



Synthesis, Characterization, Antimicrobial and Antifungal Screening of Some Novel Benzene Sulfonamide Derivatives

Rathod CP*, Dhawale SC, Pekamwar SS, Kadam NR, Rekhawar MU

¹School of Pharmacy S.R.T.M.U. Nanded.

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ABSTRACT

The present work reports the possible utility of the synthesis of some novel 4-(2-amino-3-cyano-4-(substituted-aryl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl)benzene sulfonamide (1a–1u) starting with 4-(3-oxo-cyclohex-1-enylamino)benzenesulfonamide and 4-(cyclohexenylamino) benzenesulfonamide in the synthesis of some novel 4-(quinolin-1-yl) benzenesulfonamide derivatives. The structures of the synthesized compounds were confirmed by IR, ¹H NMR, ¹³C NMR and mass spectral data. Some of the newly synthesized compounds were evaluated for their in vitro antibacterial and antifungal activity.

KEYWORDS

Synthesis, sulfonamide, cyclohexanone 1, 3 dione, antibacterial, antifungal.

INTRODUCTION

Sulfonamides form a significant class of compounds in medicinal and pharmaceutical chemistry with several biological applications. The sulfonamides were the first effective chemotherapeutic agents to be available in safe therapeutic dosage ranges. The sulfonamide drugs were the first effective chemotherapeutic agents to be employed systemically for the prevention and cure of bacterial infection in human beings. Sulfonamides possess many types of biological activities and representatives of this class of pharmacological agents are widely used as antibacterial^(1,2,3), antifungal^(2,3), anticancer^(4,5,6), antioxidant⁽⁷⁾, anti-inflammatory^(7,8), antimalarial⁽⁹⁾, anti-HIV integrase⁽¹⁰⁾, and anti-carbonic anhydrase activity⁽¹¹⁾ among others. Herein we disclose the screening results of their antibacterial and antifungal activity of some of the compounds.

RESULTS AND DISCUSSION

Chemistry

The present work reports the possible utility of 4-(3-oxocyclohex-1-enylamino) benzenesulfonamide in the synthesis of 4-(quinolin-1-yl) benzenesulfonamide derivatives 1a-1u. Enaminone C was obtained by condensation of cyclohexan-1,3-dione. Compounds 1a–1u were synthesized by the one-pot condensation of the aldehyde, malononitrile and enaminone 3 in a molar ratio of (1:1:1) in refluxing ethanol containing triethylamine as catalyst.

Biological Screening

In-vitro Antibacterial Screening of Synthesized Compounds

The *in vitro* antimicrobial activity of the newly synthesized benzenesulfonamide derivatives was tested against *B. subtilis*, *E. coli*, *S. typhi* and *S. aureus* using nutrient agar medium. The antifungal activity of synthesized Benzene sulfonamide derivatives was tested against *Aspergillus flavus* and *Aspergillus niger* using Czapek Dox Agar Medium by using disc diffusion method and *Candida albicans* using Yeast extract malt extract agar medium by using

*Address for Correspondence:

Rathod C.P

School of pharmacy S.R.T.M.U.

Nanded,

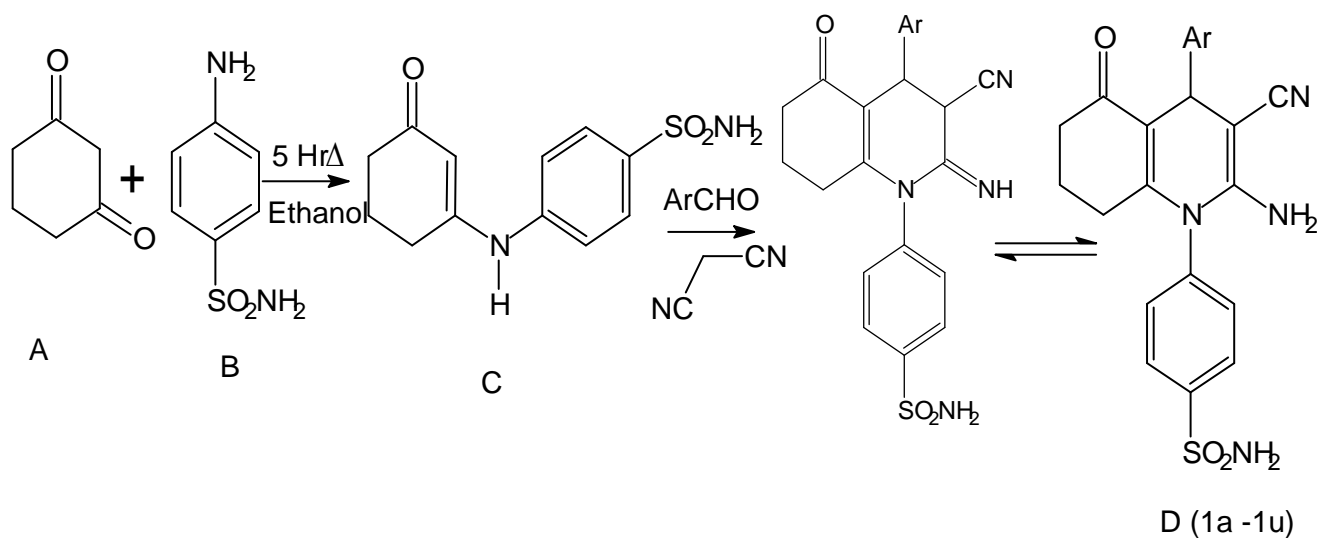
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E-Mail Id: cprathod.mpharma@rediffmail.com

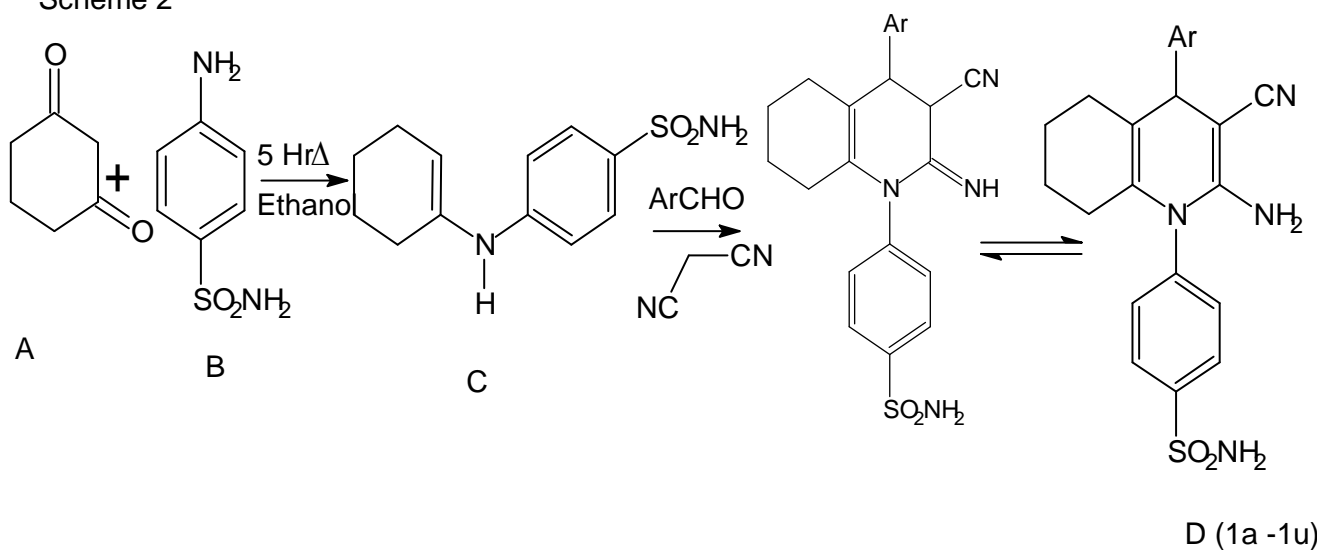
Agar plate method. The results are given in Tables as follows.

Scheme of synthesis:

Scheme 1



Scheme 2



1a, Ar = C₆H₄CH₃-2
 1b, Ar = C₆H₄CH₃-3
 1c, Ar = C₆H₄CH₃-4
 1d, Ar = C₆H₃(CH₃)₂-2,4
 1e, Ar = C₆H₄F-2
 1f, Ar = C₆H₃F₂-3,4
 1g, Ar = C₆H₄CH₃-2

1h, Ar = C₆H₄CH₃-3
 1i, Ar = C₆H₄CH₃-4
 1j, Ar = C₆H₃(CH₃)₂-2, 4
 1k, Ar = C₆H₄F-2
 1l, Ar = C₆H₄F-3
 1m, Ar = C₆H₄F-4
 1n, Ar = C₆H₃F₂-3,4

1o, Ar = C₆H₄OH-3
 1p, Ar = C₆H₄NO₂-3
 1q, Ar = C₆H₄NO₂-4
 1r, Ar = C₆H₄Cl-4
 1s, Ar = C₆H₃Cl₂-2,4
 1t, Ar = C₆H₃(CH₃)₂-3,4
 1u, Ar = C₆H₃(CH₃)₂-2,4

Table 1: Characterization data of synthesized novel benzenesulfonamide derivatives

| Comp | Molecular Formula | M.P. (°C) | Mol. Wt. | App. | R _f | Solubility | Yield (%) |
|------|---|-----------|----------|-----------------|----------------|------------------|-----------|
| 1a | C ₂₃ H ₂₂ N ₄ O ₃ S | 150-152 | 434.510 | Brownish Yellow | 0.54 | Methanol DMSO | 75.94 |
| 1b | C ₂₃ H ₂₂ N ₄ O ₃ S | 158-160 | 434.510 | Brown | 0.46 | Methanol DMSO | 78.30 |
| 1c | C ₂₃ H ₂₂ N ₄ O ₃ S | 162-164 | 434.510 | Brown | 0.34 | Methanol DMSO | 82.62 |
| 1d | C ₂₄ H ₂₄ N ₄ O ₃ S | 170-172 | 448.537 | Pink | 0.38 | Methanol DMSO | 69.51 |
| 1e | C ₂₂ H ₁₉ FN ₄ O ₃ S | 180-182 | 438.474 | Pink | 0.45 | Methanol DMSO | 75.43 |
| 1f | C ₂₂ H ₁₈ F ₂ N ₄ O ₃ S | 152-154 | 456.465 | Brown | 0.52 | Methanol DMSO | 76.22 |
| 1g | C ₂₃ H ₂₄ N ₄ O ₂ S | 152-154 | 420.526 | Orange | 0.48 | Methanol DMSO | 72.71 |
| 1h | C ₂₃ H ₂₄ N ₄ O ₂ S | 150-152 | 420.526 | Yellowish Grey | 0.46 | Methanol DMSO | 69.42 |
| 1i | C ₂₃ H ₂₄ N ₄ O ₂ S | 152-154 | 420.526 | Orange | 0.42 | Methanol DMSO | 78.80 |
| 1j | C ₂₄ H ₂₆ N ₄ O ₂ S | 152-154 | 434.553 | Yellowish Brown | 0.35 | Methanol DMSO | 78.04 |
| 1k | C ₂₂ H ₂₁ FN ₄ O ₂ S | 222-224 | 424.490 | Brown | 0.48 | Methanol DMSO | 72.08 |
| 1L | C ₂₂ H ₂₁ FN ₄ O ₂ S | 154-156 | 424.490 | Pink | 0.52 | Methanol DMSO | 76.64 |
| 1m | C ₂₂ H ₂₁ FN ₄ O ₂ S | 160-162 | 424.490 | Yellowish Brown | 0.50 | Methanol DMSO | 78.66 |
| 1n | C ₂₂ H ₂₀ F ₂ N ₄ O ₂ S | 148-150 | 442.481 | Pink | 0.46 | Methanol DMSO | 68.26 |
| 1o | C ₂₂ H ₂₀ N ₄ O ₄ S | 220-222 | 436.483 | Brown | 0.42 | Methanol DMSO | 73.24 |
| 1p | C ₂₂ H ₁₉ N ₅ O ₅ S | 212-214 | 465.481 | Yellowish Brown | 0.45 | Methanol DMSO | 70.82 |
| 1q | C ₂₂ H ₁₉ N ₅ O ₅ S | 284-286 | 465.481 | Yellowish Brown | 0.52 | Methanol DMSO | 76.35 |
| 1r | C ₂₂ H ₁₉ ClN ₄ O ₃ S | 280-282 | 454.929 | Yellowish Brown | 0.38 | Methanol DMSO | 68.58 |
| 1s | C ₂₂ H ₁₈ Cl ₂ N ₄ O ₃ S | 290-292 | 489.374 | Brown | 0.58 | Methanol | 67.87 |
| 1t | C ₂₄ H ₂₄ N ₄ O ₅ S | 148-150 | 480.536 | Yellowish Brown | 0.40 | Methanol DMSO | 72.67 |
| 1u | C ₂₄ H ₂₄ N ₄ O ₅ S | 174-176 | 480.536 | Grey | 0.46 | Methanol DMSO | 65.84 |

Table 2: Antibacterial activity of some synthesized benzene sulfonamide derivatives

| Sr. No. | Compound Code | Zone of Inhibition in mm | |
|---------|---------------|--------------------------|--------------------------|
| | | <i>Escherichia Coli</i> | <i>Bacillus Subtilis</i> |
| 1 | 1a | 22 | 27 |
| 2 | 1e | 20 | 24 |
| 3 | 1g | 24 | 32 |
| 4 | 1h | 15 | 18 |
| 5 | 1i | 26 | 28 |
| 6 | 1j | 18 | 24 |
| 7 | 1l | 34 | 15 |
| 8 | 1m | 38 | 22 |
| 9 | 1n | 22 | 28 |
| 10 | 1q | 24 | 30 |
| 11 | 1r | 16 | -- |
| 12 | 1u | -- | 36 |
| 13 | Standard. | 32 | 30 |
| 14 | Control | 06 | 06 |

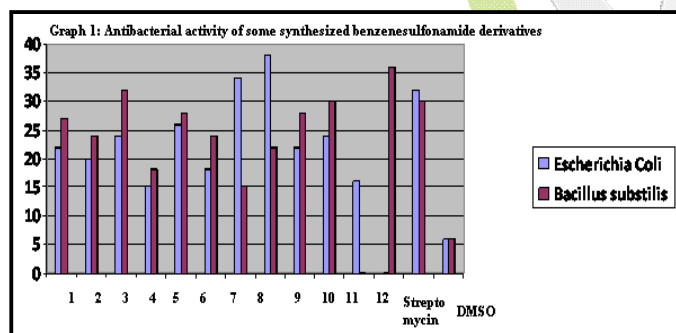


Table 3: Antibacterial Data of synthesized compounds using agar plate method.

| Sr. No. | Compound Code | Zone of Inhibition in mm | |
|---------|---------------|--------------------------|------------------------------|
| | | <i>Salmonella typhi</i> | <i>Staphylococcus aureus</i> |
| 1 | 1f | 22 | 27 |
| 2 | 1s | 20 | 24 |
| 3 | 1p | 24 | 32 |

| | | | |
|---|------------|----|----|
| 4 | 1t | 15 | 18 |
| 5 | 1o | 26 | 28 |
| 6 | Blank | 18 | 24 |
| 7 | Penicillin | 18 | 36 |

Antifungal Activity

The antifungal activity of synthesized Benzenesulfonamide derivatives was tested against *Aspergillus flavus* and *Aspergillus Niger* using Czapek Dox Agar Medium by using disc diffusion method. Fluconazole were used as standard drug for antibacterial antifungal studies. The results of the antifungal screening for the compounds were given in Tables.

Table 4: Antifungal activity of some synthesized benzene sulfonamide derivatives.

| Sr. No. | Compound Code | Zone of Inhibition in mm | |
|---------|---------------|---------------------------|--------------------------|
| | | <i>Aspergillus Flavus</i> | <i>Aspergillus Niger</i> |
| 1 | 1a | 22 | 40 |
| 2 | 1e | 26 | 25 |
| 3 | 1g | 10 | 18 |
| 4 | 1h | 32 | 24 |
| 5 | 1i | 30 | 22 |
| 6 | 1j | 25 | 30 |
| 7 | 1l | 28 | -- |
| 8 | 1m | 24 | 27 |
| 9 | 1n | -- | 12 |
| 10 | 1q | 25 | 30 |
| 11 | 1r | -- | 21 |
| 12 | 1u | 30 | 36 |
| 13 | (Fluconazole) | 28 | 26 |
| 14 | (DMSO) | 06 | 06 |

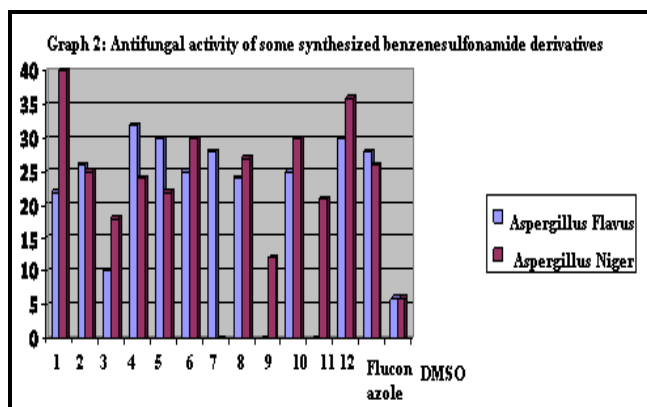


Table 5: Anti-fungal data of synthesized compounds for candida albicans.

| Sr No. | Compound code | Candida Albicans (Zone of Inhibition in mm) |
|--------|------------------------|---|
| 1 | 1a | -- |
| 2 | 1b | 15 |
| 3 | 1c | -- |
| 4 | 1d | 16 |
| 5 | 1u | 13 |
| 6 | Blank (DMSO) | -- |
| 7 | Standard (Fluconazole) | 46 |

The synthesized benzenesulfonamide derivatives were evaluated for their *in vitro* antibacterial and antifungal potential. The antibacterial activity of synthesized compounds was evaluated against *E. coli* and *B. subtilis* using disc diffusion method and results are presented as zone of inhibition (mm) in table 2. The streptomycin was used as a control. All compounds have shown positive antibacterial activity against *E. coli* and *B. subtilis*. The compound 1m (38 mm) showed greatest activity among the all synthesized compounds against *E. coli*, while the compounds 1u (36 mm) showed stronger activity than the standard against *B. subtilis*. The compound 1h (15 mm) showed the weakest activity among the synthesized compounds against *E. coli*, while the compounds 1L (15 mm) showed the weakest activities against *B. subtilis*. The antibacterial activity of five synthesized compound was carried out by using Agar Cup Method against

penicillin as a standard and results are presented as zone of inhibition (mm) in table 3. The compound 1o (26 mm) showed greatest activity among the all synthesized compounds against *Salmonella typhi*, while the compounds 1p (32mm) showed stronger activity than the standard against *Staphylococcus aureus*. The compound 1t (15 mm) showed the weakest activity among the synthesized compounds against *Salmonella typhi*, while the compounds 1t (18 mm) showed the less significant activity against *Staphylococcus aureus*.

The antifungal activity of synthesized compounds was evaluated against *A. niger* and *A. flavous* using disc diffusion method and results are presented as zone of inhibition (mm) in table 4. The Fluconazole was used as a standard and results are presented as zone of inhibition (mm) in table 4. All compounds have shown positive antifungal activity against *A. niger*. The compound 1h (32 mm) showed greater activity than the standard Fluconazole (28mm) against *A. flavous* while the compounds 1a (40 mm) showed higher activity than the standard Fluconazole (26mm) against *A. niger*. The compounds 1g (10 mm) showed the weakest activity among the synthesized compounds against *A. flavous* while the compound 1n (12mm) showed the weakest activities against *A.niger*. The five synthesized compound were screened for the antifungal activity against fungal species for candida albicans and results are presented as zone of inhibition (mm) in table 5. Compounds 1b (15mm), 1d (16mm), and 1u (13mm) showed the moderate antifungal activities among the selected five synthesized compounds.

EXPERIMENTAL

Chemistry

Melting points ($^{\circ}\text{C}$, uncorrected) were determined in open capillaries on a Gallenkamp melting point apparatus (Sanyo Gallenkamp, Southborough, UK). Precoated silica gel plates (silica gel 0.25 mm, 60G F254; Merck, Germany) were used for thin layer chromatography, dichloromethane/methanol (9.5:0.5) mixture was used as a developing

solvent system and the spots were visualized by ultraviolet light and/or iodine. IR Spectra were recorded in KBr disk on "Shimadzu FTIR model no. I.R.Affinity 1" and are reported in cm^{-1} . ^1H NMR and ^{13}C NMR spectral were recorded using NMR Spectrometer. Varian Inc, Mercury Plus 300 MHz NMR Spectrometer with tetramethylsilane (TMS) as the internal standard in CDCl_3 . and peak multiplicities are designed as follows: s, singlet; d, doublet; t, triplet; m, multiplet. Mass spectra were reported at "Indian Institute of Technology, Bombay" with Mass spectrometer, Make: Varian Inc. USA; Model: 410 prostar Binary LC with 500 MS IT PDA Detectors. Chemicals and solvents were purchased from corresponding companies (SDFCL, Merk Pvt.Ltd.) and were used in the experimentation without further purification.

4-(3-Oxo-cyclohex-1-enylamino) benzenesulfonamide (C):

A mixture of cyclohexane-1,3-dione or Cyclohexanone (A) (1.12 g, 0.01 mol) and sulfanilamide (B) (1.72 g, 0.01 mol) in ethanol (30 mL) was refluxed for 3 h. The reaction mixture was cooled and then poured onto cold water the obtained solid was crystallized from ethanol to give 3. Yield, 80%; m.p. 232–234.

4-[2-Amino-3-cyano-4-(substituted-aryl)-5-oxo-5, 6, 7, 8- tetrahydroquinolin-1(4H)-yl]benzenesulfonamide (1a–1u) :

A solution of enaminone (C) (2.66 g, 0.01 mol), the appropriate aldehyde (0.01 mol) and malononitrile (0.66 g, 0.01 mol) in absolute ethanol (30 mL) containing 3 drops of triethylamine, was refluxed for 5 h. The solid obtained after concentration and cooling, was filtered and crystallized from dioxane to give 1a–1u, respectively.

1. 4-(2-amino-3-cyano-4 (2-methylphenyl)-5-oxo-5, 6, 7, 8-tetrahydroquinolin-1(4H)-yl) benzene sulfonamide, **1a**. Yield 75.94% ; mp 150-152°C; IR (KBr, cm^{-1}) : 3182, 3265, 3356, 3469 (NH_2 , stretching), 2281, 2339, ($\text{C}\equiv\text{N}$, stretching), 1141, 1184, 1247, 1317 (SO_2 , stretching), 3035, 3062 (Ar-CH), 2949 (aliphatic- CH), 1614 (C = O). Anal. Calcd. For

$\text{C}_{23}\text{H}_{22}\text{N}_4\text{O}_3\text{S}$: C, 63.58; H, 5.10; N, 12.89; O, 11.05; S, 7.38.

2. 4-(2-amino-3-cyano-4-(3 methylphenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide, **1b**. Yield 78.30 % ; mp 158-160 °C IR (KBr, cm^{-1}) : 3211, 3280, 3356 (NH_2 , stretching), 2117, 2167, 2293 ($\text{C}\equiv\text{N}$, stretching), 1136, 1149, 1165, 1193, 1259, 1284, 1301 (SO_2 , stretching), 3028, 3045 (Ar-CH) , 2954 (aliphatic- CH), 1587, 1614, 1633, 1643 (C= O). Anal. Calcd. For $\text{C}_{23}\text{H}_{22}\text{N}_4\text{O}_3\text{S}$: C, 63.58; H, 5.10; N, 12.89; O, 11.05; S, 7.38.

3. 4-(2-amino-3-cyano-4-(4-methylphenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1c**. Yield 82.62 % ; mp 162-164 °C IR (KBr, cm^{-1}) : 3186, 3265, 3356 (NH_2 , stretching), 2189 ($\text{C}\equiv\text{N}$, stretching), 1141, 1184, 1247, 1317 (SO_2 , stretching), 3034, 3061 (Ar-CH) , 2949 (aliphatic- CH), 1573, 1612 (C= O). Anal. Calcd. For $\text{C}_{23}\text{H}_{22}\text{N}_4\text{O}_3\text{S}$: C, 63.58; H, 5.10; N, 12.89; O, 11.05; S, 7.38.

4. 4-(2-amino-3-cyano-4-(2,4-dimethylphenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1d**. Yield 69.51 %; mp 170-172 °C IR (KBr, cm^{-1}) : 3372, 3358 (NH_2 , stretching), 2222, 2191, 2169, ($\text{C}\equiv\text{N}$, stretching), 1157 , 1182, 1251, 11296, 1309 (SO_2 , stretching), 3039, 3084 (Ar-CH), 2962, 2981 (aliphatic- CH), 1612 (C= O). Anal. Calcd. For $\text{C}_{24}\text{H}_{24}\text{N}_4\text{O}_3\text{S}$: C, 60.26; H, 4.37; F, 4.33; N, 12.78; O, 10.95; S, 7.31.

5. 4-(2-amino-3-cyano-4-(2-fluorophenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1e**. Yield 75.43 %; mp 180-182 °C IR (KBr, cm^{-1}) : 3356 (NH_2 , stretching), 2220, ($\text{C}\equiv\text{N}$, stretching), 1184 , 1247, 1317, (SO_2 , stretching), 3034, 3062 (Ar-CH), 2949, (aliphatic- CH), 1612, 1687 (C= O). Anal. Calcd. For $\text{C}_{22}\text{H}_{19}\text{FN}_4\text{O}_3\text{S}$: C, 64.27; H, 5.39; N, 12.49; O, 10.70; S, 7.15.

6. 4-(2-amino-3-cyano-4-(3,4-difluorophenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1f**. Yield 76.22 %; mp 152-

154 °C **IR** (KBr, cm⁻¹) : 3120, 3230, 3325 (NH₂, stretching), 2127, 2179, 2289 (C≡N, stretching), 1141, 1161, 1186, 1222, 1257, 1284, (SO₂, stretching), 3028, 3074, (Ar-CH), 2937, 2947, 2972 (aliphatic-CH), 1573, 1612, 1645 (C=O). Anal. Calcd. For C₂₂H₁₈F₂N₄O₃S: C, 57.89; H, 3.97; F, 8.32; N, 12.27; O, 10.52; S, 7.02.

7. 4-(2-amino-3-cyano-4-(2-methylphenyl)-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1g**. Yield 72.71%; mp 152-154 °C **IR** (KBr, cm⁻¹) : 3462, 3477, 3373 (NH₂, stretching), 2281, 2233, 2204 (C≡N, stretching), 1186 (SO₂, stretching), 3066, 3089 (Ar-CH), 2962 (aliphatic-CH). **MS** (m/z): 420. **¹H NMR**: Solvent (CD₃OD) δ 3.321-3.300 [s, 2H, R-NH₂], δ 4.846 [m, 8H, AR-NH₂], δ 6.710-6.662 [m, 4H, AR-H], δ 7.607-7.559 [m, 4H, ARH]. **¹³C NMR**: δ153.62 (C in aromatic), δ131.330 (C=C in alkenes), δ129.04 (C=C in alkenes), δ 114.60 (C=C), δ 114.50 (C=C), δ 49.96 (C-C). Anal. Calcd. For C₂₃H₂₄N₄O₂S: C, 65.69; H, 5.75; N, 13.32; O, 7.61; S, 7.62.

8. 4-(2-amino-3-cyano-4-(3-methylphenyl)-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1h**. Yield 69.42%; mp 150-152 °C **IR** (KBr, cm⁻¹) : 3462, 3375 (NH₂, stretching), 2281, 2204 (C≡N, stretching), 1186, 1332 (SO₂, stretching), 3088, 3066 (CH-Aromatic), 2866, 2831 (CH-aliphatic), 900 (S-N). Anal. Calcd. For C₂₃H₂₄N₄O₂S: C, 65.69; H, 5.75; N, 13.32; O, 7.61; S, 7.62.

9. 4-(2-amino-3-cyano-4-(4-methylphenyl)-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1i**. Yield 78.80%; mp 152-154 °C **IR** (KBr, cm⁻¹) : 3375, 3477, 3269 (NH₂, stretching), 2281, 2231 (C≡N, stretching), 3089, 3066 (CH-aromatic), 2868, 2833 (CH-aliphatic). **MS** (m/z): 420. **¹H NMR**: Solvent (CD₃OD) δ 3.321-3.300 [s, 2H, R-NH₂], δ 4.827 [m, 8H, AR-NH₂], δ 6.714-6.666 [m, 4H, AR-H], δ 7.611-7.564 [m, 4H, ARH]. **¹³C NMR**: δ153.62 (C in aromatic), δ131.32 (C=C in alkenes), δ129.06 (C=C in alkenes), δ 114.65 (C=C), δ 114.55 (C=C), δ 49.99 (C-C).

Anal. Calcd. For C₂₃H₂₄N₄O₂S: C, 65.69; H, 5.75; N, 13.32; O, 7.61; S, 7.62.

10. 4-(2-amino-3-cyano-4-(2,4-dimethylphenyl)-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1j**. Yield 78.04%; mp 152-154 °C **IR** (KBr, cm⁻¹) : 3383, 3477, 3323 (NH₂, stretching), 2235, 2279 (C≡N, stretching), 3055, 3068, 3041 (CH Aromatic), 2852, 2899, 2868 (CH aliphatic), 1184, 1334 (SO₂). **MS** (m/z): 434. **¹H NMR**: Solvent (CD₃OD) δ 3.321-3.300 [s, 2H, R-NH₂], δ 4.827 [m, 8H, AR-NH₂], δ 6.714-6.666 [m, 4H, AR-H], δ 7.611-7.564 [m, 4H, ARH]. **¹³C NMR**: δ153.60 (C in aromatic), δ131.33 (C=C in alkenes), δ129.06 (C=C in alkenes), δ 114.66 (C=C), δ 114.55 (C=C), δ 49.73 (C-C), δ 50.01 (C-O). Anal. Calcd. For C₂₄H₂₆N₄O₂S: C, 66.33; H, 6.03; N, 12.89; O, 7.36; S, 7.38.

11. 4-(2-amino-3-cyano-4-(2-fluorophenyl)-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1k**. Yield 72.08%; mp 222-224 °C **IR** (KBr, cm⁻¹) : 3477, 3375 (NH₂, stretching), 2222, 2281 (C≡N, stretching), 1188, 1313 (SO₂, stretching), 1095 (C-F, stretching), 900.76 (S-N, stretching). Anal. Calcd. For C₂₂H₂₁FN₄O₂S: C, 62.25; H, 4.99; F, 4.48; N, 13.20; O, 7.54; S, 7.55.

12. 4-(2-amino-3-cyano-4-(3-fluorophenyl)-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzene Sulfonamide **1L**. Yield 76.64%; mp 154-156 °C **IR** (KBr, cm⁻¹) : 3477, 3383 (NH₂, stretching), 2235 (C≡N, stretching), 1182.36 (SO₂, stretching), 1010.70 (C-F, stretching), 2831.50 (C-H, aliphatic). **MS** (m/z): 424.2. **¹H NMR**: Solvent (CD₃OD) δ 3.320-3.299 [s, 2H, R-NH₂], δ 4.909-4.843 [m, 8H, AR-NH₂], δ 6.709-6.661 [m, 4H, AR-H], δ 7.606-7.558 [m, 4H, AR-H]. **¹³C NMR**: δ153.62 (C in aromatic), δ131.34 (C=C in alkenes), δ129.04 (C=C in alkenes), δ 114.58 (C=C), δ 114.47 (C=C), δ 49.94 (C-C). Anal. Calcd. For C₂₂H₂₁FN₄O₂S: C, 62.25; H, 4.99; F, 4.48; N, 13.20; O, 7.54; S, 7.55.

13. 4-(2-amino-3-cyano-4-(4-fluorophenyl)-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzene

sulfonamide **1m**. Yield 78.66%; mp 160-162°C **IR** (KBr, cm⁻¹): 3375, 3477 (NH₂, stretching), 2279, 2206.57 (C≡N, stretching), 1186 (SO₂, stretching), 1002 (C-F, stretching), 898 (S-N, stretching), 3066, 3039 (CH, aromatic), 2833.43 (CH aliphatic). **MS** (m/z): 424. **¹H NMR**: Solvent (CD₃OD) δ 3.321-3.300 [s, 2H, R-NH₂], δ 4.827 [m, 8H, AR-NH₂], δ 6.714-6.666 [m, 4H, AR-H], δ 7.611-7.564 [m, 4H, ARH]. **¹³C NMR**: δ153.61(C in aromatic), δ131.32 (C=C, in alkenes), δ129.06(C=C in alkenes), δ 114.65 (C=C), δ 114.54 (C=C), δ 49.99 (C-C). Anal.Calcd.For C₂₂H₂₁FN₄O₂S: C, 62.25; H, 4.99; F, 4.48; N, 13.20; O, 7.54; S, 7.55.

14. 4-(2-amino-3-cyano-4-(3,4-difluorophenyl)-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1n**. Yield 68.26%; mp 148-150°C **IR** (KBr, cm⁻¹): 3477, 3375 (NH₂, stretching), 2233.57 (C≡N, stretching), 1184 (SO₂, stretching), 1072 (C-F, stretching), 898 (S-N, stretching). Anal.Calcd.For C₂₂H₂₀F₂N₄O₂S: C, 59.72; H, 4.56; F, 8.59, N, 12.66; O, 7.23; S, 7.25.

15. 4-(2-amino-3-cyano-4-(3-hydroxyphenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1o**. Yield 68.26%; mp 220-222°C **IR** (KBr, cm⁻¹): 3265, 3358 (NH₂, stretching), 2177, 2216, (C≡N, stretching), 1149, 1170, 1190, 1251, 1278, 1321, 1340, (SO₂, stretching), 3464 (OH), 3091 (Ar-CH), 2953(aliphatic- CH), 1587, 1612, 1649 (C= O). Anal.Calcd.For C₂₂H₂₀N₄O₄S : C, 60.54; H, 4.62; N, 12.84; O, 14.66; S, 7.35.

16. 4-(2-amino-3-cyano-4-(3-nitrophenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzene sulfonamide **1p**. Yield 70.82%; mp 212-214°C **IR** (KBr, cm⁻¹): 3224, 3342 (NH₂, stretching), 2181, 2216, 2283, (C≡N, stretching), 1165, 1192, 1255, 1267, 1348, (SO₂, stretching), 3091 (Ar-CH), 2960, 2912 (aliphatic-CH), 1587, 1620, 1649 (C= O). Anal.Calcd.For C₂₂H₁₉N₅O₅S: C, 56.77; H, 4.11; N, 15.05; O, 17.19; S, 6.89.

17. 4-(2-amino-3-cyano-4-(4-nitrophenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzene sulfonamide **1q**. Yield 76.35%; mp 284-286°C **IR** (KBr, cm⁻¹): 3223, 3346 (NH₂, stretching), 2193 (C≡N, stretching), 1161, 1184, 1255, 1288, 1348, 1323, (SO₂, stretching), 3078, 3103 (Ar-CH), 2953, 2910, 2985, (aliphatic- CH), 1517, 1608, (C= O). Anal.Calcd.For C₂₂H₁₉N₅O₅S :C, 56.77; H, 4.11; N, 15.05; O, 17.19; S, 6.89.

18. 4-(2-amino-3-cyano-4-(4-chlororophenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1r**. Yield 68.58%; mp 280-282°C **IR** (KBr, cm⁻¹): 3356, 3257 (NH₂, stretching), 2181, (C≡N, stretching), 1141, 1184, 1247, 1317 (SO₂, stretching), 3035, 3043, 3062 (Ar-CH), 2949 (aliphatic- CH), 1612 (C= O). Anal.Calcd.For C₂₂H₁₉ClN₄O₃S: C, 58.08; H, 4.21; Cl, 7.79; N, 12.32; O, 10.55; S, 7.05.

19. 4-(2-amino-3-cyano-4-(2,4-dichlorophenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)-yl) benzenesulfonamide **1s**. Yield 67.87%; mp 290-292°C **IR** (KBr, cm⁻¹): 3219, 3296, 3346 (NH₂, stretching), 2183, (C≡N, stretching), 1163, 1190, 1265, 1315, 1338 (SO₂, stretching), 3068, 3089 (Ar-CH), 2960, 2953 (aliphatic-CH), 1587, 1612 (C= O). Anal.Calcd.For C₂₂H₁₈Cl₂N₄O₃S: C, 53.99; H, 3.71; Cl, 14.49; N, 11.45; O, 9.81; S, 6.55.

20. 4-(2-amino-3-cyano-4-(3, 4-dimethoxyphenyl)-5-oxo-5,6,7,8 tetrahydro-quinolin-1(4H)yl) benzenesulfonamide **1t**. Yield 72.67%; mp 148-150°C **IR** (KBr, cm⁻¹): 3240, 3257, 3350 (NH₂, stretching), 2181, 2125, 2223 (C≡N, stretching), 1143, 1157, 1184, 1205, 1253, 1274 (SO₂, stretching), 3068, 3028, 3089 (Ar-CH), 2964, 2937 (aliphatic- CH), 1566 (C= O). Anal.Calcd. For C₂₄H₂₄N₄O₅S: C, 59.99; H, 5.03; N, 11.66; O, 16.65; S, 6.67.

21. 4-(2-amino-3-cyano-4-(2,3-dimethoxyphenyl)-5-oxo-5,6,7,8-tetrahydroquinolin-1(4H)yl) benzene sulfonamide **1u**. Yield 65.84%; mp 174-176°C **IR** (KBr, cm⁻¹): 3228, 3250, 3350

(NH₂, stretching), 2173,2194,2218,2254 (C≡N, stretching), 1165,1186,1226,1269,1317,1338 (SO₂, stretching), 3068, 3003 (Ar-CH), 2906,2943 (aliphatic- CH),1556,1585,1614 (C=O). Anal.Calcd.For C₂₄H₂₄N₄O₅S: C, 59.99; H, 5.03; N, 11.66; O, 16.65; S, 6.67.

CONCLUSION

Twenty one novel benzenesulfonamide derivatives have been synthesized and some of the novel synthesized compounds are screened for antibacterial and antifungal activity. These compounds showed good to moderate activity against E. coli, B. Subtilis, S Typhi, S Aurus. Aspergillus Flavus, Aspergillus Niger and Candida albicans. The biological activity of these compounds will trigger more interest in the synthesis of such compounds from the easily available starting materials.

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