

## DIFFERENT BIOLOGICAL ACTIVITIES OF SUBSTITUTED BENZIMIDAZOLE DERIVATIVES-A REVIEW ARTICLE

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Article Received on 12/01/2018

Article Revised on 04/02/2018

Article Accepted on 25/02/2018

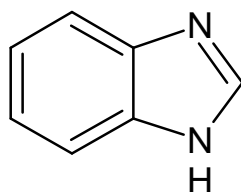
### ABSTRACT

Benzimidazole and its derivatives are considered as an important heterocyclic compound that exhibits a wide range of pharmaceutical applications including anti-cancers, anti-microbial, anti-hypertensive, anti-viral, anti-fungal, anti-HIVs, anti-convulsant, and anti-diabetics. In spite of their wide ranging activities, the synthesis of Benzimidazoles and its derivatives. The present review article focuses on substituted benzimidazole derivatives with potential activities.

**KEYWORDS:** Heterocyclic compound, benzimidazole, biological activity.

### INTRODUCTION

Benzimidazole is a important heterocyclic aromatic organic compound. Benzimidazole moieties are regarded as an important compound due to their wide range of applications in pharmaceutical chemistry. This compound is bicyclic in nature which consists of the combination of benzene and imidazole nucleus. The most well-known benzimidazole compound in nature is N-ribosyl-dimethylbenzimidazole, which treat as axial ligand for cobalt in vitamin B12.<sup>[1]</sup>



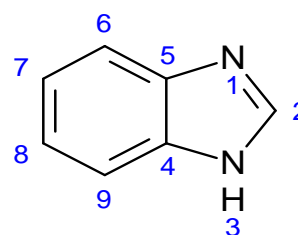
**Benzimidazole**

**Fig. 1.**

The benzimidazoles are also wellknown as benziminazoles or benzoglyoxalines. They have been also familiar also as derivatives of o-phenylenediamine, especially in the early literature. Thus, benzimidazole following to this nomenclature would be called methenyl-o-phenylenediamine and 2-methylbenzimidazole would be also called ethenyl-o-phenylenediamine. And they have been named as

derivatives of the grouping consisting the imidazole portion of the ring.<sup>[2]</sup>

The numbering system in benzimidazoles is as follows;

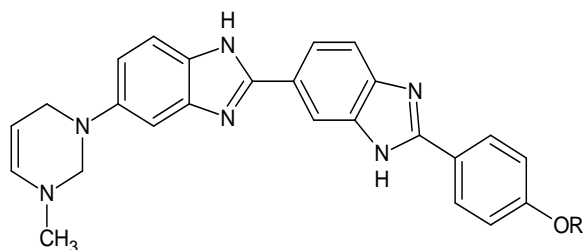


**Fig.-2.**

### LITERATURE REVIEW

#### Anticancer Activity

- Acar Çevik U, Sağlık BN, Korkut B, ozkay Y, Ilgın S et al.,(2018) synthesized 4-(5-chloro-1H-benzimidazol-2-yl)-benzoic acid benzylidene hydrazide derivatives and evaluated their anticancer activity against A549 (human lung adenocarcinoma) and MCF-7 (human breast adenocarcinoma) cells.<sup>[3]</sup>



R=ET, H

Fig-3.

- Wang Z, Deng X, Xiong S, Xiong R, Liu J, Zou L, Lei X, Cao X, Xie Z, Chen Y, Liu Y et al.,(2017) series of chrysin benzimidazole derivatives were prepared and evaluated for their anticancer activity.<sup>[4]</sup>

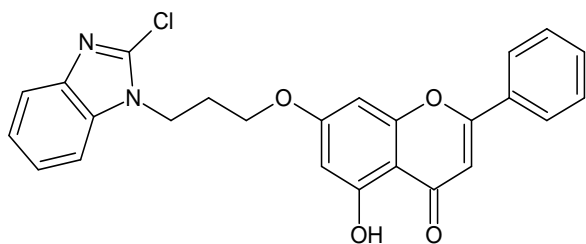
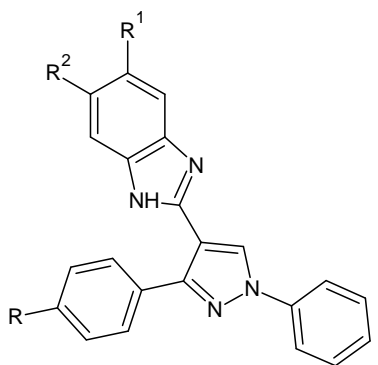


Fig.-4.

- Reddy TS, Kulhari H, Reddy VG, Bansal V, Kamal A, Shukla R et al.,(2015) series of different pyrazole containing benzimidazole derivatives have been designed, synthesized and screened for their potential anti-proliferative activity against three human tumor cell lines - lung (A549), breast (MCF-7), and cervical (HeLa).<sup>[5]</sup>



R=methyl, chloro, bromo, fluoro.

Fig.-5.

- Refaat HM et al.,(2010) synthesized 2-substituted benzimidazole derivatives and screened for anticancer activity.<sup>[6]</sup>

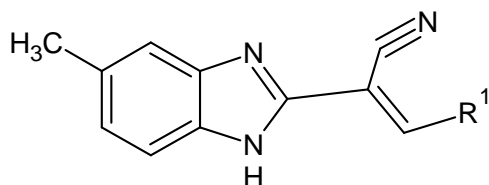
R=CH<sub>3</sub>, Cl, Br.

Fig.-6.

- Demirayak S, Kayagil I, Yurttas L et al.,(2011) synthesized 1,3-diarylpyrazino[1,2-a]benzimidazole derivatives and investigated of their anticancer activities.<sup>[7]</sup>

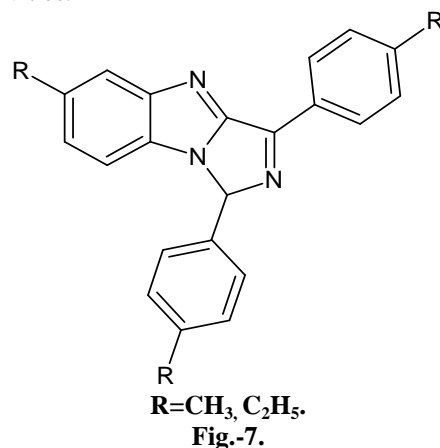
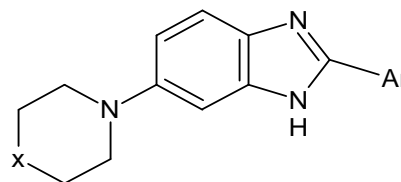
R=CH<sub>3</sub>, C<sub>2</sub>H<sub>5</sub>.

Fig.-7.

#### Antioxidant Activity

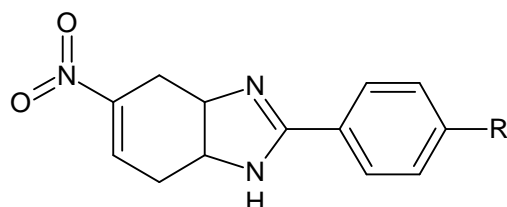
- Özil M, Parlak C, Baltaş N et al., (2018) 2-(aryl)-6-morpholin-4-yl(or 4-methylpiperazin-1-yl)-1H-benzimidazole derivatives were designed and synthesized and evaluated for *in vitro* antioxidant activities.<sup>[8]</sup>



X=Nmethyl, O.

Fig.-8.

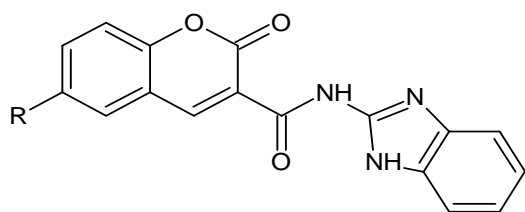
- Archie SR, Das BK, Hossain MS, Kumar UT et al.,(2017) synthesised 2-substituted-5-nitro benzimidazole derivatives and evaluated their antioxidant activity.<sup>[9]</sup>



R=Methyl, ethyl.

Fig.-9.

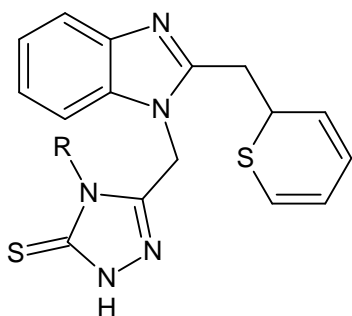
- Arora RK, Kaur N, Bansal Y, Bansal G et al.,(2014) series of 2-substituted benzimidazoles,derivatives were synthesised and evaluated for its antioxidant activity.<sup>[10]</sup>



R=methyl, chloro.

Fig.-10.

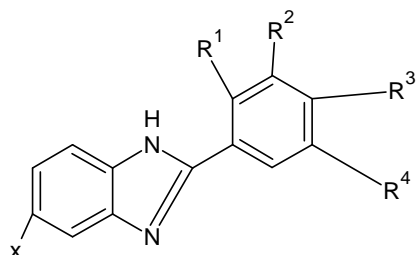
- Menteşe E, Yılmaz F, Baltaş N, Bekircan O, Kahveci B *et al.*,(2015) triheterocyclic compounds containing benzimidazole, were synthesized and screened for their antioxidant activities.<sup>[11]</sup>



R=methyl, ethyl, propyl.

Fig.-11.

- Zhou B, Li B, Yi W, Bu X, Ma L *et al.*,(2013). 2-arylbenzimidazole derivatives were synthesized and evaluated for their antioxidant activity.<sup>[12]</sup>



X, R1, R2, R3, R4= H, OH, OMe.

Fig.-12.

#### Antimicrobial Activity

- Liu HB, Gao WW, Tangadanchu VK, Zhou CH, Geng RX *et al.*,(2018) series of aminopyrimidinyl benzimidazoles were synthesized and evaluated for its antimicrobial activity.<sup>[13]</sup>

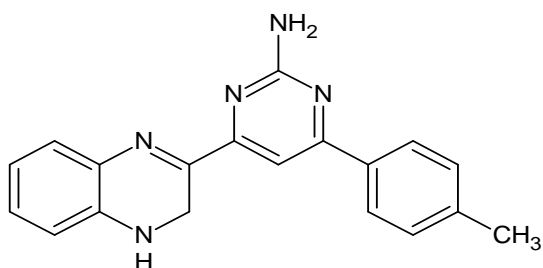
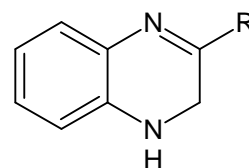


Fig.-13.

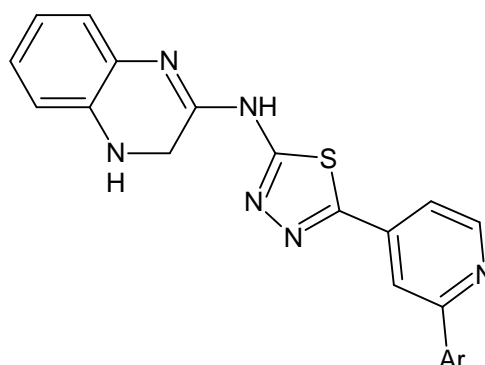
- El-Gohary NS, Shaaban MI *et al.*,(2018) benzimidazole derivatives were synthesized and evaluated for antimicrobial activity toward *Escherichia coli*, *Bacillus cereus*, *Staphylococcus aureus*, *Candida albicans*.<sup>[14]</sup>



R=H, Amino, Methyl.

Fig.-14.

- Barot KP, Manna KS, Ghate MD *et al.*,(2017) Synthesised benzimidazole derivatives and screened for its antimicrobial activity.<sup>[15]</sup>

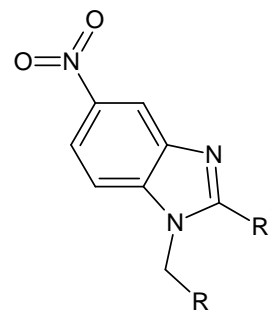


Ar=substituted hydrazide derivatives.

Fig.-15.

#### Miscellaneous Activities

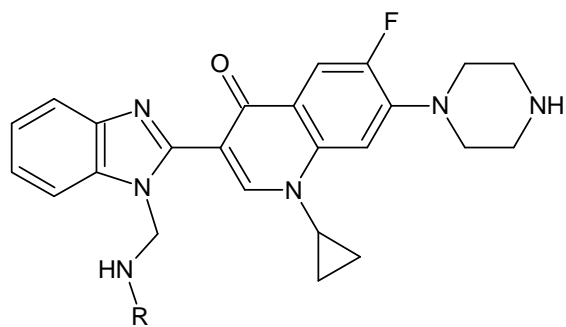
- Alam F, Dey BK, Sharma K, Chakraborty A, Kalita P *et al.*,(2017) synthesized derivatives of benzimidazole and screened for its anthelmintic activity.<sup>[16]</sup>



R=benzyl chloro, fluoro benzyl.  
R<sup>1</sup>=N N dimethyl, N N di ethyl.

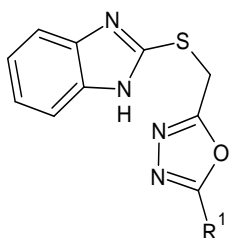
Fig.-16.

- Muluk R, Kothawade P, Kulkarni G, Ingale P *et al.*,(2017) synthesized a series of novel benzimidazole and evaluated for antidiabetic activity.<sup>[17]</sup>



R=Methyl, ethyl.  
Fig.-17.

- Shingalapur RV, Hosamani KM, Keri RS, Hugar MH *et al.*, (2010) Series of benzimidazole derivatives, a group of 4-thiazolidinones and 1,3,4-oxadiazoles containing 2-mercapto benzimidazole nucleus were synthesized and evaluated for *in vivo* anticonvulsant activity by Maximal Electroshock (MES) model and antidiabetic activity using Oral Glucose Tolerance Test (OGTT).<sup>[18]</sup>



R=hetero ring/halogenated aryl group.  
Fig.-9.

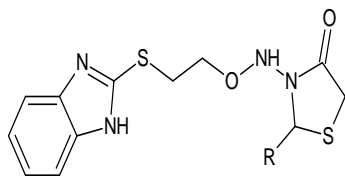


Fig.-20.

## CONCLUSION

The benzimidazole ring is an crucial pharmacophore in the field of drug discovery. Some substituted benzimidazole derivate drugs are having more effective activity. The synthesis of benzimidazole derivatives is a lucky scaffold, having a variety of therapeutic uses. From this literature review concluded that benzimidazole possess various biological activities like anticancer activity, anti parasitic activity, antioxidant activity, antimicrobial activity, antidiabetic activity, anthelmintic activity, anticonvulsant activity.

## ACKNOWLEDGMENT

I, **Shadiha Saheed K**, student of M-Pharm-Pharmaceutical Chemistry at Kerala University of Health Science like to express my thanks to Almighty for giving me power and patience to complete the review article in a successful way.

I, express my sincere gratitude to my HOD **Mr. Akash Marathakam** who is the source of inspiration for achieving greater height in the perusal of excellence. And also great pleasure to acknowledge my sincere gratitude to my project guide **Mrs.Thushara B S** for her valuable guidance during the preparation of this review

article. And also thanks to my batch mates **Midhula CC** and **Baijika P** for the co-operation to this article.

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