of 1,8-dioxo-decahydroacridine derivatives **71** using Fe₃O₄-SiO₂-N-propylbenzoguanamine-SO₃H nanocomposite, 2 mmol of dimedone **25**, 1 mmol of aromatic aldehydes **10**, and 1.2 mmol of ammonium acetate **6** or aniline **68** were added to 2.0 mL of (H₂O: EtOH 1:3) as a solvent in the presence of 6.0 mg of Fe₃O₄-SiO₂-N-propylbenzoguanamine-SO₃H at 100 °C for the appropriate time. In this study, the prepared catalyst was identified using FT-IR, FESEM, EDXA, XRD, VSA,

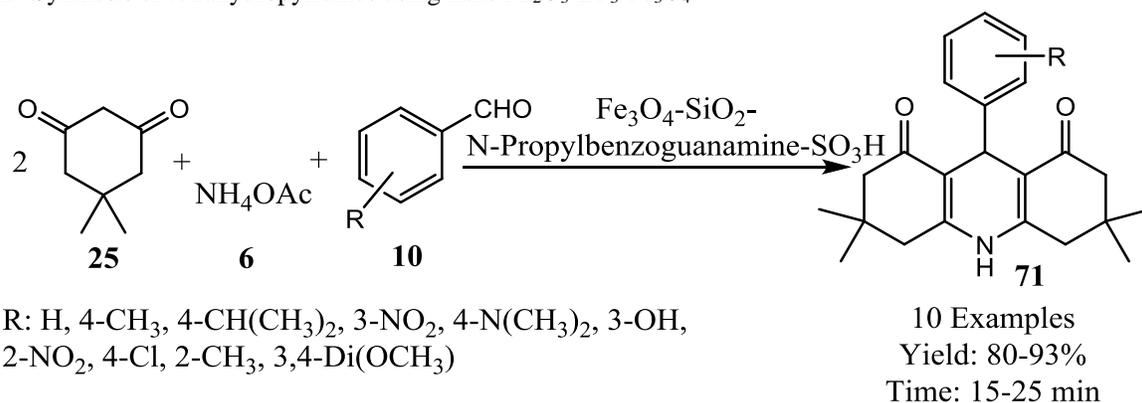
ICP-MS, and TGA techniques. The obtained tests show that the average size of $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-N-propylbenzoguanamine-SO}_3\text{H}$ was approximately 25 nm. The schematic representation of $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-N-propylbenzoguanamine-SO}_3\text{H}$ was shown in **Fig. 8**.

In 2018, Hazeri and co-workers reported a diastereoselective synthesis of fully functionalized tetrahydropyridines **75** using $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-TiO}_2$ nanocomposites at 50 °C in ethanol as a green solvent

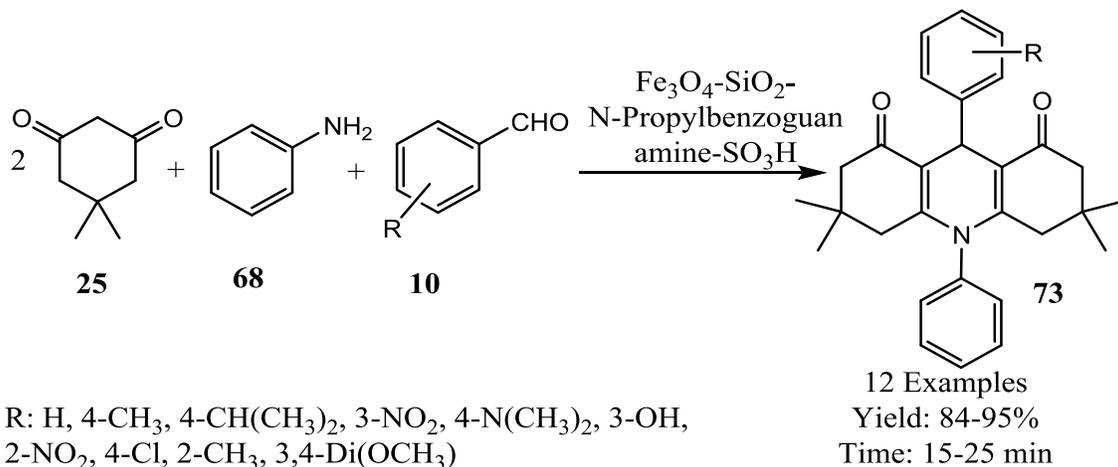
(**Scheme 28**) [35]. They used 2.0 mmol of aromatic amines **68**, 1.0 mmol of β -ketoester **74**, and 2.0 mmol of aromatic aldehydes **10** in the presence of 2 mol% of $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-TiO}_2$ nanocomposites at 50 °C for an appropriate time. The synthesized compounds were identified using $^1\text{H NMR}$ and melting point. The $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-TiO}_2$ nanocomposites show reusability for 4th runs catalytic cycles without loss of their catalytic activity.



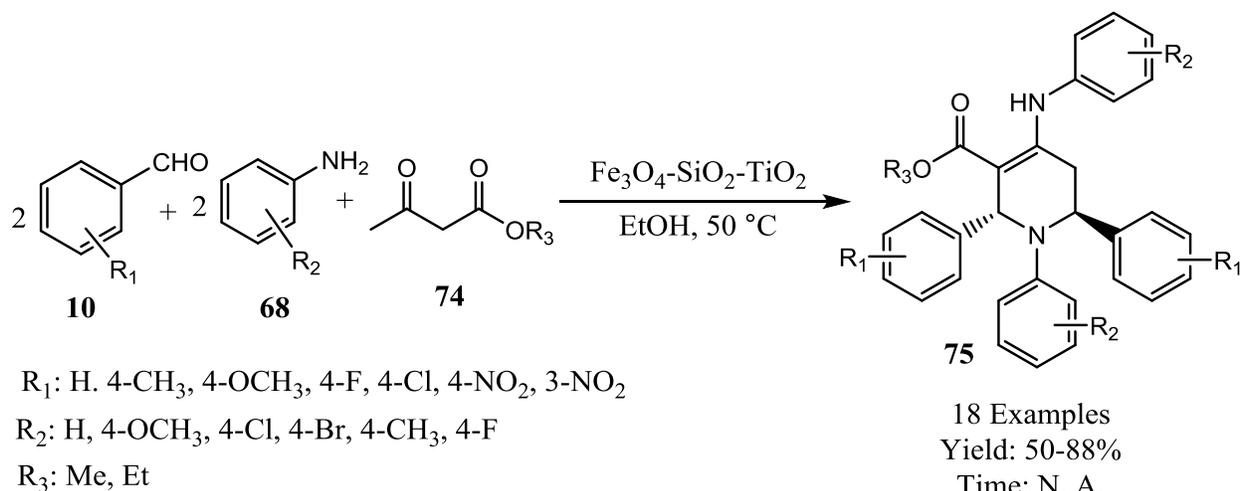
Scheme 25. Synthesis of tetrahydropyridines using nano $\text{Al}_2\text{O}_3\text{-BF}_3\text{-Fe}_3\text{O}_4$



Scheme 26. Synthesis of 1,8-dioxo-decahydroacridine derivatives using $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-N-propylbenzoguanamine-SO}_3\text{H}$ nanocomposite



Scheme 27. Synthesis of 1,8-dioxo-decahydroacridine derivatives using $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-N-propylbenzoguanamine-SO}_3\text{H}$ nanocomposite



Scheme 28. Diastereoselective synthesis of fully functionalized tetrahydropyridines using Fe₃O₄-SiO₂-TiO₂ nanocomposites

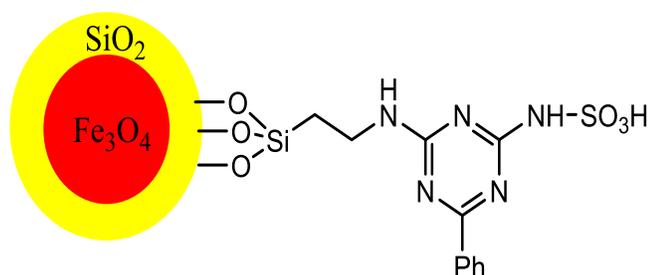
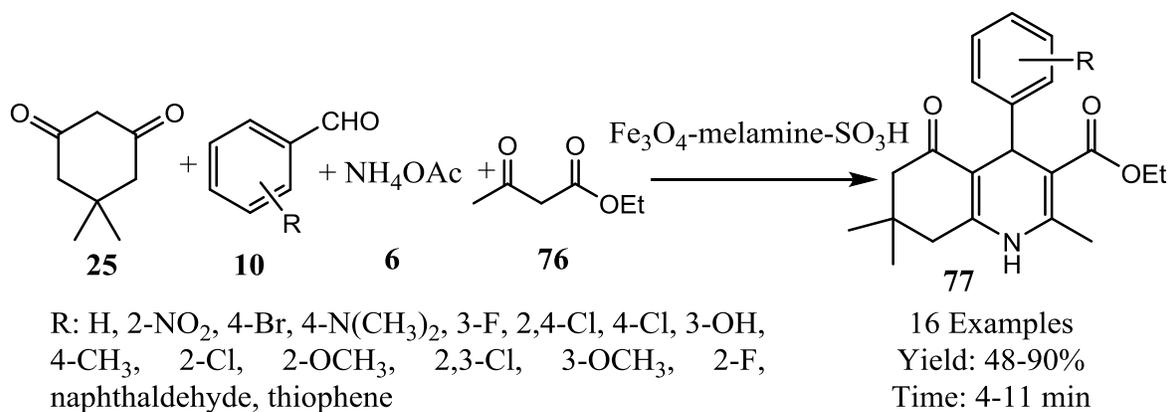


Fig. 8. The representation of the chemical structure of Fe₃O₄-SiO₂-N-propylbenzguanamine-SO₃H nanocomposite

In 2018, Khazaei and co-workers designed a new catalyst for the synthesis of hexahydroquinoline derivatives **77** (**Scheme 29**) [36]. The chemical structure of Fe₃O₄-melamine-SO₃H was shown in **Fig. 9**. The total density of sulfonic acid groups anchored on Fe₃O₄-melamine has been calculated at approximately 6.6 mmol of H⁺/g. The pointed catalyst was characterized using FT-IR, XRD, EDX, TEM, SEM,

TGA, and VSM techniques. Interestingly, they proposed the mechanism of the reaction using FT-IR spectroscopy. In addition, the Fe₃O₄-melamine-SO₃H show good reusability for 4 catalytic cycles. The metal-free, solvent-free, green chemistry and simple separation of the catalyst are other advantages of the presented method.

In 2018, Mirjalili and Aref reported nano-cellulose-BF₃-Fe₃O₄ as a bio-magnetic nanocatalyst for the synthesis of pyrimido[2,1-b]benzothiazoles **80** under solvent-free conditions (**Scheme 30**) [37]. The authors synthesized nano-cellulose from cotton. To prepare pyrimido[2,1-b]benzothiazoles, 0.06 g of nano-cellulose-BF₃-Fe₃O₄ was added to 1.0 mmol of ethyl acetoacetate **78**, 1.0 mmol of aromatic aldehyde **10**, and 1.0 mmol of 2-aminobenzothiazole **79** at 100 °C for the appropriate time. Moreover, the catalyst recycling tests were performed and displayed 4 catalytic cycles without loss of their catalytic activity.



Scheme 29. Synthesis of hexahydroquinoline derivatives using Fe₃O₄-melamine-SO₃H

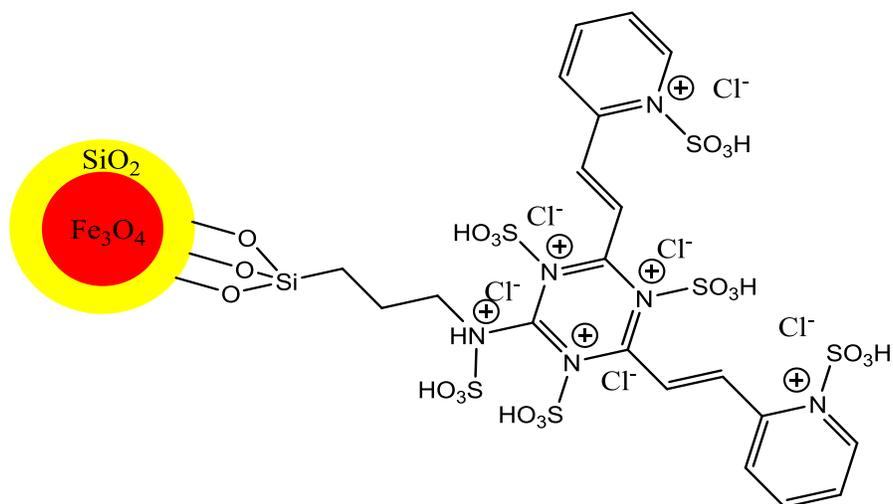
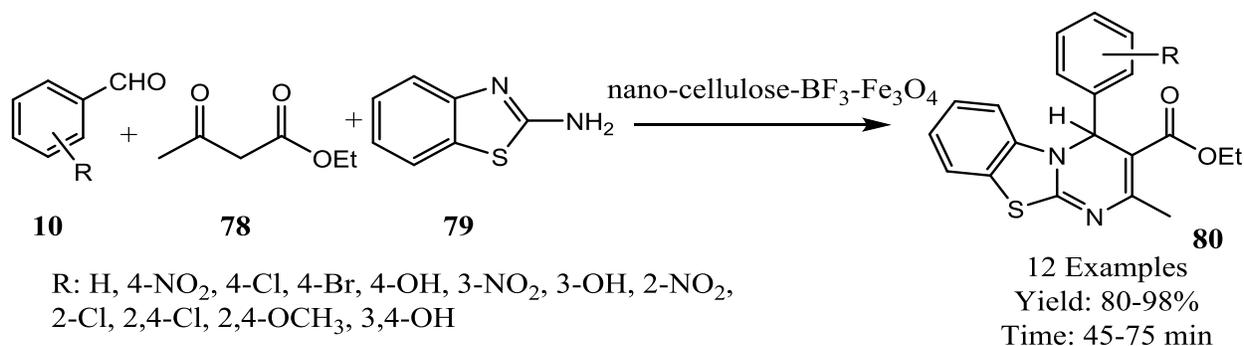


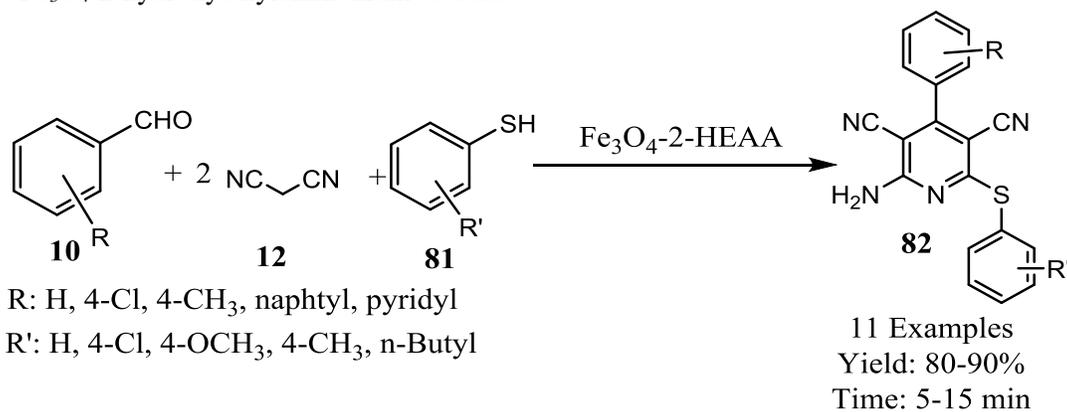
Fig. 9. The schematic representation of Fe₃O₄-melamine-SO₃H



Scheme 30. Synthesis of pyrimido[2,1-b]benzothiazoles using nano-cellulose-BF₃-Fe₃O₄ under solvent-free conditions

In 2018, Sobhani and co-workers developed a novel method for the synthesis of pyridine derivatives **82** using Fe₃O₄-2-hydroxyethylammonium acetate (HEAA) (**Scheme 31**) [38]. The schematic representation of Fe₃O₄-2-HEAA was shown in **Fig. 10**. For the preparation of 2-amino-3,5-dicarbonitrile-6-thio-pyridines in the presence of Fe₃O₄-2-hydroxyethylammonium acetate

(HEAA) (0.016 g or 1.0 mol%), 2.0 mmol of malononitrile **12**, 1.0 mmol of aromatic aldehyde **10**, and 1.0 mmol of thiol derivatives **81** were heated at 70 °C. The final product was identified using ¹H NMR, FT-IR, and melting point. The Fe₃O₄-2-hydroxyethylammonium acetate was characterized using FT-IR, XRD, SEM, TEM, and TGA analysis.



Scheme 31. Synthesis of 2-amino-3,5-dicarbonitrile-6-thio-pyridines in the presence of Fe₃O₄-2-hydroxyethylammonium acetate (HEAA)

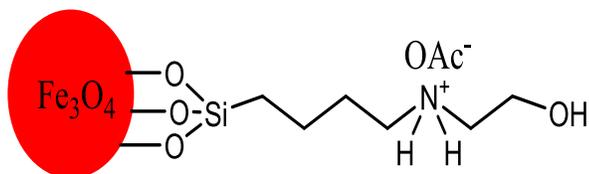


Fig. 10. The chemical structure of Fe_3O_4 -2-hydroxyethylammonium acetate (HEAA)

In 2017, Abdi and co-workers reported a green and efficient method for the synthesis of 1'Hspiro[isindoline-1,2'-quinazolin]3,4'(3'H)dione derivatives **86** in the presence of Fe_3O_4 -graphene oxide- SO_3H at 120°C under solvent-free conditions (**Scheme 32**) [39]. They used 1.4 mmol of isatoic anhydride **83**, 1.0 mmol of isatin **85**, 1.5 mmol of primary amine **84** using 1.0 g of Fe_3O_4 -graphene oxide- SO_3H at 120°C under solvent-free conditions. The 1'Hspiro[isindoline-1, 2'-quinazolin] 3, 4' (3'H)dione derivatives were identified using ^1H NMR, ^{13}C NMR, CHNS, and melting point. In addition, the Fe_3O_4 -graphene oxide- SO_3H was characterized by TEM, SEM, FT-IT, EDX, XRD, and TGA analysis. The reusability of the catalyst for 4 runs, short reaction times, and high yields of the products were some of the advantages of the current methodology.

In 2017, Beyki and Fallah-Mehrjerdi reported a powerful method for the synthesis of 2,3-dihydroquinazolin-4(1H)-ones **89** using Fe_3O_4 - SiO_2 - SO_3H at room temperature under solvent-free conditions (**Scheme 33**) [40]. The chemical structure of Fe_3O_4 - SiO_2 - SO_3H was shown in **Fig. 11**. In the current research, 0.05 g of Fe_3O_4 - SiO_2 - SO_3H was added to the 1.0 mmol of anthralinamide **87** and 1.0 mmol of aldehydes or ketones **88** for the appropriate time. The prepared catalyst was characterized using FT-IR, SEM,

EDX, and XRD analysis. The authors compared the prepared catalyst with other reported catalysts such as $\text{Ga}(\text{OTf})_3$, $\text{Sc}(\text{OTf})_3$, NH_4Cl , $\text{H}_3\text{PW}_{12}\text{O}_{40}$, $\text{SiCl}_2 \cdot 6\text{H}_2\text{O}$, ZrCl_4 , and $\text{Y}(\text{OTf})_2$. Their comparison shows that the Fe_3O_4 - SiO_2 - SO_3H nanocatalyst was superior in terms of the time of the reaction and yield of the final product.

In Taiwan, in 2017, Dam and co-workers reported an efficient and novel method for the preparation of dihydroquinazolinone derivatives **94** using Fe_3O_4 -DOPA(dopamine)- SnO_2 nanocatalyst at 80°C in water (**Scheme 34**) [41]. To synthesize dihydroquinazolinone derivatives **94**, 1.0 mmol of isatoic anhydride **90**, 1.5 mmol of ammonium acetate **92**, and 1.0 mmol of amine derivatives **91** in the presence of 10 mg of Fe_3O_4 -DOPA- SnO_2 nanocatalyst in 5.0 mL of water were heated at 80°C . The Fe_3O_4 -DOPA- SnO_2 nanocatalyst was characterized by FT-IR, TEM, HRTEM, SEM, EDX, XPS, XRD, VSM, and TGA analysis. Good to excellent yield, relatively reaction times, green solvent, and reusability of the catalyst for 5 catalytic cycles are some of the advantages of the presented method.

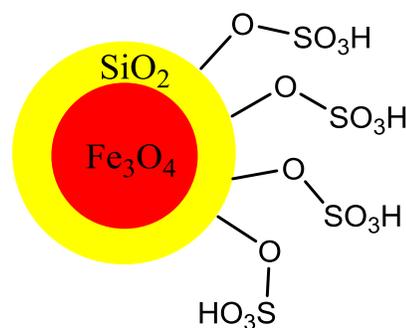
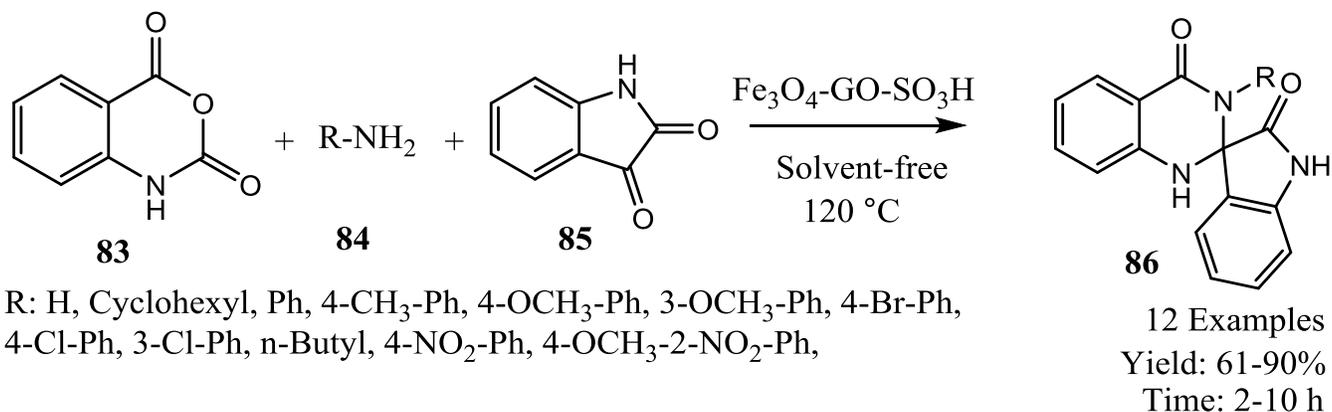
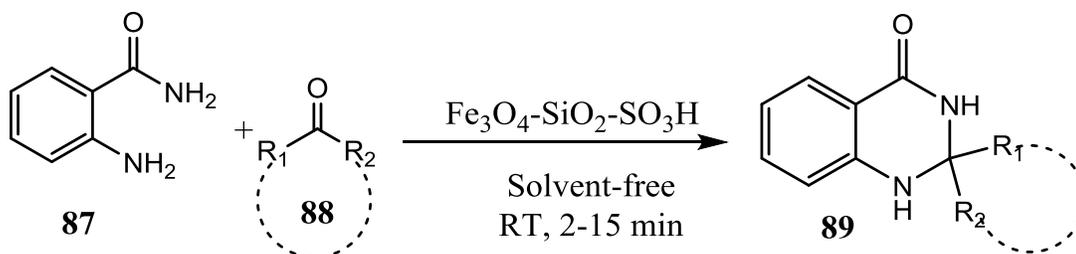


Fig. 11. The schematic representation of Fe_3O_4 - SiO_2 - SO_3H



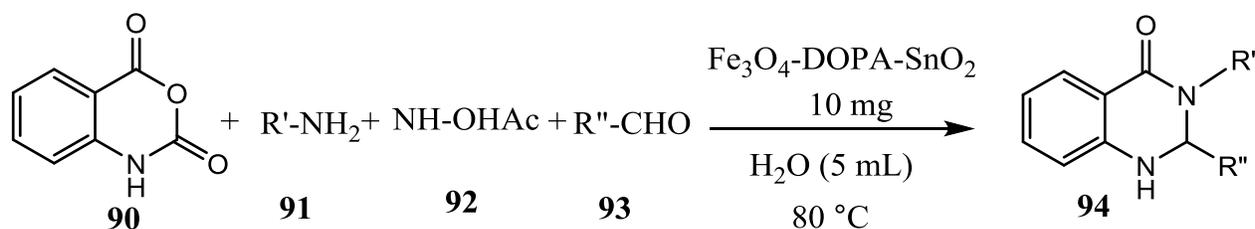
Scheme 32. Synthesis of 1'Hspiro [isindoline-1, 2'-quinazolin] 3, 4' (3'H)dione derivatives using Fe_3O_4 -graphene oxide- SO_3H



A: Benzaldehyde (a); 4-CH₃-a; 4-OCH₃-a; 2-OH-a;
 2-Cl-a; 3-Cl-a; 4-Cl-a; 3-NO₂-a; 2,5-DiOCH₃-a;
 3,4-DiOH-a; 4-Dimethylamino-a; Cinnamaldehyde;
 2-Furaldehyde; Cyclopentanone(hexanone) (heptanone);
 Isatin; 1,4-Cyclohexandione

19 Examples
 Yield: 82-96%
 Time: 2-15 min

Scheme 33. Synthesis of 2,3-dihydroquinazolin-4(1H)-ones using $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-SO}_3\text{H}$



R': 4-methylphenyl; benzimidazole

R'': 4-Cl, 4-CN, 4-Br, 4-OCH₃, 4-CH₃, 2-NO₂, 2-CH₃, H,
 2-OCH₃-PhCHO, aliphatic aldehyde, tricyclic aldehyde

16 Examples
 Yield: 82-92%
 Time: 45-90 min

Scheme 34. Preparation of dihydroquinazolinone derivatives using $\text{Fe}_3\text{O}_4\text{-DOPA(dopamine)-SnO}_2$ nanocatalyst

In 2017, Fattahi and co-workers reported a one-pot and efficient synthesis of novel pyrimido[5',4':5,6]pyrido[2,3-d]pyrimidines **96** catalyzed by magnetic iron oxide nanoparticles (**Scheme 35**) [42]. They reacted 2.0 mmol of 1,3-dimethylbarbituric acid **95**, 1.0 mmol of aromatic aldehydes **10**, and 1.2 mmol of ammonium acetate **6** in the presence of 0.075 g of Fe_3O_4 nanoparticles in 5 mL of ethanol: water (1:1). The obtained products were approved using ¹³C NMR, ¹H NMR, FT-IR, and melting point.

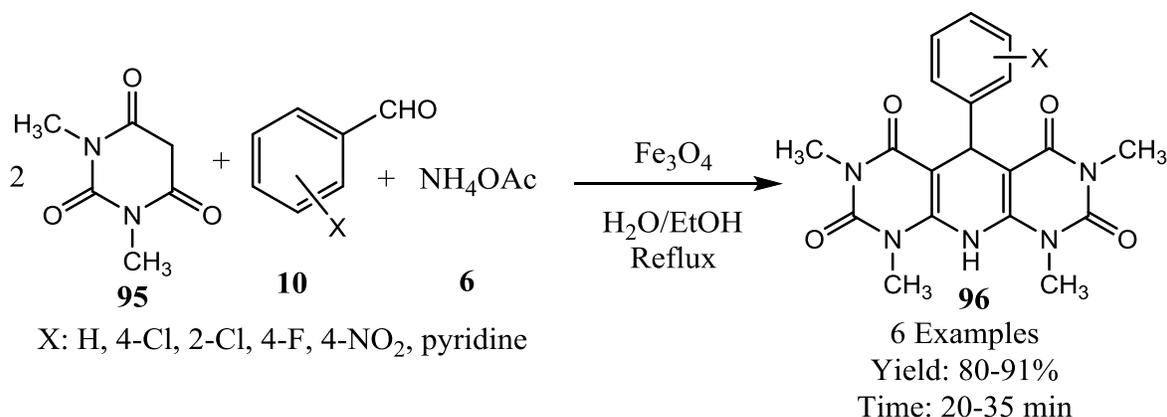
In 2017, Mirjalili and Tafti developed a novel method for the synthesis of quinoxaline derivatives **99** using nano-Kaoline-BF₃-Fe₃O₄ nanocomposite in grinding conditions (**Scheme 36**) [43]. The final products were identified using ¹H NMR, ¹³C NMR, FT-IR, and melting point. In addition, the nano-Kaoline-BF₃-Fe₃O₄ nanocomposite was characterized using FT-IR, SEM, EDX, TEM, VSM, TGA, BET, and XRD analysis.

In 2017, Naeimi and Lahouti developed a one-pot synthesis of spiroacridines **102** using $\text{Fe}_3\text{O}_4\text{-CS-SO}_3\text{H}$ under ultrasound irradiation (**Scheme 37**) [44]. In this three-component and one-pot reaction, a mixture of

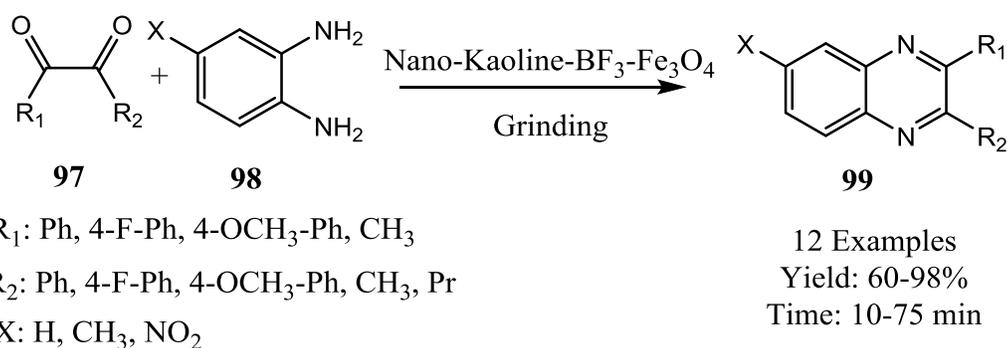
acenaphthoquinone **101**, dimedone **25**, and different anilines **100** were used in the presence of $\text{Fe}_3\text{O}_4\text{-CS-SO}_3\text{H}$ in ethanol: water under ultrasound irradiation. They characterized the $\text{Fe}_3\text{O}_4\text{-CS-SO}_3\text{H}$ using XRD, VSM, SEM, FT-IR, and EDX techniques. They found that in comparison with conventional methods, ultrasound irradiation has several worthwhile advantages including the high yield of final products, short reaction times, and reusability of the catalyst.

In China, Zhang and co-workers designed a new ionic liquid supported on Fe_3O_4 nanoparticles for the synthesis of polyhydroquinolines **105** at 70 °C under solvent-free conditions (**Scheme 38**) [45]. The prepared catalyst was characterized using XRD, TGA, FT-IR, TEM, and VSM analysis. The $\text{Fe}_3\text{O}_4\text{-SiO}_2\text{-BIL}$ shows the reusability for six catalytic cycles without loss of their catalytic activity.

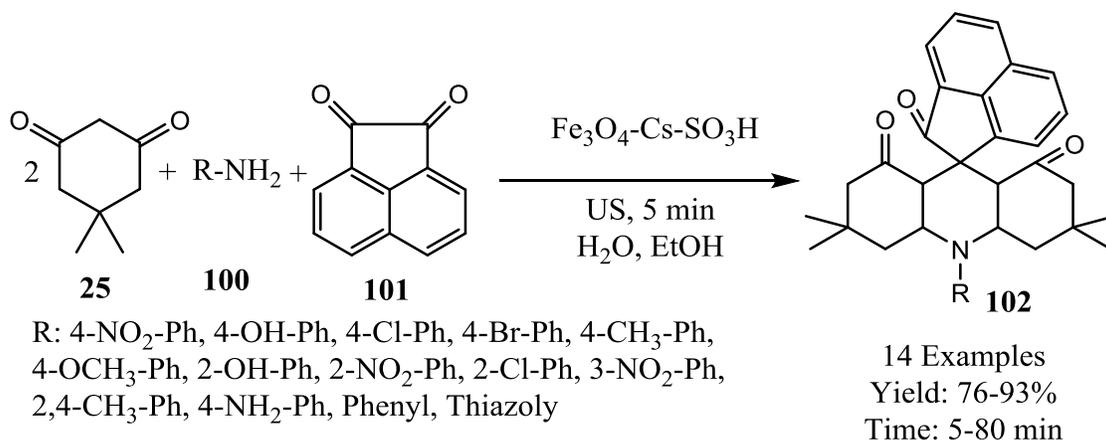
This group also checked the synthesis of Knoevenagel condensation of aromatic aldehydes with malonitrile derivatives (**Scheme 39**). The schematic representation of the prepared catalyst was shown in **Fig. 12**.



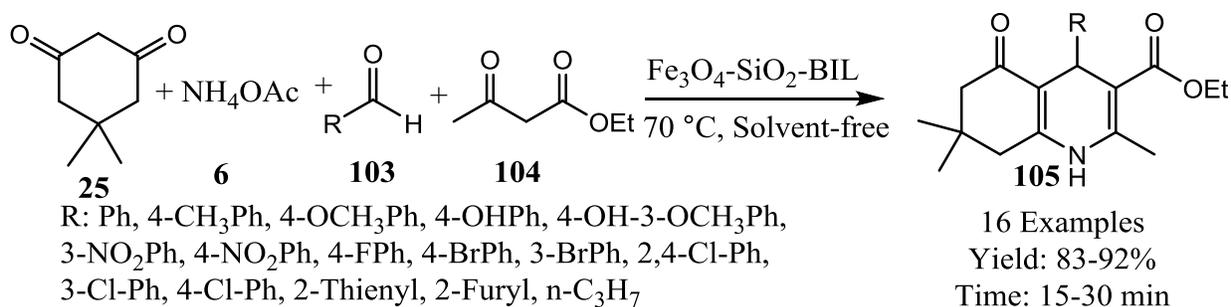
Scheme 35. Synthesis of pyrimido[5',4':5,6]pyrido[2,3-d]pyrimidines using Fe_3O_4 nanoparticles



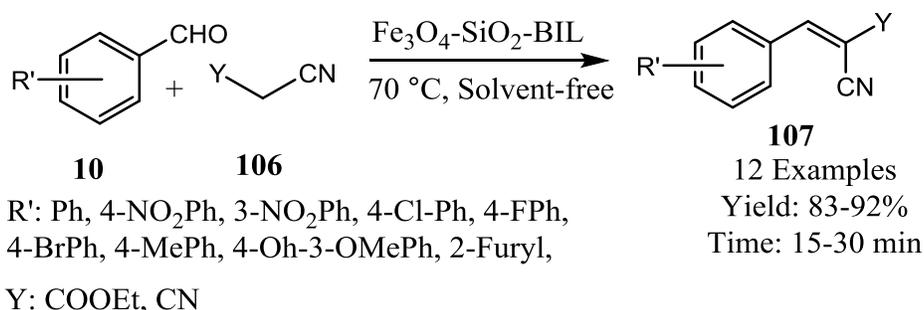
Scheme 36. Synthesis of quinoxaline derivatives using nano-Kaoline- BF_3 - Fe_3O_4 nanocomposite



Scheme 37. One-pot synthesis of spiroacridines using Fe_3O_4 -CS- SO_3H under ultrasound irradiation



Scheme 38. Synthesis of polyhydroquinolines using Fe_3O_4 - SiO_2 -BIL



Scheme 39. The Knoevenagel condensation by Fe₃O₄-SiO₂-BIL

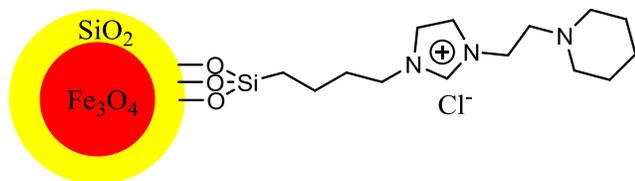


Fig. 12. The chemical structure of Fe₃O₄-SiO₂-BIL

There is a lot of review paper concerning heterocyclic compounds in the research database. But the six-membered heterocyclic compounds containing nitrogen atom has grown into a wide subject area in recent years. The main limitation of this review paper was focusing on nitrogen atom compounds while a lot of heterocyclic compounds with other atoms such as oxygen, and sulfur are in natural and drug compounds. The strength of this review paper is the discussion of the 6-membered ring compounds containing the nitrogen atom catalyzed by magnetically heterogeneous nanocatalysts based on Fe₃O₄ nanoparticles.

4. Conclusions

In conclusion, a comprehensive review of the catalytic methods was developed for the synthesis of six-membered compounds (heterocyclic compounds) containing nitrogen catalyzed by functionalized Fe₃O₄ nanoparticles. The application of these catalytic systems in the synthesis of different organic reactions such as cyclization, condensation, ring-opening, and addition reactions have been investigated in the presence of functionalized Fe₃O₄ nanoparticles. In addition, it is described in all of the reactions, that the magnetic catalytic systems could be separated by the use of the external magnet. Notably, in all of the reactions, a new carbon-nitrogen bond was formed to form six-membered organic compounds.

Authors Contributions

Faris H. Mohammed proposed the idea, searched databases, downloaded PDFs, extracted data, and wrote the draft. Aseel M. Aljeboree proposed the idea, searched databases, downloaded PDFs, and extracted

data. Alrazzak Nour Abd was the head of the group and rewrote the final draft and submission. Ayad F. Alkaim extracted data and drew the chemical compounds. Yasir Salam Karim extracted data and drew the chemical compounds. Sarah A. Hamood revised English and revised based on the editors' comments. Ahmed B. Mahdi and Mohammed Abed Jawad proposed the idea, searched the databases, and downloaded PDFs. Salam Ahjel extracted data.

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References

- [1] R.B. Sosman, J.C. Hostetter, THE OXIDES OF IRON. I. SOLID SOLUTION IN THE SYSTEM FE₂O₃—FE₃O₄, J. Am. Chem. Soc., 38 (1916) 807-833.
- [2] E. Vitaku, D.T. Smith, J.T. Njardarson, Analysis of the Structural Diversity, Substitution Patterns, and Frequency of Nitrogen Heterocycles among U.S. FDA Approved Pharmaceuticals, J. Med. Chem., 57 (2014) 10257-10274.
- [3] M.M. Heravi, V. Zadsirjan, Prescribed drugs containing nitrogen heterocycles: an overview, RSC Adv., 10 (2020) 44247-44311.
- [4] S. Mohan, L. Sim, D.R. Rose, B.M. Pinto, Probing the active-site requirements of human intestinal N-terminal maltase-glucoamylase: Synthesis and enzyme inhibitory activities of a six-membered ring nitrogen analogue of kotalanol and its de-O-sulfonated derivative, Bioorg. Med. Chem., 18 (2010) 7794-7798.
- [5] J.S. Carey, D. Laffan, C. Thomson, M.T. Williams, Analysis of the reactions used for the preparation of drug candidate molecules, Org. Biomol. Chem., 4 (2006) 2337-2347.

- [6] M. Vatanparast, Z. Shariatinia, Revealing the role of different nitrogen functionalities in the drug delivery performance of graphene quantum dots: a combined density functional theory and molecular dynamics approach, *Journal of Materials Chemistry B*, 7 (2019) 6156-6171.
- [7] Q. Deng, X. Meng, Recent Advances in the Cycloaddition Reactions of 2-Benzylidene-1-benzofuran-3-ones, and Their Sulfur, Nitrogen and Methylene Analogues, *Chemistry – An Asian Journal*, 15 (2020) 2838-2853.
- [8] H. Dong, W. Li, J. Sun, S. Li, M.L. Klein, Understanding the Boron–Nitrogen Interaction and Its Possible Implications in Drug Design, *J. Physic. Chem. B*, 119 (2015) 14393-14401.
- [9] Y.-J. Wu, N.A. Meanwell, Geminal Diheteroatomic Motifs: Some Applications of Acetals, Ketals, and Their Sulfur and Nitrogen Homologues in Medicinal Chemistry and Drug Design, *J. Med. Chem.*, 64 (2021) 9786-9874.
- [10] S. Dongbang, D.N. Confair, J.A. Ellman, Rhodium-Catalyzed C–H Alkenylation/Electrocyclization Cascade Provides Dihydropyridines That Serve as Versatile Intermediates to Diverse Nitrogen Heterocycles, *Acc. Chem. Res.*, 54 (2021) 1766-1778.
- [11] T.P. Andrejević, B. Warzajtis, B.Đ. Glišić, S. Vojnovic, M. Mojicevic, N.L. Stevanović, J. Nikodinovic-Runic, U. Rychlewska, M.I. Djuran, Zinc(II) complexes with aromatic nitrogen-containing heterocycles as antifungal agents: Synergistic activity with clinically used drug nystatin, *J. Inorg. Biochem.*, 208 (2020) 111089.
- [12] M. Savari, A. Varasteh-Moradi, S.Z. Sayyed-Alangi, Z. Hossaini, R. Zafarmehrabian, Ag/Fe₃O₄/TiO₂@MWCNTs as a reusable organometallic nanocatalyst promoted green synthesis of new pyridobenzoazepines: Study of biological activity and reduction of organic pollutants, *Appl. Organomet. Chem.*, 36 (2022) e6510.
- [13] S. Motahari, A. Khazaei, Preparation and Characterization of Co (II) Supported on Modified Magnetic Fe₃O₄ Nanoparticles and Its Application as Catalyst for the Synthesis of 2-Amino-3-Cyanopyridines, *Polycyclic Aromat. Compd.*, (2021) 1-12.
- [14] E. Ezzatzadeh, F.Z. Hargalani, F. Shafaei, Bio-Fe₃O₄-MNPs Promoted Green Synthesis of Pyrido[2,1-a]isoquinolines and Pyrido[1,2-a]quinolines: Study of Antioxidant and Antimicrobial Activity, *Polycyclic Aromat. Compd.*, (2021) 1-16.
- [15] Z. Hossaini, N. Tabarsaei, S. Khandan, P. Valipour, M. Ghorchibeigi, ZnO/Ag/Fe₃O₄ nanoparticles supported on carbon nanotubes employing Petasites hybridus rhizome water extract: A novel organometallic nanocatalyst for the synthesis of new naphthyridines, *Appl. Organomet. Chem.*, 35 (2021) e6114.
- [16] A. Bamoniri, A. Mazoochi, S.A. Pourmousavi, Synthesis of 2,3-dihydroperimidines in the Presence of Nano- γ -Al₂O₃/BF₃ and Nano- γ -Al₂O₃/BF₃/Fe₃O₄ as Catalysts under Different Conditions, *Journal of Nanostructures*, 11 (2021) 554-567.
- [17] A. Moussa, A. Rahmati, Synthesis and characterization of silica-coated Fe₃O₄ nanoparticle@silylpropyl triethylammonium polyoxometalate as an organic–inorganic hybrid heterogeneous catalyst for the one-pot synthesis of tetrahydrobenzimidazo[2,1-b]quinazolin-1(2H)-ones, *Appl. Organomet. Chem.*, 34 (2020) e5894.
- [18] M. Mahmoodi Fard Chegeni, A. Bamoniri, A.A. Taherpour, One-pot synthesis of 2H-indazolo[2,1-b]phthalazine-triones via nano γ -Al₂O₃/BF₃/Fe₃O₄ as an efficient catalyst and theoretical DFT study on them, *J. Heterocycl. Chem.*, 57 (2020) 2801-2814.
- [19] M. Ghoorchibeigi, K. Larijani, P.A. Azar, K. Zare, I. Mehregan, ZnO/Fe₃O₄ nanoparticles promoted green synthesis of pyrazolo pyrimidinones: Study of antioxidant activity, *J. Heterocycl. Chem.*, 57 (2020) 3612-3621.
- [20] A. Zare, M. Barzegar, Dicationic ionic liquid grafted with silica-coated nano-Fe₃O₄ as a novel and efficient catalyst for the preparation of uracil-containing heterocycles, *Res. Chem. Intermed.*, 46 (2020) 3727-3740.
- [21] Z. Hossaini, A. Noushin, P. Valipour, M. Ghazvini, M.H. Rostam, Reusable Fe₃O₄/ZnO/MWCNTs Magnetic Nanocomposites Promoted Synthesis of New Naphthyridines, *Polycyclic Aromat. Compd.*, (2020) 1-20.
- [22] F. Shafaei, S. Sharafian, Green Synthesis of Imidazole Derivatives via Fe₃O₄-MNPs as Reusable Catalyst, *J. Heterocycl. Chem.*, 56 (2019) 2644-2650.
- [23] N. Faal Hamedani, M. Ghazvini, L. Azad, A. Noushin, Green synthesis of pyrido[2,1-a]isoquinolines and pyrido[1,2-a]quinolins using Fe₃O₄-MNPs as efficient nanocatalyst: Study of antioxidant activity, *J. Heterocycl. Chem.*, 57 (2020) 337-345.

- [24] M.A. Ashraf, Z. Liu, W.-X. Peng, C. Gao, New Copper Complex on Fe₃O₄ Nanoparticles as a Highly Efficient Reusable Nanocatalyst for Synthesis of Polyhydroquinolines in Water, *Catal. Lett.*, 150 (2020) 683-701.
- [25] M. Kalhor, Z. Zarnegar, Fe₃O₄/SO₃H@zeolite-Y as a novel multi-functional and magnetic nanocatalyst for clean and soft synthesis of imidazole and perimidine derivatives, *RSC Adv.*, 9 (2019) 19333-19346.
- [26] S. Andalibi Salem, A. Khazaei, J.Y. Seyf, N. Sarmasti, M. Mahmoudiani Gilan, Preparation of Magnetic Cu(II) Nano-structure (Based on Nano-Fe₃O₄) and Application to the Synthesis of Hexahydroquinoline Derivatives, *Polycyclic Aromat. Compd.*, 41 (2021) 319-332.
- [27] E. Babaei, B.B.F. Mirjalili, One Pot Aqueous Media Synthesis of 1,3-Oxazine Derivatives Catalyzed by Reusable Nano-Al₂O₃/BF₃/Fe₃O₄ at Room Temperature, *Polycyclic Aromat. Compd.*, 41 (2021) 518-525.
- [28] A. Karimiyan, S. Rostamizadeh, An Efficient Route to Imidazo[1,2-a]Pyridines and Pyrido[1,2-a]Pyrimidines via One-Pot Four-Component Reaction Using Fe₃O₄-GO-SO₃H, *Polycyclic Aromat. Compd.*, 41 (2021) 1483-1494.
- [29] N. Basavegowda, K. Mishra, Y.R. Lee, Fe₃O₄-decorated MWCNTs as an efficient and sustainable heterogeneous nanocatalyst for the synthesis of polyfunctionalised pyridines in water, *Materials Technology*, 34 (2019) 558-569.
- [30] P. Farokhian, M. Mamaghani, N.O. Mahmoodi, K. Tabatabaieian, A.F. Shojaie, An expeditious one-pot synthesis of pyrido[2,3-d]pyrimidines using Fe₃O₄-ZnO-NH₂-PW12O₄₀ nanocatalyst, *J. Chem. Res.*, 43 (2019) 135-139.
- [31] B.B.F. Mirjalili, R. Soltani, Nano-kaolin/Ti⁴⁺/Fe₃O₄: a magnetic reusable nano-catalyst for the synthesis of pyrimido[2,1-b]benzothiazoles, *RSC Adv.*, 9 (2019) 18720-18727.
- [32] M.A. Amrollahi, F. Vahidnia, Decoration of β-CD-ZrO on Fe₃O₄ magnetic nanoparticles as a magnetically, recoverable and reusable catalyst for the synthesis of 2,3-dihydro-1H-perimidines, *Res. Chem. Intermed.*, 44 (2018) 7569-7581.
- [33] E. Babaei, B.B.F. Mirjalili, One-pot synthesis of five substituted tetrahydropyridines using nano-Al₂O₃/BF₃/Fe₃O₄ as a highly efficient nano-catalyst, *Res. Chem. Intermed.*, 44 (2018) 3493-3505.
- [34] M. Gholami Dehbalaei, N. Foroughifar, H. Pasdar, A. Khajeh-Amiri, N-Propyl benzoguanamine sulfonic acid supported on magnetic Fe₃O₄ nanoparticles: a novel and efficient magnetically heterogeneous catalyst for the synthesis of 1,8-dioxo-decahydroacridine derivatives, *New J. Chem.*, 42 (2018) 327-335.
- [35] N. Hazeri, S. Salahi, M. Lashkari, M.T. Maghsoodlou, E. Esmaeili-Shahri, E. Mollashahi, Facile Diastereoselective Synthesis of Functionalized Tetrahydropyridines Using Fe₃O₄/SiO₂/TiO₂ Nanocomposites, *Org. Prep. Proced. Int.*, 50 (2018) 375-383.
- [36] A. Khazaei, N. Sarmasti, J. Yousefi Seyf, Anchoring high density sulfonic acid based ionic liquid on the magnetic nano-magnetite (Fe₃O₄), application to the synthesis of hexahydroquinoline derivatives, *J. Mol. Liq.*, 262 (2018) 484-494.
- [37] B.B.F. Mirjalili, F. Aref, Nano-cellulose/BF₃/Fe₃O₄: a magnetic bio-based nano-catalyst for the synthesis of pyrimido[2,1-b]benzothiazoles under solvent-free conditions, *Res. Chem. Intermed.*, 44 (2018) 4519-4531.
- [38] S. Sobhani, F. Nasserri, F. Zarifi, Unique role of 2-hydroxyethylammonium acetate as an ionic liquid in the synthesis of Fe₃O₄ magnetic nanoparticles and preparation of pyridine derivatives in the presence of a new magnetically recyclable heterogeneous catalyst, *J. Iran. Chem. Soc.*, 15 (2018) 2721-2732.
- [39] M. Abdi, S. Rostamizadeh, N. Zekri, An Efficient and Green Synthesis of 1'Hspiro[isoinoline-1,2'-quinazoline]-3,4'(3'H)-dione Derivatives in the Presence of Nano Fe₃O₄-GO-SO₃H, *Polycyclic Aromat. Compd.*, 39 (2019) 413-424.
- [40] M. Beyki, M. Fallah-Mehrjardi, Silica Sulfuric Acid-coated Fe₃O₄ Nanoparticles as a Highly Efficient and Reusable Solid Acid Catalyst for the Green Synthesis of 2,3-Dihydroquinazolin-4(1H)-ones under Solvent-free Conditions, *Lett. Org. Chem.*, 15 (2018) 39-44.
- [41] B. Dam, R.A. Patil, Y.-R. Ma, A.K. Pal, Preparation, characterization and catalytic application of nano-Fe₃O₄-DOPA-SnO₂ having high TON and TOF for non-toxic and sustainable synthesis of dihydroquinazolinone derivatives, *New J. Chem.*, 41 (2017) 6553-6563.
- [42] M. Fattahi, A. Davoodnia, M. Pordel, Efficient one-pot synthesis of some new pyrimido[5',4':5,6]pyrido[2,3-d]pyrimidines catalyzed by magnetically recyclable Fe₃O₄ nanoparticles, *Russ. J. Gen. Chem.*, 87 (2017) 863-867.

[43] B.B.F. Mirjalili, M. Dehghani Tafti, Synthesis of quinoxalines promoted by eco-friendly nano-kaoline/BF₃/Fe₃O₄ nano-catalyst under grinding condition, *Sci. Iranica*, 24 (2017) 3014-3021.

[44] H. Naeimi, S. Lahouti, Sonochemical one pot synthesis of novel spiroacridines catalyzed by magnetically functionalized Fe₃O₄ nanoparticles with chitosan as a reusable effective catalyst, *RSC Adv.*, 7 (2017) 2555-2562.

[45] Q. Zhang, X.-M. Ma, H.-X. Wei, X. Zhao, J. Luo, Covalently anchored tertiary amine functionalized ionic liquid on silica coated nano-Fe₃O₄ as a novel, efficient and magnetically recoverable catalyst for the unsymmetrical Hantzsch reaction and Knoevenagel condensation, *RSC Adv.*, 7 (2017) 53861-53870.