

**أبحاث جامعة سيناء المنشورة
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[Investigation of Racah Parameters and \$\gamma\$ -ray Protection Features of Borate-Glasses Containing Fe₂O₃ and Er₂O₃ for Optical and Radiation Shielding Applications](#)

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Design, synthesis and biological screening of some diaryl acetamide derivatives as potential cytotoxic agents on HL-60(TB) cell line

Bioorganic Chemistry • Article • 2025 •

DOI: 10.1016/j.bioorg.2025.108999



Abdelhameid, Mohammed K.^a ; Hara, Mohammed A.^b; Ramadan, Mohamed;
Taher, Ehab S.^{b,c}; A.Ramadan, Mostafa^d;

^b 

+2 authors

^a Department of Pharmaceutical Organic Chemistry, Faculty of Pharmacy,
Cairo University, Cairo, 11562, Egypt

Abstract

In this study, novel diaryl acetamide derivatives were designed and evaluated as potential cytotoxic agents against the leukemic cells HL-60

(TB) with tyrosine kinase FMS-3(FLT-3) enzyme inhibitory activity. The designed molecules were synthesized and investigated for the FLT3 enzyme inhibition. Compounds 5a, 15b, and 16b showed significant inhibitory activity against the FLT-3 enzyme. Assessment of cytotoxicity on HL-60 cells revealed that compound 15b exhibited superior activity than Quizartinib (AC220). Additionally, compounds 5a and 15b effectively arrested the cell cycle at the G₁ phase upon testing on HL-60 cells, suggesting potential abilities in blocking HL-60 cells' proliferation. The annexin-V stain assay demonstrated that compounds 5a, 15b, and 16b induced apoptosis in HL-60 cells. Furthermore, the ELISA assay showed that the blocking of cell cycle proliferation of HL-60 cells was mediated via the induction of cell cycle regulatory proteins p53 and p21. Meanwhile, the observed apoptosis induction was mediated by increasing apoptotic mediators, Bax/Bcl-2 ratio, and up-regulating caspase-3. Molecular docking studies revealed the binding affinities and interactions of the newly synthesized compounds, along with the reference compound Quizartinib, with the FLT-3 enzyme binding site. © 2025 Elsevier Inc.

Author keywords

Diaryl acetamide; Dimedone; FLT-3 kinase inhibitors; HL-60 (TB); Leukemia; Oxadiazole; Quizartinib; Thiazole; Triazole

Indexed keywords

EMTREE drug terms

2 [[4 (4 chlorophenyl)thiazol 2 yl)amino] n [4 [(5,5 dimethyl 3 oxocyclohex 1 en 1 yl)amino]phenyl]acetamide; 2 [[5 (4 chlorophenyl) 1,3,4 oxathiazol 2 yl)thio] n [4 [(5,5 dimethyl 3 oxocyclohex 1 en 1 yl)amino]phenyl]acetamide; 2 [[5 (4 chlorophenyl) 4 phenyl 4h 1,2,4 triazol 3 yl)thio] n [4 [(5,5 dimethyl 3 oxocyclohex 1 en 1 yl)amino]phenyl]acetamide; 5,5 dimethyl 1,3 cyclohexanedione; acetamide derivative; antineoplastic agent; caspase 3; cytotoxic agent; diaryl acetamide derivative; Flt3 ligand; protein Bax; protein bcl 2; protein p21; protein p53; quizartinib; unclassified drug

EMTREE medical terms

alkylation; antineoplastic activity; antiproliferative activity; Article; binding affinity; carbon nuclear magnetic resonance; controlled study; drug design; drug screening; drug synthesis; enzyme binding; enzyme inhibition; enzyme linked immunosorbent assay; flow cytometry; G1 phase cell cycle checkpoint; HL-60 cell line; human; human cell; in vitro study; molecular docking; proton nuclear magnetic resonance

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5,5 dimethyl 1,3 cyclohexanedione	126-81-8
protein bcl 2	219306-68-0

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Funding text 2

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Corresponding authors

Corresponding author

M.K. Abdelhameid

Affiliation

Department of Pharmaceutical Organic Chemistry, Faculty of Pharmacy, Cairo University, Cairo, 11562, Egypt

Email address

mohamed.sayed@pharma.cu.edu.eg

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