

Microwave Assisted Synthesis of Some Novel Sulphonamide bearing Pyrazolone Core Structure

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ABSTRACT

We have described some novel Sulphonamide bearing pyrazoline derivatives synthesized by conventional method as well as microwave assisted method of synthesis. The reaction of 4-(3-methyl-5-oxo-4,5-dihydro-1-*H*-pyrazol-1-yl)benzenesulphonamide with substituted benzaldehyde in the presence of Methanol as solvent and piperidine as catalyst, generated a series of substituted pyrazolone derivatives 4a-m. The structures of all synthesized compounds are well characterized by Mass spectroscopy, FT-IR, ¹H NMR and elemental analysis. After obtaining experimental data regarding the yield and the time taken for the synthesis by both the approaches, convenient and microwave assisted method, it was ascertained that the microwave assisted method is more suitable for synthesis of pyrazolone derivatives 4a-m.

Keywords: Pyrazolone; Arylidene; Sulphonamide

1. INTRODUCTION

Pyrazolone, a five-membered-ring lactam, is a derivative of pyrazole that has an additional keto (=O) group. It has a molecular formula of C₃H₄NO. The chemistry of pyrazolone began in 1883 when Knorr reported the first pyrazolone derivative. The reaction of phenyl hydrazine and ethyl acetoacetate resulted in novel structure identified in 1887 as 1-phenyl-3-methyl-5-pyrazolone [6]. The Knorr pyrazole synthesis is the reaction of hydrazine with 1,3-dicarbonyl compounds to provide the pyrazole or pyrazolone ring system. The prototype molecule, antipyrine was synthesized for clinical use in 1883. The pyrazolone nucleus has been known to exist in three tautomeric structures [7].

All these compounds are characterized by the presence of a phenyl group attached to nitrogen atom in the 1- position and a methyl group in 3- position. Phenyl group in 1- position and a methyl group in 3- position seem to be essential for antipyretic activity. Several 4,4-dimethyl derivatives, as well as Pyrazole Blue and tartrazine are derived from formula II whereas from structure III several pyrazolone dyes have been derived. When pyrazolones were discovered, they were only known as non steroidal anti-inflammatory agents (or drugs) – NSAID, but in recent times, they are known to exhibit antioxidant, anticancer, antibacterial and several other pharmacological actions [6,8]. Pyrazolones are

very important class of heterocycles due to their wide applications in pharmacological and biological activities [3,5]. Large numbers of 2-pyrazolin-5-ones have been used as therapeutics agents such as analgesics and antipyretics [1,2,4].

Microbial development of resistance, as well as economic incentives, has resulted in research and development in the search for new antibiotics in order to maintain a pool of effective drugs at all time. It is important to find out newer, safer and more effective antibiotics with broad spectrum of activity. Although several antifungal agents and the azole class of drugs are currently available there is clearly a critical need for the development of new specific antimicrobial agents.

Heterocycles containing a pyrazole / pyrazolone ring system are found to exhibit a wide spectrum of biological activities, including antibacterial and antifungal activities. Due to their easier preparation and rich biological activity, pyrazolone framework plays an essential role and represents an interesting template for combinatorial and medicinal chemistry. Similarly, pyrazole derivatives have showed significant biological activities, such as anti-microbial [16], analgesic [17], anti-inflammatory [18] and anticancer [19] activities.

This gave a great impetus to the search for potential pharmacologically active drugs carrying pyrazole substituents. Increasing antibiotic resistance in microbial populations has necessitated the search for alternate cellular targets for new and existing antimicrobial agents. It is well established that small modifications in the structure of the targets are altering their biological character as well as their physiochemical properties. A detailed literature survey on antimicrobial activity of various types of compounds clearly indicates that presence of certain pharmacophore such as pyrazole in any molecule plays an important role in enhancing activity.

Keeping in view of this and in continuation of our search on biologically potent molecules [20-23], we hereby report the synthesis and antimicrobial property of some new pyrazolone derivatives. Multi drug resistance is widespread with specific relevance to Gram positive and Gram negative bacteria. Infections caused by these organisms create a serious challenge to the community.

The therapeutic problem is more pronounced in patients with immuno-compromised system or those undergoing anticancer therapy substantiating the need for design and development of novel less toxic potent antimicrobial agents. Inflammation is a nonspecific immune response in which the body reacts to infection, localized irritation, free radicals, other injury or disease [24].

Antipyrine, 2,3-dimethyl-1-phenyl-3-pyrazolin-5-one, was the first pyrazolone derivative used in the management of pain and inflammation. Simultaneous administration of several drugs to treat inflammatory conditions that might be associated with some microbial and fungal infections may cause serious health problems, especially in patients with compromised liver or kidney functions.

So the discovery of a dual anti-fungal and anti-microbial agent with potential activity and fewer adverse effects not only results in a pharmaco economic agent but also leads to better patient compliance. This study was aimed at the synthesis, characterization and screening of some new pyrazolone derivatives by most convenient microwave assisted method.

2. EXPERIMENTAL

All the reactions were carried out in domestic microwave synthesizer. Melting points of all the synthesized compounds have been recorded by open capillary method. Microanalyses were performed on Euro EA Elemental Analyser, at National facility for drug discovery (NFDD) center Saurashtra university Rajkot. Reaction was monitored by thin-layer chromatography (TLC) on silica gel (Merck 60 F₂₅₄) using (3:7) Ethyl acetate: Hexane as solvent system.

The IR spectra were recorded on a Shimadzu FT-IR-8400 instrument using KBr pellet method. Mass spectra were recorded on Shimadzu GC-MS-QP-2010 model using Direct Injection Probe technique. ¹H NMR was determined in CDCl₃/DMSO solution on a Bruker Ac 400 MHz spectrometer. Commercial grade solvents and reagents were used without further purification.

2. 1. Chemistry

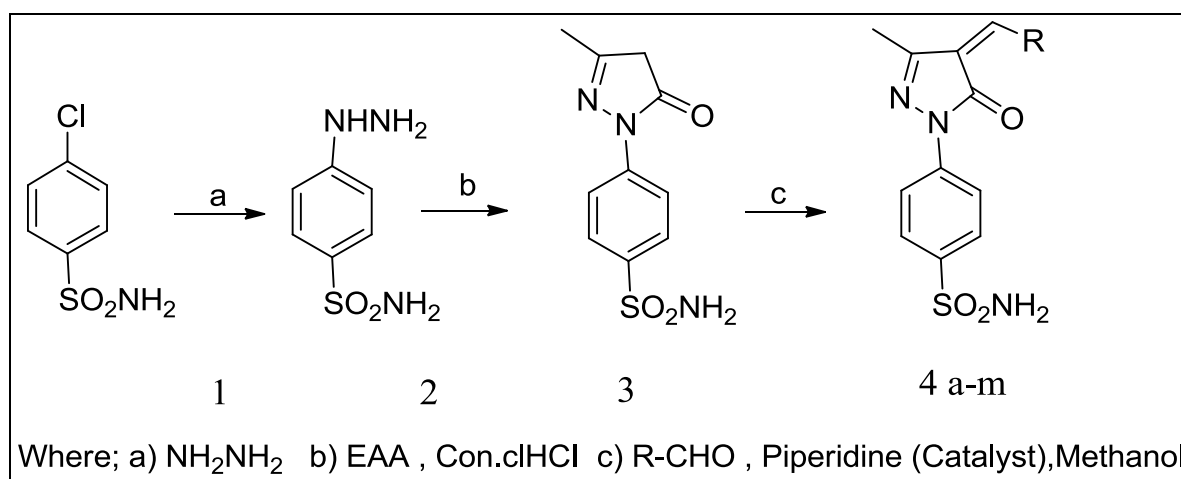
4-hydrazinyl benzene sulfonamide (2) was synthesized by refluxing 4-chlorosulphonamide (compound -1) with hydrazine hydrate. Obtained compound on reaction with ethyl acetoacetate by using few drops of hydrochloric acid as a catalyst gives (compound - 3). The arylidene was synthesized by refluxing (compound-3) with corresponding aldehydes in Methanol using piperidine as a catalyst for an appropriate time to give compound 4a-m different aromatic aldehydes were chosen for synthesized series of diverse pyrazolidinone derivatives.

Compounds with the long chain aldehyde (e.g. 4-(hexyloxy) benzaldehyde) and halogen substituted aldehyde and bulky group (e.g. anthraldehyde; 3,4,5-tri methoxy benzaldehyde; 2,5-dimethoxy benzaldehyde) were also synthesized. Structure of all synthesized compound in (Scheme 1) were confirmed on the basis of IR, ¹H NMR, mass and element analysis. The IR spectrum of compound 4b showed the absorption bands at 3317, 3088, 2976, 1672, 1101, which were due to N-H, Aromatic =CH, CH₃ stretch, C=O, C-O, and absorption band at 1159, 1330 is due to the S=O group.

¹H NMR spectrum of 4b showed singlet at 2.35 which is due to presence of CH₃ group, singlet at 3.37 due to tri methoxy group, proton resonate at 7.73 is due to presence of SO₂NH₂ group, and 8.12, 8.14, 8.20 were due to the aromatic proton. Mass spectrum of compound 4b gives the molecular ion peak at Mass (m/e): 431 (M⁺). Which is confirmed the molecular formula of compound C₂₀H₂₁N₃O₆S. Elemental analysis indicated the % of the elements very close to the theoretical values.

2. 2. General procedure for the synthesis of new derivatives of 4-(unsubstituted phenyl-3-methyl-5-oxo-4, 5-dihydro-1-H-pyrazol-1-yl) benzenesulphonamide

Mixture of 4-(3-methyl-5-oxo-4,5-dihydro-1-*H*-pyrazol-1-yl)benzenesulphonamide (0.01 M) and corresponding aldehyde (0.01M) were taken Methanol using piperidine as a catalyst which are irradiated in microwave under 400 watt for 10-15 minutes. The reaction was monitored by TLC. After completion of reaction, the reaction mixture was allowed to cool. The precipitate formed upon cooling, was filtered off, and washed with diethyl ether and dried in vacuum. Purification of The compound of premeditated series was carried out by crystallization in Methanol.



2. 3. Analytical data

4-(4-(4-(hexyloxy) benzylidene)-3-methyl-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4a)

Crystallization: Ethanol (yield 87 %), colour: yellow crystals. M.P.: 160-162 °C. Anal. Calcd for $\text{C}_{23}\text{H}_{27}\text{N}_3\text{O}_4\text{S}$ (441.17): C, 62.56; H, 6.16; N, 9.52; O, 14.49; S, 7.26. Found: C, 61.759; H, 5.632; N, 9.182. S, 6.911. ^1H NMR (DMSO- d_6) δ ppm: 0.89 (3H, s), 1.32-1.43 (6H, d), 1.74 (2H, s), 2.35 (3H,s), 4.12 (2H, s), 7.151 (2H,s), 7.35 (2H,s), 7.82-7.89 (3H,d), 8.14 (2H,s), 8.7 (2H,s); IR V_{max} cm^{-1} (KBr): 3350 (NH), 2997 (CH_3 stretch), 1737, (C=O ketone), 1643, 1587 (C=C), 1157,1327 (S=O), 1406, (CH_3 bend), 1051 (C-O); Mass (m/e): 441 (M^+).

4-(3-methyl-5-oxo-4-(3,4,5-trimethoxybenzylidene)-4,5-dihydro-1H-pyrazol-1-yl)benzene sulfonamide; (4b)

Crystallization: Ethanol (yield 85 %), colour: orange crystals. M.P.: 182-180 °C. Anal. Calcd for $\text{C}_{20}\text{H}_{21}\text{N}_3\text{O}_6\text{S}$ (431.12): C, 55.67; H, 4.91; N, 9.74; O, 22.25; S, 7.43. Found: C, 49.359; H, 5.632; N, 9.082. S, 10.091. ^1H NMR (DMSO- d_6) δ ppm: 2.34 (3H,s), 3.88 (9H,s), 7.37 (2H,s), 7.82-7.90 (3H, d), 8.11-8.20 (4H,d); IR V_{max} cm^{-1} (KBr): 3317 (NH), 3088 (Aromatic =CH) 2976 (CH_3 stretch), 1672, (C=O ketone), 1500, 1587 (C=C), 1159,1330 (S=O), 1425, (CH_3 bend), 1101 (C-O); Mass (m/e):431 (M^+).

4-(4-(2,5-dimethoxybenzylidene)-3-methyl-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4c)

Crystallization: Ethanol (yield 80 %), colour: orange crystals. M.P.: 178-180 °C. Anal. Calcd for $\text{C}_{19}\text{H}_{19}\text{N}_3\text{O}_5\text{S}$ (401.10): C, 56.85; H, 4.77; N, 10.47; O, 19.93; S, 7.99. Found: C, 50.606; H, 4.707; N, 11.841. S, 11.039. ^1H NMR (DMSO- d_6) δ ppm: 2.36 (3H,s), 3.369 (6H,s), 7.36 (2H,s), 7.40 (1H,s), 7.88-7.91 (2H,d), 8.14-8.16 (2H,d), 8.25-8.30 (3H,m); IR V_{max} cm^{-1} (KBr): 3288 (NH), 3078 (Aromatic =CH) 2833 (CH_3 stretch), 1681, (C=O ketone), 1500, 1591 (C=C), 1159,1323 (S=O), 1460, (CH_3 bend), 1095(C-O); Mass (m/e): 401 (M^+).

4-(4-(anthracen-9-ylmethylene)-3-methyl-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4d)

Crystallization: Ethanol (yield 83 %), colour : Red crystals. M.P.: 176-178 °C. Anal. Calcd for $\text{C}_{25}\text{H}_{19}\text{N}_3\text{O}_3\text{S}$ (441.10): C, 68.01; H, 4.34; N, 9.52; O, 10.87; S, 7.26 Found: C, 66.914; H, 5.149; N, 9.564. S, 9.620. ^1H NMR (DMSO- d_6) δ ppm: 1.24 (3H,s),7.30-7.4 (2H,d), 7.58-

7.62 (4H,dd), 7.93 (1H,s), 8.14-8.16 (2H,s), 8.19-8.77 (5H,m), 8.81-8.88 (2H,m); IR V_{\max} cm^{-1} (KBr): 3345, 3223 (NH), 3093 (Aromatic =CH) 2833 (CH_3 stretch), 1699 (C=O ketone), 1618, 1591 (C=C), 1155,1344 (S=O), 1496, (CH_3 bend), Mass (m/e): 441 (M^+).

4-(4-(3,4-dimethoxybenzylidene)-3-methyl-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4e)

Crystallization: Ethanol (yield 80 %), colour: red-orange crystals. M.P.: 180-182 °C. Anal. Calcd for $\text{C}_{19}\text{H}_{19}\text{N}_3\text{O}_5\text{S}$ (401.10): C, 56.85; H, 4.77; N, 10.47; O, 19.93; S, 7.99. Found: C, 51.609; H, 6.016; N, 8.777. S, 11.039. ^1H NMR (DMSO-d_6) δ ppm: 1.36 (3H,s), 3.48 (6H, s), 7.32 (2H,s), 7.38 (1H,s), 7.80-7.86 (2H, d), 8.12-8.14 (2H,d), 8.28-8.39 (3H,m); IR V_{\max} cm^{-1} (KBr): 3284 (NH), 3097 (Aromatic =CH) 1693, (C=O ketone), 1552, 1597 (C=C), 1155,1330 (S=O), 1095 (C-O); Mass (m/e):401 (M^+).

4-(3-methyl-5-oxo-4-(thiophen-2-ylmethylene)-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4f)

Crystallization: Ethanol (yield 84 %), colour: yellow crystals. M.P.: 172-174 °C. Anal. Calcd for $\text{C}_{15}\text{H}_{13}\text{N}_3\text{O}_3\text{S}_2$ (347.040): C, 51.86; H, 3.77; N, 12.10; O, 13.82; S, 18.46. Found: C, 51.911; H, 4.718; N, 12.218. S, 20.694. ^1H NMR (DMSO-d_6) δ ppm: 2.36 (3H,s), 7.36-7.40 (3H,d), 7.89-7.91 (2H,d), 8.14-8.16 (2H,d), 8.25-8.29 (3H,m); IR V_{\max} cm^{-1} (KBr): 3284 (NH), 3097 (Aromatic =CH) 1693, (C=O ketone), 1552, 1597 (C=C), 1155,1330 (S=O), 1095 (C-O); Mass (m/e): 347 (M^+).

4-(4-(4-chlorobenzylidene)-3-methyl-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4g)

Crystallization: Ethanol (yield 76 %), colour: yellow crystals. M.P.: 194-192 °C. Anal. Calcd for $\text{C}_{17}\text{H}_{14}\text{ClN}_3\text{O}_3\text{S}$ (375.044): C, 54.33; H, 3.75; Cl, 9.43; N, 11.18; O, 12.77; S, 8.53. Found: C, 53.911; H, 4.118; N, 11.218. S, 9.694. ^1H NMR (DMSO-d_6) δ ppm: 2.12 (3H,s), 7.33 (2H,s), 7.75-7.82 (3H, d), 8.25-8.39 (6H,m); IR V_{\max} cm^{-1} (KBr): 3279 (NH), 3080 (Aromatic =CH) 1707, (C=O ketone), 1487, 1593 (C=C), 1155,1335 (S=O); Mass (m/e): 375 (M^+).

4-(3-methyl-4-(4-methylbenzylidene)-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4h)

Crystallization: Ethanol (yield 79 %), colour: yellow crystals. M.P.: 168-166 °C. Anal. Calcd for $\text{C}_{18}\text{H}_{17}\text{N}_3\text{O}_3\text{S}$ (355.02): C, 60.83; H, 4.82; N, 11.82; O, 13.50; S, 9.02. Found: C, 59.911; H, 4.781; N, 12.218. S, 8.694. ^1H NMR (DMSO-d_6) δ ppm: 2.15 (3H,s), 6.99 (3H,s), 7.12 (2H,s), 7.14-7.24 (2H,m), 7.68-8.2(6H,m); IR V_{\max} cm^{-1} (KBr): 3285 (NH), 3086 (Aromatic =CH) 1772, (C=O ketone), 1494, 1593 (C=C), 1159,1329 (S=O); Mass (m/e): 375 (M^+).

4-(3-methyl-4-(naphthalen-1-ylmethylene)-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4i)

Crystallization: Ethanol (yield 76 %), colour: yellow crystals. M.P.: 186-188 °C. Anal. Calcd for $\text{C}_{21}\text{H}_{17}\text{N}_3\text{O}_3\text{S}$ (391.10): C, 64.43; H, 4.38; N, 10.73; O, 12.26; S, 8.19. Found: C, 62.45; H, 4.118; N, 11.20. S, 8.60. ^1H NMR (DMSO-d_6) δ ppm: 1.28 (3H,s), 7.28-7.4 (2H,d), 7.48-7.52 (4H,m), 8.11-8.14 (2H,s), 8.39-8.48 (6H,m); IR V_{\max} cm^{-1} (KBr): 3254 (NH), 3186 (Aromatic =CH) 1674, (C=O ketone), 1494, 1595 (C=C), 1159,1325 (S=O); Mass (m/e): 391(M^+).

4-(3-methyl-4-(4-nitrobenzylidene)-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4j)

Crystallization: Ethanol (yield 68 %), colour: dark brown crystals. M.P.: 170-172 °C. Anal. Calcd for C₁₇H₁₄N₄O₅S (386.068): C, 52.84; H, 3.65; N, 14.50; O, 20.70; S, 8.30. Found: C, 50.75; H, 4.00; N, 15.68. S, 9.50. ¹H NMR (DMSO-d₆) δ ppm : 2.18 (3H,s), 7.28 (2H,s), 7.78-7.91 (3H, m), 8.40-8.69 (6H,m); IR V_{max} cm⁻¹ (KBr): 3342 (NH), 3211 (Aromatic =CH) 1691, (C=O ketone), 1487, 1591 (C=C), 1159,1342 (S=O); Mass (m/e): 386.(M⁺).

4-(4-(3-methoxybenzylidene)-3-methyl-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4k)

Crystallization: Ethanol (yield 80 %), colour: orange crystals. M.P.: 252-250 °C. Anal. Calcd for C₁₈H₁₇N₃O₄S (371.09): C, 58.21; H, 4.61; N, 11.31; O, 17.23; S, 8.63 Found: C, 55.45; H, 5.24; N, 12.45. S, 10.24. ¹H NMR (DMSO-d₆) δ ppm: 2.32 (3H,s), 3.4 (3H,s), 6.77 (2H,s), 7.11-7.39 (4H,m), 7.52-8.12 (5H,m); IR V_{max} cm⁻¹ (KBr): 3257 (NH), 3078 (Aromatic =CH) 2922 (CH₃ stretch), 1680, (C=O ketone), 1595 (C=C), 1159,1332 (S=O), 1460, (CH₃ bend), 1037 (C-O); Mass (m/e) :371(M⁺).

4-(4-(4-methoxybenzylidene)-3-methyl-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4l)

Crystallization: Ethanol (yield 74 %), colour: orange crystals. M.P.: 168-170 °C. Anal. Calcd for C₁₈H₁₇N₃O₄S (371.09): C, 58.21; H, 4.61; N, 11.31; O, 17.23; S, 8.63 Found: C, 58.00; H, 4.24; N, 12.00. S, 9.24. ¹H NMR (DMSO-d₆) δ ppm: 2.35 (3H,s), 3.86 (3H,s), 6.78-6.80 (2H,d), 7.14-7.16 (2H,d), 7.36-7.88 (4H, m), 8.13-8.72 (3H,m); IR V_{max} cm⁻¹ (KBr): 3341 (NH), 3261 (Aromatic =CH) 2922 (CH₃ stretch), 1676, (C=O ketone), 1577(C=C), 1155, 1327 (S=O), 1460, (CH₃ bend), 1030 (C-O); Mass (m/e): 371 (M⁺).

4-(4-(3-bromo-4-methoxybenzylidene)-3-methyl-5-oxo-4,5-dihydro-1H-pyrazol-1-yl)benzenesulfonamide; (4m)

Crystallization: Ethanol (yield 83 %), colour: brown crystals. M.P.: 182-180 °C. Anal. Calcd for C₁₈H₁₆BrN₃O₄S (449.00): C, 48.01; H, 3.58; Br, 17.74; N, 9.33; O, 14.21; S, 7.12. Found: C, 55.45; H, 5.24; N, 12.45. S, 10.24. ¹H NMR (DMSO-d₆) δ ppm: 2.34 (3H,s), 2.89 (3H,s), 7.02-7.04 (2H,d), 7.12-7.16 (3H,m), 7.66-7.89 (3H,m), 7.92-8.12 (2H,m); IR V_{max} cm⁻¹ (KBr): 3252 (NH), 3078 (Aromatic =CH) 2922 (CH₃ stretch), 1666, (C=O ketone), 1589 (C=C), 1157,1332 (S=O), 1460, (CH₃ bend), 1099(C-O); Mass (m/e): 449 (M⁺).

Table 1. Optimization of (%) yield for the microwave assisted synthesis of 4a, 4b, 4c and 4e using different solvents and catalyst.

Entry as	Solvent	catalyst	Time ^a (min) microwave	Yield ^b %
4a	Methanol	Piperidine	10	88
		TEA	15	75
4a	Chloroform	Piperidine	18	80
		TEA	25	74

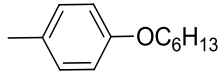
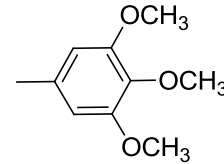
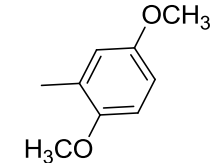
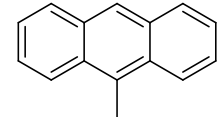
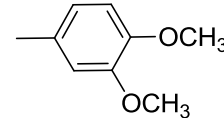
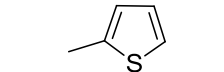
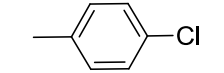
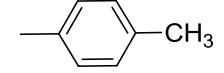
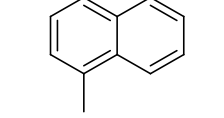
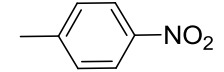
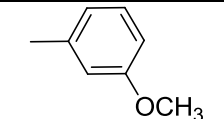
4a	Ethyl Acetate	Piperidine	35	68
		TEA	38	55
4a	DMF	K ₂ CO ₃	8	65
		C ₅ CO ₃	8	68
4a	Methanol	HCl	30	48
4a	Acetic acid	-	25	45
4b	Methanol	Piperidine	8	92
		TEA	10	85
4b	Chloroform	Piperidine	15	82
		TEA	18	76
4b	Ethyl Acetate	Piperidine	20	70
		TEA	22	69
4b	DMF	K ₂ CO ₃	5	69
		C ₅ CO ₃	5	68
4b	Methanol	HCl	28	56
4b	Acetic acid	-	30	52
4c	Methanol	Piperidine	12	90
		TEA	10	88
4c	Chloroform	Piperidine	18	86
		TEA	15	80
4c	Ethyl Acetate	Piperidine	20	74
		TEA	20	72
4c	DMF	K ₂ CO ₃	6	68
		C ₅ CO ₃	6	64
4c	Methanol	HCl	25	50
4c	Acetic acid	-	30	54
4e	Methanol	Piperidine	8	79
		TEA	12	72

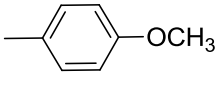
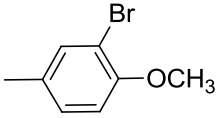
4e	Chloroform	Piperidine	14	70
		TEA	18	64
4e	Ethyl Acetate	Piperidine	25	58
		TEA	28	54
4e	DMF	K ₂ CO ₃	6	54
		C _s CO ₃	6	52
4e	Methanol	HCl	30	48
4e	Acetic acid	-	25	45

Table 2. Comparison of yield (%) of the Arylidene derivatives obtained using microwave assisted as well as conventional method of synthesis.

compound	Yield (%) obtained by the method of synthesis	
	Microwave assisted method	Conventional method
4a	88	74
4b	92	75
4c	90	78
4d	82	62
4e	88	76
4f	79	63
4g	78	68
4h	70	48
4i	78	63
4j	77	54
4k	85	66
4l	86	62
4m	80	68

Table 3. Physical data of the pyrazolone derivatives obtained using microwave assisted method of synthesis 4(a-m).

Compounds	Substitution R	Molecular formula	Molecular weight	Color	Yield (%)	M.P. °C
4a		C ₂₃ H ₂₇ N ₃ O ₄ S	441.17	yellow	87 %	160-162
4b		C ₂₀ H ₂₁ N ₃ O ₆ S	431.12	orange	85 %	182-180
4c		C ₁₉ H ₁₉ N ₃ O ₅ S	401.10	orange	80 %	178-180
4d		C ₂₅ H ₁₉ N ₃ O ₃ S	441.10	Red	83 %	176-178
4e		C ₁₉ H ₁₉ N ₃ O ₅ S	401.10	red-orange	80 %	180-182
4f		C ₁₅ H ₁₃ N ₃ O ₃ S ₂	347.04	Yellow	84 %	172-174
4g		C ₁₇ H ₁₄ ClN ₃ O ₃ S	375.04	Yellow	76 %	194-192
4h		C ₁₈ H ₁₇ N ₃ O ₃ S	355.02	Yellow	79 %	168-166
4i		C ₂₁ H ₁₇ N ₃ O ₃ S	391.10	Yellow	76 %	186-188
4j		C ₁₇ H ₁₄ N ₄ O ₅ S	386.06	dark brown	68 %	252-250
4k		C ₁₈ H ₁₇ N ₃ O ₄ S	371.09	Orange	80 %	174-176

4l		$C_{18}H_{17}N_3O_4S$	371.09	Orange	74 %	168-170
4m		$C_{18}H_{16}BrN_3O_4S$	449.00	Brown	83 %	182-180

We have synthesized 4a, 4b, 4c, and 4e in diverse solvents like Methanol, chloroform, Ethyl Acetate, and DMF by using different basic catalyst such as piperidine, TEA as an organic base and K_2CO_3 , C_5CO_3 as an inorganic base under microwave irradiation. Synthesis was also carried out in acidic media by using methanol as solvent, few drops of HCl as catalyst. We have also use gl. Acetic acid for the optimisation of yield. We observed that reaction carried out in DMF and inorganic base taken lesser time but yield observed very low, but in case of methanol as solvent and piperidine as basic organic catalyst, instead of, chloroform, DMF and Ethyl Acetate as solvents and other acidic or basic catalyst. The time taken for the completion of reaction was 3-9 minutes and the % yields observed was 85-90 % under microwave assisted method. Furthermore, we carried out the same synthesis by using methanol as solvent and piperidine as catalyst under conventional method. The time taken for synthesis of compounds 4a-m by conventional method was around 6-8 hours and the % yield observed was about 60-75 %. All the synthesized compounds were characterized by TLC, Melting point, elemental analysis, IR 1H NMR.

3. CONCLUSION

Novel Sulphonamide Arylidene derivatives were synthesized in rationally good yields by microwave assisted method. This is environmentally compassionate technique gives higher fraction of yields and lesser reaction time and easy work up method.

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