

# Synthesis and Biological Evaluation of 2-Aryl - 6, 7-Difluorophenyl-1,3,4-Oxadiazolo (3,2-a) (1,3,5)-Triazine-5-(6H,7H)-Thiones

Kiran Mishra\*

Department of Applied Sciences, Chandigarh Engineering College, CGC, Landran, Mohali, Punjab, India

## Abstract

2-Aryl-6,7-difluorophenyl-1,3,4-oxadiazolo (3,2-a) (1,3,5)- triazine-5-(6H,7H)- thiones (2a-j) have been orchestrated by the cyclo expansion of para-fluorophenyl 4-fluoro phenyl isothiocyanate and 2-(4-fluorobenzylidene) to (1a-j) 2-amino - 5-phenyl-1,3,4-oxadiazole in dry toluene all the integrated compounds were very much described by their elemental analysis and spectral data. The integrated compounds have been screened for their antifungal action against *Phytophthora infestans* and *Colletotrichum falcatum*. Results demonstrated that greatest antifungal activity was appeared by the compounds 2b, 2c, 2g and 2h. These compounds demonstrated 99%, 98%, 99% and 97% hindrance of *Phytophthora infestans* and 97%, 96%, 98% and 97% *Colletotrichum falcatum* at 1000 ppm respectively.

**Keywords:** Fungitoxicity; Oxadiazole; Triazine; Thiones

## Introduction

A few 1,3,4-oxadiazole derivatives are known not different kind of valuable natural exercises including herbicidal [1], fungicidal [2,3], bactericidal [4], insecticidal [5], etc. Perhaps toxophoric significance of which have been all around pushed in numerous pesticides [6,7]. Numerous sorts of 1,3,5-triazine derivatives have significance in horticulture as herbicides and fungicides of these, Simanize (1) [2-chloro-4,6-bis (ethyl amino) - 1,3,5-triazine], Atrazine [2-chloro - 4-ethylamino-6-isopropylamino-1,3,5-triazine], Prometryne [2-methylthio-4,6-bis (isopropyl amino)- 1,3,5-triazine], Ametryne [2-Methylthio-4-ethylamino-6-(isopropylamino)- 1,3,5-triazine], Dyrene [2,4-dichloro-6-(2-chloroanilino)- 1,3,5-triazine] and Methoprotryne [2-Methylthio-4-isopropylamino-6(3-methoxypropylamino)- 1,3,5-triazine] are all the more remarkable. This significance of 1,3,4-oxadiazole derivatives and 1,3,5-triazine derivatives have incited us to synthesize. Some novel title compounds (2a-j). Thus, antifungal action of synthesize compounds have been screened against growths *Phytophthora infestans* and *Colletotrichum falcatum*.

## Experimental

Melting points were taken in open capillary tubes and are uncorrected. The IR spectra were recorded in KBr on Perkin-Elmer-720 spectrophotometer. The <sup>1</sup>H NMR spectra were recorded in CDCl<sub>3</sub> on Varian A-60D spectrophotometer. The chemical shifts are recorded in ppm downfield from TMS, which are utilized as an inner standard (Figure 1).

## Synthesis of compounds (2a-j)

**2-(4-fluorobenzylidene amino)-5-phenyl-1, 3, 4-oxadiazole (1a-j):** A mixture of 2-amino-5-phenyl-1,3,4-oxadiazole 0.02 mol and 4-fluorobenzaldehyde 0.02mol in total ethanol was refluxed for 4 hrs. and separated while hot. The filtrate after cooling outfitted the wanted item which was recrystallized from ethanol as yellowish needles. All the prepared compounds very much concurred with their antifungal information.

**7-(4-Fluorophenyl-6,7-dihydro-2-phenyl-6-fluorophenyl)-1,3,4-oxadiazolo(3,2-a)-s-triazine-5-thione (2aj):** A mixture of 2-(4-fluorobenzylidene) amino-5-phenyl-1,3,4-oxadiazole 0.01mol and 4-fluorophenyl isothiocyanate 0.01mol) was refluxed in dry toluene for 6 hrs. What's more, the dissolvable was refined of under decreased

weight. The residue thus acquired was washed with little measures of ethanol took after by water and the product was recrystallized from ethanol was sparkling yellowish needles. Yield, melting point, molecular formula and elemental analysis of this and in addition that of alternate compounds of this class are recorded in Table 1.

## Antifungal activity

The compounds (2a-j) were screened for their antifungal movement against *Phytophthora infestans* and *Colletotrichum falcatum* by known technique at the three concentrations Viz., 1000, 100, 10 ppm. The screening information of compounds is recorded in Table 2. Results were contrasted with commercial fungicide DithaneM-45 tested under similar conditions. The percentage inhibition has been calculated by the formula:

$$\% \text{ of inhibition} = (C - T) \times 100 / C$$

Where C and T are diameter (in mm) of fungus colony in control and treated plates respectively.

## Results and Discussion

The 2-Aryl-6,7-difluorophenyl-1,3,4-oxadiazolo (3,2-a) (1,3,5)- triazine-5-(6H,7H)- thiones compounds were screened against *Phytophthora infestans* and *Colletotrichum falcatum* for antifungal activity and their screening data have been summarized in Table 2. Perusal of the screening results demonstrates that all the tested compounds (2a-j) hindered more than 65% growth of the both the test parasites at 1000 ppm concentrations of these, the most dynamic compounds 2b and 2g showed the fungicidal action almost equivalent to that of Dithane M-45 at 1000 ppm concentrations, and restrained 40-44% development of the contagious species even at 10 ppm concentrations. The screened compounds 2b and 2g were highly

\*Corresponding author: Kiran Mishra, Department of Applied Sciences, Chandigarh Engineering College, CGC, Landran, Mohali-140 307, Punjab, India, Tel: 01723984200; E-mail: [drmishrakiran@gmail.com](mailto:drmishrakiran@gmail.com)

Received April 26, 2017; Accepted May 02, 2017; Published June 09, 2017

Citation: Mishra K (2017) Synthesis and Biological Evaluation of 2-Aryl - 6, 7-Difluorophenyl-1,3,4-Oxadiazolo (3,2-a) (1,3,5)-Triazine-5-(6H,7H)-Thiones. Chem Sci J 8: 153. doi: [10.4172/2150-3494.1000153](https://doi.org/10.4172/2150-3494.1000153)

Copyright: © 2017 Mishra K. This is an open-access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.

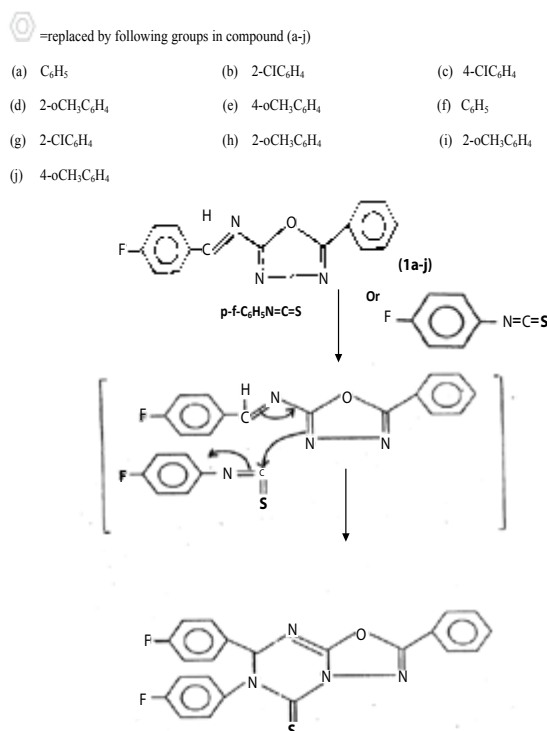


Figure 1: The chemical shifts are recorded in ppm downfield from TMS.

Compd. No.	Ar	Yield %	M.P. (°C)	Molecular Formula	Found (Calcd.)%		
					C	N	S
2a	C <sub>6</sub> H <sub>5</sub>	76	236	C <sub>12</sub> H <sub>14</sub> F <sub>2</sub> N <sub>4</sub> OS	62.8(62.83)	13.3(13.34)	07.6(07.60)
2b	2-ClC <sub>6</sub> H <sub>4</sub>	80	250	C <sub>22</sub> H <sub>13</sub> ClF <sub>2</sub> N <sub>4</sub> OS	58.1(58.11)	13.3(13.34)	07.0(07.60)
2c	4-ClC <sub>6</sub> H <sub>4</sub>	70	240	C <sub>22</sub> H <sub>13</sub> ClF <sub>2</sub> N <sub>4</sub> O <sub>2</sub> S	58.1(58.16)	13.3(13.34)	07.0(07.03)
2d	2-oCH <sub>3</sub> C <sub>6</sub> H <sub>4</sub>	75	185	C <sub>23</sub> H <sub>16</sub> F <sub>2</sub> N <sub>4</sub> O <sub>2</sub> S	61.3(61.30)	13.3(13.34)	07.1(07.14)
2e	4-oCH <sub>3</sub> C <sub>6</sub> H <sub>4</sub>	78	242	C <sub>23</sub> H <sub>16</sub> F <sub>2</sub> N <sub>4</sub> O <sub>2</sub> S	61.3(61.35)	13.3(13.34)	07.1(07.10)
2f	C <sub>6</sub> H <sub>5</sub>	74	235	C <sub>22</sub> H <sub>14</sub> F <sub>2</sub> N <sub>4</sub> OS	62.8(62.86)	13.3(13.31)	07.6(07.63)
2g	2-ClC <sub>6</sub> H <sub>4</sub>	78	248	C <sub>22</sub> H <sub>13</sub> ClF <sub>2</sub> N <sub>4</sub> OS	58.1(58.15)	12.3(12.35)	07.0(07.03)
2h	4-ClC <sub>6</sub> H <sub>4</sub>	73	240	C <sub>22</sub> H <sub>13</sub> ClF <sub>2</sub> N <sub>4</sub> OS	58.1(58.17)	12.3(12.32)	07.0(07.07)
2i	2-oCH <sub>3</sub> C <sub>6</sub> H <sub>4</sub>	76	189	C <sub>23</sub> H <sub>16</sub> F <sub>2</sub> N <sub>4</sub> O <sub>2</sub> S	61.3(61.31)	13.3(12.45)	07.1(07.12)
2j	4-oCH <sub>3</sub> C <sub>6</sub> H <sub>4</sub>	72	240	C <sub>23</sub> H <sub>16</sub> F <sub>2</sub> N <sub>4</sub> O <sub>2</sub> S	61.3(61.37)	12.4(12.42)	07.1(07.09)

Table 1: The Yield, melting point, molecular formula and elemental analysis of 7-(4-Fluorophenyl)-6,7-dihydro-2-phenyl-6-fluorophenyl)-1,3,4-oxadiazolo(3,2-a)-s-triazine-5-thione (2a-j) and in addition, alternate compounds of this class are shown below.

Comp. No.	Ar	Average % inhibition after 96 hours					
		Phytophthora infestans			Colletotrichum falcatum		
		1000 ppm	100 ppm	10 ppm	1000 ppm	100 ppm	10 ppm
2a	C <sub>6</sub> H <sub>5</sub>	66	37	20	65	35	19
2b	2-ClC <sub>6</sub> H <sub>4</sub>	99	58	40	97	53	40
2c	4-ClC <sub>6</sub> H <sub>4</sub>	98	56	38	96	52	40
2d	2-oCH <sub>3</sub> C <sub>6</sub> H <sub>4</sub>	72	43	30	74	45	33
2e	4-CH <sub>3</sub> C <sub>6</sub> H <sub>4</sub>	67	39	23	68	41	24
2f	C <sub>6</sub> H <sub>5</sub>	71	42	30	69	39	26
2g	2-ClC <sub>6</sub> H <sub>4</sub>	99	65	44	98	63	43
2h	4-ClC <sub>6</sub> H <sub>4</sub>	97	60	41	97	60	39
2i	2-oCH <sub>3</sub> C <sub>6</sub> H <sub>4</sub>	71	42	29	72	43	31
2j	4-oCH <sub>3</sub> C <sub>6</sub> H <sub>4</sub>	69	40	24	70	42	26
Dithane M-45		100	80	66	100	85	68

Table 2: Antifungal activity of compounds (2a-j) and Dithane M-45.

harmful to *Phytophthora infestans* and *Colletotrichum falcatum* at higher concentrations (1000 ppm) the general results are not all that empowering as one would anticipate from the combined execution of the two boilable nuclei 1,3,4-oxadiazole and s-triazine this may be credited to the halfway immersion in the s-triazine nucleus resulting about the loss of planarity of the oxadiazolo -s-triazine ring system. This assumption is bolstered by the before perception that minimal size and planarity of molecule frequently improve its pesticidal activity. It is in any case, critical that the presentation of chloromethoxy and methyl groups in aryl moiety of these compounds tend to contention the growths fungi toxicity and that presentation of methoxy or methyl group at ortho position is more viable than that para-position. Likewise, the presentation of chloro group. Fungicidal activity changed imperceptibly with the fungal species.

#### Acknowledgements

The authors are very thankful to the Director, CDRI, Lucknow for providing basic, unearthy information and to the Director, IARI, New Delhi for lab facilities to evaluate antifungal activity.

#### References

1. Maienfisch P, Hall RG (2004) The Importance of Fluorine in the Life Science Industry. CHIMIA International Journal for Chemistry 58: 93-99.
2. Weinke KE, Lauber JJ, Greenwald BW, Preiser FA (1969) Thiabendazole, a new systemic fungicide. Proceedings of the Fifth British Insecticide and Fungicide Conference, Brighton, London, UK.
3. Gibson MS (1962) A bond energy scheme for aliphatic and benzenoid compounds. Tetrahedron 18: 1377.
4. Summers LA (1976) One- electron transfer properties and herbicidal activity of diquaternary salts of 2,4-di-(4-pyridyl)-1,3,5-triazines. Tetrahedron 32: 615.
5. Kumar V, Averill BA (1988) Facile synthesis of N-10 flavin disulfides. J Hetrocycl Chem 25: 253.
6. Jumat S, Nadia S, Ayad H, Hiba IEY (2010) Synthesis and antibacterial activity of some new 1,3,4-oxadiazole and 1,3,4-thiadiazole derivatives. J Appl Sci Res 6: 866-870.
7. Selvakumar K, Anandarajgopal K, Rajamanickam V, Ajaykumar TM, Jesindha B (2011) 5-(2-Aminophenyl)-1,3,4-Oxadiazole-2(3h)-Thionederivatives: Synthesis, Characterization and Antimicrobial Evaluation. Intl J of Pharm Sci Rev and Res 6: 64-67.