Anticancer Evaluation of Innovated Macrocyclic Sulfazan and Triazan Compounds as anticancer- Breast Drugs

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Abstract

Triazan with Macro cyclic Sulfazan as a novel class of compounds were attended and developed for the primary stage in organic chemistry in 2019 internationally by (Professor Dr. Nagham Aljamali), who began working on studying of these chemical compounds in the field of cancerous tumors and comparing infected cells with healthy cells by using many materials prepared from the interaction of chemical compounds that include donor atoms that can attack and treat malignant tumor cells, such as sulfur and nitrogen atoms, which are considered as a measure of the ability of drugs to treat Diseases through knowledge of the ability of triazan and formazan compounds to penetrate cellular membranes, while there are special medicinal compounds that often lose this ability, and in this case it was found that antibiotics become highly effective and their activity increases in the presence of groups of Formazan, Triazan and Sulfazan. Sulfazan compounds differ in their mode of action and activity in the life system depending on their composition. For example, inert (stable) metal complexes have good effectiveness against microbes, fungi, and viruses. They are also used to control the spread of cancerous tissues. All spectral studies appeared good results and strong evidences for their structures and all results of anticancer studies indicated to high efficiency of these compounds on cancer cells of breast.

Keywords: azan, Macrocyclic Sulfazan, Triazan, cancer, tumor, drug.

1. Introduction

Macrocyclic Sulfazan Compounds: Novel conceived Triazan and Sulfazan composites were innovated by (investigator Prof. Dr. Nagham) (1-3) in April 2019 for the primary stage internationally (1-3). The basics in addition to original approaches for synthesis of these novel derivatives and steps of reaction (5-7) have been positioned by Dr. Nagham as a novel procedure in organic chemistry field.

Triazan Compounds: Also it is Novel conceived Triazan composite was innovated by (investigator Prof. Dr. Nagham) (1-3) in April 2019 for the primary stage

internationally (1-3). The lack of studies and references related to it, its composition has a periodic arrangement of (-N = N-N-) in the periodic arrangement) (5-7) according to the type of derivative, It is prepared under different conditions and catalysts bestowing to the type of reactant of the amino compounds in fact(47-53), and the researcher provided sources and research papers on the methods of its preparation(1-3), and these innovative compounds were recently applied, so the academic (Dr. Nagham) operated on serial trainings For submissions of these novel composites such as fungal assay studies(5-7), bacterial assay, and cancer and oncological

studies to provide information on the influence of these composites. The dynamic groups in these compounds have different submissions.

2. Appliances and Trials:

All mechanism of preparation with instrumental analysis carried out in Tehran Universities and some of them in center of chemical lab in pharmacy college , Also anticancer Studies for two Bio-compounds.

Designation Procedures:

Designation of Triazole [1]:

1,4-Dicarboxy benzene (0.01 mol , 1.66g) was retorted with (NH2NH-CS-NH2) (0.02 mol, 1.82g) in 45 ml (C2H5-OH)in basic medium (5% alkali solution) in alternation and condensation step for 35 hrs, then ring fused step via condensation reaction and cyclization step by cyclization of ester group with thiosemicarbazide , the product filtered, dried , recrystallized, according to procedures(1-3).

Designation of Inventive Macrocyclic Sulfazan- Triazan [2]:

Chaos.1: Designation of Triazole [1] as a drug

Starting composite [1]was (0.01 mol, 2.76g) dissolved in basic solution from (alkali solution) then responded with (0.02 mol, 5.80g) of di-azo salt of Trimethoprime by way of sundry steps in basic solution to construction invented Macrocyclic Sulfazan-Triazan subsequently (60 hrs), the merchandise cleaned ,dehydrated ,eroded by refined water, recrystallized to revenue Invented Macrocyclic Sulfazan-Triazan bestowing to techniques(1-3).

Designation of Inventive Macrocyclic Sulfazan- Triazan [3]:

Starting composite [1]was (0.001 mol, 0.276g) dissolved in basic solution from (Piprydine) then reacted with (0.002 mol,0.216g) of diazo salt of Benzene-1,4-by way of sundry steps in basic solution to construction invented Macrocyclic Sulfazan-Triazan subsequently (52 hrs), the merchandise cleaned ,dehydrated ,eroded by refined water, recrystallized to revenue Invented Macrocyclic Sulfazan-Triazan bestowing to techniques (1-3).



Chaos.2: Designation of Conceived Macrocyclic Sulfazan-Triazan [2, 3] as a drugs



3. RESULTS AND DISCUSSION:

Invented Triazan and Macrocyclic Sulfazan have been Designated in equivalent process that shadowed and conceived by the academic Dr. Nagham in year 2019 via series of studies (1-3) that got a patent for invention of Triazan and Macrocyclic Sulfazan (1, 2) :

Spectral Part :

FT.IR- Supernatural Suggestion of Macrocyclic Triazan and Sulfazan : It Statistic. (1): Infrared [1]

designated to foundation our Conceived Sulfazan by exterior of gang at :(2400) cm-1 for (SH) mercapto cluster in triazole-thiol [1] which is vanished and other bands appeared as an alternative of it characterized in (-S-N=N-) Sulfazan and (-N-N=N-) Triazan group at : {(1379,1429,1477), (1406,1438,1498)} cm-1 respectively, and band at (1311) for band (N-N-N) in Macrocyclic Sulfazan and Triazan compounds [2, 3] correspondingly, statistics (1-3) :



Statistic. (2): Infrared [2]



Statistic. (3): Infrared [3]



H.NMR – Rang Suggestion of Macrocyclic Sulfazan, Triazan: It designated to foundation our Conceived triazole by entrance of highest at : δ (4.36) for (SH) mercapto cluster and peak at (5.55) for (NH) of amine group of **Statistic. (4): Resonance [1]**

Triazole amine [1] that vanished in Triazan and Macrocyclic Sulfazan [2, 3] due to formation Triazan and Sulfazan [2, 3] Statistics (4-6) :



Statistic. (5): Resonance [2]



Statistic. (6): Resonance [3]



C.NMR - rang Suggestion of Triazan and Macrocyclic Sulfazan : It indicated to formation our Invented Sulfazan compounds by appearance of peaks at : δ (106.0-141.9) for signals of Aromatic carbon atoms in Triazan and Sulfazan compounds [2, 3], also peak at : δ (66.0) for signal of carbon atom of **Statistic. (7): Carbon-Resonance [2]** methoxy group (-OCH3) in Sulfazan compound [2], other peak at : δ (21.56) for signal of carbon atom of methylene group (-CH2) in Sulfazan compound [2], all other signals clarified according to reference(30), all data signals in Statistic. (7):



Anti-malignance Inspection (1-3):

Triazan besides to Sulfazan {2, 3} were selected for anticancer evaluation (breast cancer) by dint of MTT-Investigation of dualistic sorts of Cells (MCF-7) as Malignancy cell besides (MCF-10 A)as well cell affording to revisions [1-3], all statistics (8, 9), Stands (1,2). The results gave good data indicated to high inhibition of tumor cells. Stand. (1): Cytotoxic Commotion of Sulfazan-Triazan Composite [2] continuously Breast Malignancy Cells (MCF-7) and Well Cells (MCF-10A) at the equivalent meditation exhausting 24 hrs., MTT trial 370c.

Dose	Proportion (%) in place of every selected cells				
$(\mu g.mL^{-1})$					
	MCF-7 / IC50= 12.76		MCF-10A / IC50 = 196.82		
Triazan -Sulfazan {2}	Malignant cells MCF-7		Common cells MCF-10A		
	Cell Viability	Cell -Inhibition	Cell Viability	Cell-Inhibition	
15.62	90.63	9.37	85.43	14.57	
31.25	87.81	12.19	89.75	10.25	
62.5	80.92	19.08	91.64	8.36	
125.0	54.78	45.22	93.32	6.68	
250	47.48	52.52	94.58	5.42	
500	43.02	56.98	95.76	4.24	
Control	100		96.20	3.80	

Stand. (2): Cytotoxic Commotion of Triazan-Sulfazan Composite [3] continuously Breast Malignancy Cells (MCF-7) and Well Cells (MCF-10A) at the equivalent meditation exhausting 24 hrs., MTT trial 370c.

$\frac{\text{Dose}}{(mam L^{-1})}$	Proportion (%) in place of every selected cells				
(µg.mL)	MCF-7 / $IC_{50} = 14.45$		MCF-10A / IC ₅₀ = 194.98		
Triazan -Sulfazan	Malignant cells MCF-7		Common cells MCF-10A		
{3 }	Cell -Viability	Cell- Inhibition	Cell -Viability	Cell -Inhibition	
15.62	90.55	9.45	89.01	10.99	
31.25	86.43	13.57	90.71	9.29	
62.5	80.36	19.64	93.48	6.52	
125.0	76.81	23.19	94.99	5.01	
250	50.44	49.56	95.20	4.8	
500	45.71	54.29	95.67	4.33	
Control	100		96.45	3.55	

Fig. 8: Anti-malignancy commotion of Triazan-Sulfazan {2} on (MCF-7) at 500µg/ml Fig. 9: Anti-malignancy commotion of Triazan-Sulfazan {2}on (MCF-A 10) at 500µg/ml





4. Conclusions:

We distinguished that Macrocyclic Triazan-Sulfazan [2] abstains higher competence on Malignancy cells and extraordinary spontaneity than Triazan-Sulfazan [3] owing to attendance of paired sets of Triazan in addition to Sulfazan groups (-S-N=N) and (-N-N=N-) linked with trimethoprim in same compound.

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