

Synthesis of Substituted Benzimidazole Derivatives using Alanine as Catalyst

Gurumeet Wadhawa¹, Smita Tandale¹, Maryappa Sonawale¹, Yashwant Gaikwad¹,
Sohani Hake², Pranali Satpute², Kajal Kharje²

Department of Chemistry, Veer Wajekar ASC College, Phunde¹
Students, Department of Chemistry Veer Wajekar ASC College, Phunde²

Abstract: Both benzimidazole and its derivatives have a wide range of biological actions. They possess a range of fascinating biological traits, such as anticancer, antidiabetic, antiviral, and antibacterial. In the presence of alanine and sonication, orthophenylene diamine was coupled with an aromatic aldehyde to create novel benzimidazole derivatives. The solution made use of a novice course that was also financially viable.

Keywords: O-Phenylenediamine, Catalyst, Green Route.

I. INTRODUCTION

Benzimidazole and its equivalents demonstrate a variety of biological actions in the pharmaceutical industry. The benzimidazole group is present in a wide variety of bioactive compounds, including those having anticancer, antiviral, and antihypertensive properties. They demonstrate antiviral action against the herpes (HSV-1) virus, HIV, and influenza viruses [1-6]. Moreover, they are employed in clinical medicine as an antiviral, antitumor, and anti-ulcer drug. Treatment options for them include the management of ischemia repression, traumatic hypertension, and obesity [7-10]. This benzimidazole make it feasible to develop highly desired, widely used organic compound chemical processing techniques that are simple, successful, environmentally safe, and commercially viable. It is possible to make 2-substituted benzimidazole in a variety of ways. The majority of the time, O-phenyldiamines is condensed with a range of carbonyl substances, such as carboxylic acid, nitriles, imides, orthoesters, orthoformate, and orthoacetate [11-19]. Reacting it with various aromatic aldehydes and o-Phenylenediamine while employing cobalt (II), magnesium chloride, chloride hexahydrate, ferric chloride, and ferrous sulphate as catalysts is a quick and practical technique to create 2-arylbenzimidazole. In the presence of strong oxidising agents such I₂/KI/K₂CO₃, N-halosuccinimide (Cl, Br, I), Yb (OTf)₃, PEG-100, and (NH₄)₂H₂PW₁₂O₄ as well as mineral acids like hydrochloric acid and polyphosphoric acid, O-Phenylenediamine condensation with carbonyl compounds occurs [18-22]. Without considerably boosting activity, the method works with substituted o-Phenylenediamine, aromatic, unsaturated, and aliphatic aldehydes. It converts Phenylenediamine and aldehydes into variously substituted benzimidazole in good yields without the need of solvents. A few methods made no claims that they were ubiquitous or flexible enough to work with various beginning materials. These techniques consume more organic solvent, produce less, are not recyclable, have lower purity, take longer, and cause more pollution.

II. EXPERIMENTAL

Using the basic paraffin Bath method, the melting point is determined. ¹H NMR spectra (CDCl₃) were acquired using a Bruker Advance I 300 NMR spectrophotometer, with TMS serving as the internal standard. Nujol mull IR spectra in the frequency region of 4000-450 cm⁻¹ were recorded using a Shimadzu FTIR spectrophotometer. Use of S.d.fine chemicals. On silica gel-G plates, the purity of the compound was analysed using thin-layer chromatography. [28-30]

Experimental procedure for synthesis of 2 substituted benzimidazole derivative

A mixture of 10 mmol of substituted aromatic aldehyde, 10 mmol of orthophenyldiamine, and 5 mmol of alanine was treated to a chemical reaction at 84 0 °C for the specified period in a sonicator. Using thin-layer chromatography, the reaction is observed. After the reaction is complete, water is added, and then the product is extracted with organic

solvent and purified using silica. The product was roughly 90% pure when it was obtained. There are no adverse effects. The catalyst is recycled five to six times.

Observation Table

Sr. No	Aromatic Group	Time (Min)	(%) Yield	MP (0C)
1	2-phenyl-1H-benzimidazole	85	75	290
2	2-(2-chlorophenyl)-1H-benzimidazole	75	80	134
3	2-(3-chlorophenyl)-1H-benzimidazole	75	96	229
4	2-(4-methoxyphenyl)-1H-benzimidazole	95	86	221
5	2-(2-nitrophenyl)-1H-benzimidazole	70	95	254
6	2-(3-nitrophenyl)-1H-benzimidazole	70	92	301
7	2-(4-nitrophenyl)-1H-benzimidazole	70	96	303
8	2-(2-methoxyphenyl)-1H-benzimidazole	95	60	267
9	2-(tetrahydrofuran-2-yl)-1H-benzimidazole	80	80	285

III. DISCUSSION

We have created a superior method for the synthesis of benzimidazole derivatives in terms of time, purity, yield, and cost. The catalyst employed in this process is recyclable. Low-priced and easily accessible catalyst.

ACKNOWLEDGMENT

All Authors are Thankful to the Principal of College Dr.P.G.Pawar and the RayatShikshan Sanstha For Providing the Support for Research work.

REFERENCES

- [1]. Middleton, R.W.; Wimberley D. G. J. Het. Chem. 1980, 17, 1757.
- [2]. Kumar R, Joshi YC E-journal of chemistry, 2007, 4(4):606-610.
- [3]. Sukalovic V., Andric D., Roglic G., Sladjana K.R., Schratteholz A. and Soskis V., Eur. J. Med. Chem., 2005, 40, 481.
- [4]. Hashem Sharghietal Journal of Heterocyclic Chemistry Volume 45, Issue 5 September/October 2008 Pages 1293–1298
- [5]. J S Yadav, etal Canadian Journal of Chemistry, 2008, 86, 124-128.
- [6]. Elder field, R. C.; Kreysa, F. J. J. Am. Chem. Soc. 1948, 70, 44.
- [7]. B. S. Praveen, Reddy, P.; Sondhi, S. M.; Lown, J. W. Pharmacol. Ther. 1999, 84,(1) 1-111.
- [8]. Sluka, J.; Novak, J.; Budesinsky, Z. Collect. Czech. Chem. Commun. 1976, 41, 3628.
- [9]. Nitin A. Mirgane, Vitthal S. Shivankar, Sandip B. Kotwal, Gurumeet C. Wadhawa, Maryappa C. Sonawale, Degradation of dyes using biologically synthesized zinc oxide nanoparticles, Materials Today: Proceedings, Volume 37, Part 2021, 849-853, ISSN 2214-7853, <https://doi.org/10.1016/j.matpr.2020.06.037>.

- [10]. Nitin A. Mirgane, Vitthal S. Shivankar, Sandip B. Kotwal, Gurumeet C. Wadhawa, Maryappa C. Sonawale, the Waste pericarp of ananas comosus in green synthesis zinc oxide nanoparticles and their application in wastewater treatment, *Materials Today: Proceedings*, Volume 37, Part 2, 2021, 886-889, ISSN 2214-7853, <https://doi.org/10.1016/j.matpr.2020.06.045>.
- [11]. Shubhada S. Nayak, Nitin A. Mirgane, Vitthal S. Shivankar, Kisan B. Pathade, Gurumeet C. Wadhawa, Adsorption of methylene blue dye over activated charcoal from the fruit peel of plant *Hydnocarpus pentandra*, *Materials Today: Proceedings*, Volume 37, Part 2, 2021, 2302-2305, ISSN 2214-7853, <https://doi.org/10.1016/j.matpr.2020.07.728>.
- [12]. Patil, D.D.; Mhaske, K.D.; Wadhawa, C.G., Antibacterial and Antioxidant study of *Ocimum basilicum* Labiatae (sweet basil), *Journal of Advanced Pharmacy Education & Research* (2011) 2, 104-112.
- [13]. Dinanath PD, Gurumeet WC, 2013. Antibacterial, antioxidant and anti-inflammatory studies of leaves and roots of *Solanum xanthocarpum*. *Unique J Ayurvedic Herb Med* (2013) ;(3):59-63.
- [14]. Dynashwar K. Mhaske, Dinanth D. Patil, Gurumeet C. Wadhawa. Antimicrobial activity of methanolic extract from rhizome and roots of *Valerianawallichii*. *International Journal on Pharmaceutical and Biomedical Research*, 2011; 2(4):107- 111
- [15]. Patil DD, Mhaske DK, Gurumeet MP, Wadhawa C. Antibacterial and antioxidant, anti-inflammatory study of leaves and bark of *Cassia fistula*. *Int J Pharm* 2012; 2(1):401-405.
- [16]. G. C. Wadhawa, M. A. Patare, D. D. Patil and D. K. Mhaske, Antibacterial, antioxidant and anti-inflammatory studies of leaves and roots of *Anthocephalus kadamba*. *Universal Journal of Pharmacy*, 2013.
- [17]. Shubhada S. Nayak, Nitin A. Mirgane, Vitthal S. Shivankar, Kisan B. Pathade, Gurumeet C. Wadhawa, Degradation of the industrial dye using the nanoparticles synthesized from flowers of plant *Ceropegia attenuata*, *Materials Today: Proceedings*, Volume 37, Part 2, 2021, Pages 2427-2431, ISSN 2214-7853, <https://doi.org/10.1016/j.matpr.2020.08.274>.
- [18]. G. C. Wadhawa, V. S. S. Hivankar, Y. A. Gaikwad, B. L. Ingale, B. R. Sharma, S. S. Hande, C. H. Gill and L. V. Gavali, *Eur. J. Pharm. Med. Res.*, 3, 556 (2016).
- [19]. Thaipongl K, Boonprakob1 U, Cisneros-Zevallos L and Byrne DH. Hydrophilic and lipophilic antioxidant activities of guava fruits. *Southeast Asian J trop med public health* 2005; 36: 254-257. [PMID:16438219]
- [20]. Jia Z., Tang M., Wu J., the determination of flavonoid contents in mulberry and their scavenging effects on superoxide radicals *Food Chem*; 1999;64:555-599.
- [21]. Roberta Re, Pellegrini N, Proteggente A, Pannala A, Yang M, Rice-Evans C. Antioxidant activity applying an improved ABTS radical cation decolorization assay *FreeRadical Biol Med*; 1999;26:1231-1237.
- [22]. Békro YA, Békro JAM, Boua BB, Tra BFH, Ehilé EE, Phytochemical Screening, Antioxidant and Antibacterial activity of *Lepidium sativum* seeds from Morocco. *Rev. Sci. Nat.* 2007; 4: 217-225.
- [23]. Huang B., Ban X.Q., He J.S., Tong J., Tian J., Wang Y.W., Hepatoprotective and antioxidant activity of ethanolic extracts of edible lotus (*Nelumbo nucifera Gaertn.*) leaves *Food Chem.* 2010;120:873-878.
- [24]. Oyaizu M. Jpn Assessment of the Quality and Shelf-Life in Enriched n3 PUFA Raw Beef Patties Using Dry Soybean Sprouts as Antioxidant *J Nutr*1986; 103:413- 419.
- [25]. Fadili K., Amalich S., N'dedianhoua S K., Bouachrine M., Mahjoubi M., El Hilali F., Zair T. Phytochemical Screening and Antioxidant Activity of Moroccan *Thymus satureioides* Extracts *Inter J Innov Sci Res*, 2015;17:24-33.
- [26]. Harborne JB., In: *Phytochemical Methods* (Chapman and Hall, London); 1973.
- [27]. Schulz H, Baranska M. Identification, and quantification of valuable plant substances by IR and Raman spectroscopy. *Vibrational Spectroscopy.* 2007; 43: 13-25. [<https://doi.org/10.1016/j.vibspec.2006.06.001>]
- [28]. Binet ML, Commereuc S, Chalchat JC, Lacoste J. Oxidation of polyterpenes: a comparison of poly α , and poly β , pinenes behaviours: Part I photo-oxidation. *Journal of Photochemistry and Photobiology A: Chemistry.* 1999; 125: 45-53. [[https://doi.org/10.1016/S1010-6030\(98\)00412-2](https://doi.org/10.1016/S1010-6030(98)00412-2)]
- [29]. Graßmann J. Terpenoids as Plant Antioxidants. *Vitamins & Hormones.* 2005; 75: 505-535. [[https://doi.org/10.1016/S0083-6729\(05\)72015-X](https://doi.org/10.1016/S0083-6729(05)72015-X)]

- [30]. Patil, Dinanath D., Gurumeet C. Wadhava, and Arun K. Deshmukh. "One Pot Synthesis of Nitriles from Aldehydes and Hydroxylamine Hydrochloride Using Ferrous Sulphate in DMF Under Reflux Condition." *Asian Journal of Chemistry* 24.3 (2012): 1401.
- [31]. Patil DD, Mhaske DK, Wadhawa GC. Green Synthesis of 3,4-dihydropyrimidinone using ferrous sulphate as recyclable catalyst, *Journal of Pharmaceutical Research and Opinion*, 2011; 1(6): 172–174.
- [32]. Nayak, Shubhada S., et al. "Green synthesis of the plant assisted nanoparticles from *Euphorbia neriifolia* L. and its application in the degradation of dyes from industrial waste." *Plant Science Today* 8.2 (2021): 380-385.
- [33]. S. S. Nayak, G. C. Wadhawa, V. S. Shivankar, R. Inamadar, and M. C. Sonawale, "Phytochemical Analysis and Dpph Antioxidant Activity of Root and Bark of *Syzygium Stocksii* (Duthie) Plant," *Eur. J. Mol. Clin. Med.*, 7(10), 2021, 2655–2668
- [34]. Shubhada S. Nayak, Gurumeet C. Wadhawa, Vitthal S. Shivankar, Dinanath D. Patil, Maryappa C. Sonawale, Nitin A. Mirgane, Tin oxide plant assisted nanoparticle catalyzed green synthesis of imidazole derivatives, *Materials Today: Proceedings*, Volume 37, Part 2, 2021, Pages 2490-2494, ISSN 2214-7853, <https://doi.org/10.1016/j.matpr.2020.08.301>.