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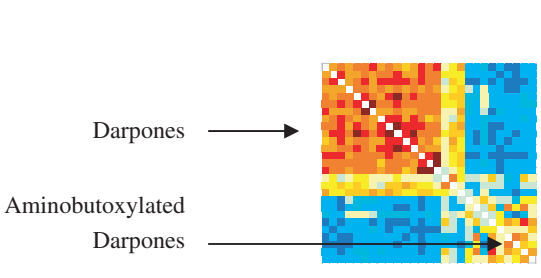
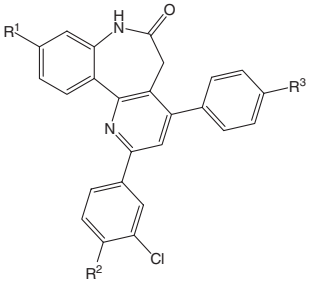
Chemistry
Letters

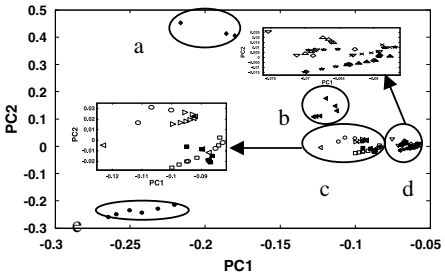
Bioorganic & Medicinal Chemistry Letters Vol. 17, No. 7, 2007

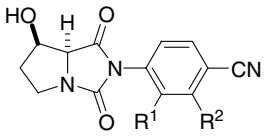
Contents

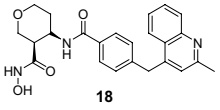
ARTICLES











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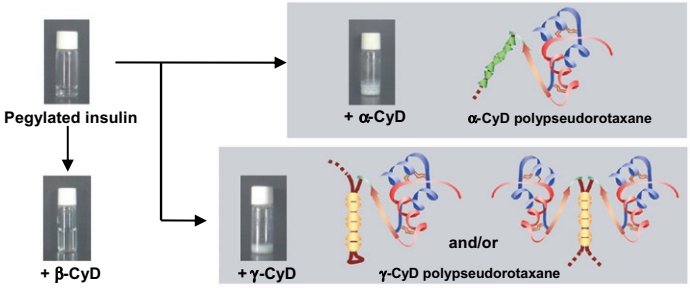
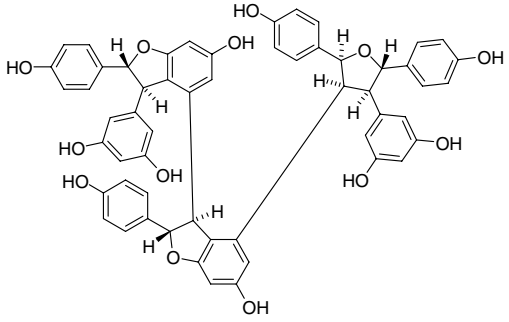
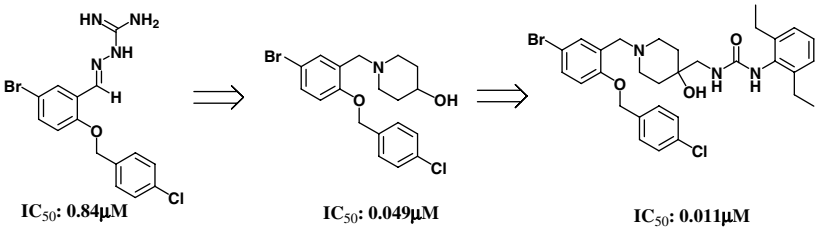
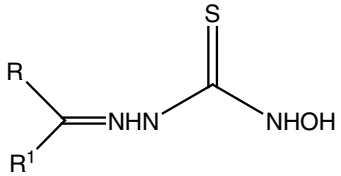
p-TACE IC_{50} = 1.0 nM
WBA IC_{50} = 130 nM





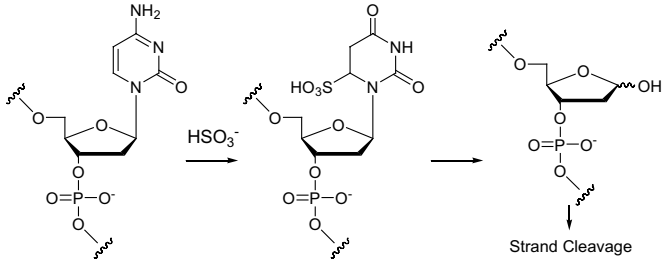


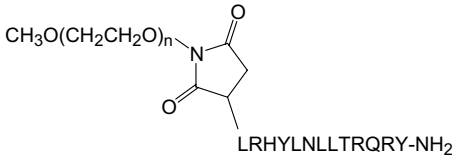
Optimizing the guanylhydrazone of 2-(4-chlorobenzyloxy)-5-bromobenzaldehyde, which is a hit from high-throughput screening (HTS), let to discover the potent 4-hydroxypiperidine derivatives as a CCR5 receptor antagonist.



10a-10r

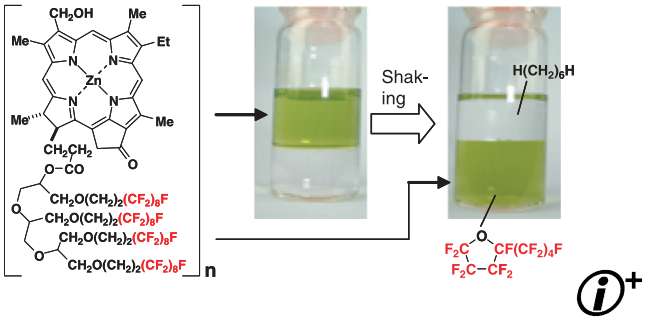
Eighteen *N*-hydroxythiosemicarbazones were prepared and tested for their in vitro activity against *Mycobacterium tuberculosis* H37Rv using the agar dilution method.

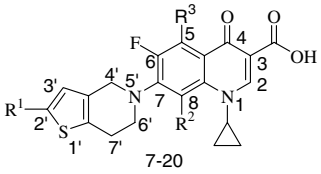




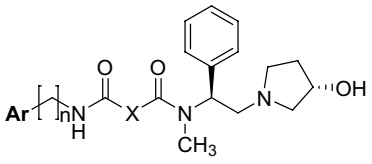
PEG SAR studies identify a potent appetite suppressant.

Synthetic zinc chlorophyll possessing four perfluorooctyl groups self-aggregated in a hexane solution and the resulting highly ordered *J*-aggregates were dissolved in fluorous solvents by biphasic extraction.



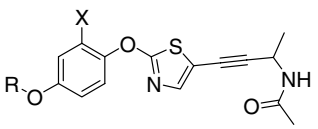


Synthesis and antibacterial activity of a number of substituted 4,5,6,7-tetrahydro-thieno[3,2-*c*]pyridine quinolones is reported. The antibacterial activities were evaluated in standard in vitro MIC assay method. Some of the compounds showed in vitro (MIC) antibacterial activity comparable to those of Gatifloxacin, Ciprofloxacin, and Sparfloxacin.

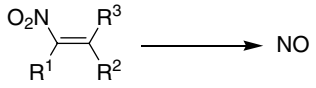


A novel series of malonamide derivatives was synthesized. These amides were shown to be potent and selective κ opioid receptor agonists.

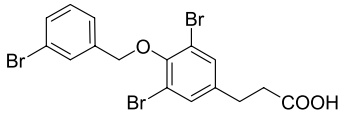
N_1-arylsulfonyl-1,3-dihydro-2*H*-benzimidazol-2-one as a novel template for the design of new anti-HIV agents is reported.



Optimization studies aimed at improving the potency and selectivity profiles of a recently discovered series of acetyl-CoA carboxylase (ACC) inhibitors are described. The incorporation of 2-position phenyl ring substituents resulted in dramatic and general ACC2 selectivity enhancements (>3000-fold) within the lead series.

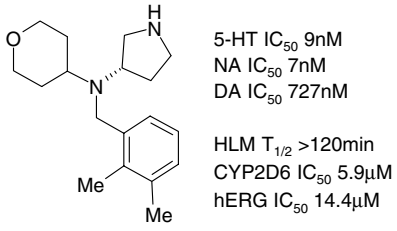


Nitroalkenes decompose to small amounts of NO as judged by chemiluminescence NO detection.

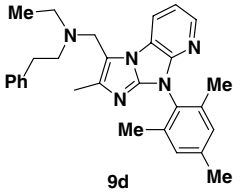


11f

Based on the concept of 'indirect antagonism' of nuclear receptors, a series of thyroid hormone receptor (TR) antagonists were prepared with improved affinity compared with what was previously described. The results of a binding assay for the human TR and reporter cell assay revealed, within this series, that an *m*-bromobenzoyl substituent (**11f**) was optimal in terms of affinity and antagonist activity.

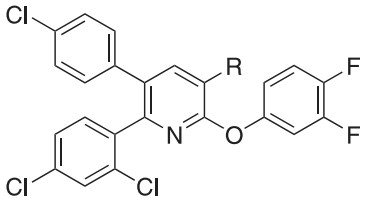


A series of amino-pyrrolidines with potent dual 5-HT/NA reuptake inhibition, good metabolic stability and weak CYP2D6 inhibition is described.



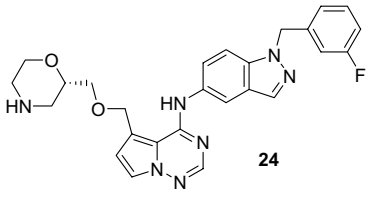
9d

The synthesis and SAR of tetraazacyclopenta[*a*]indenes, pharmacokinetic properties and the anxiolytic activity of an orally dosed exemplary compound **9d** (K_i 8.0 nM) are reported.



The synthesis and biological optimization for 5,6-diarylpyridines as CB1 receptor inverse agonists is described.

Novel C-5 substituted pyrrolotriazines were optimized for dual EGFR and HER2 protein tyrosine kinase inhibition. The lead compound exhibited promising oral efficacy in both EGFR and HER2 driven human tumor xenograft models. It is hypothesized that its C-5 morpholine side chain binds in the ribose phosphate portion of the ATP binding pocket.



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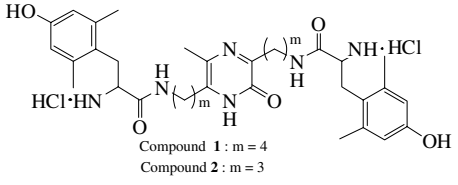
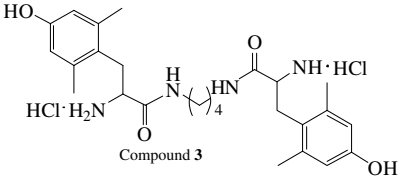
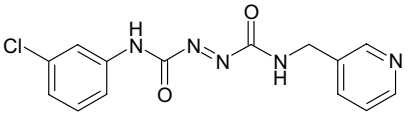


Compound **1** : m = 4
Compound **2** : m = 3



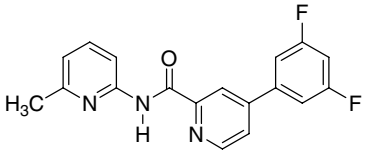
Compound **3**

Compound **4** : Endomorphin 2 H-Tyr-Pro-Phe-Phe-NH_2

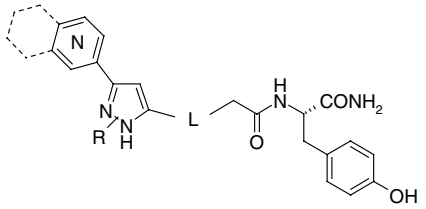


6eC
IC_{50}=15 μM

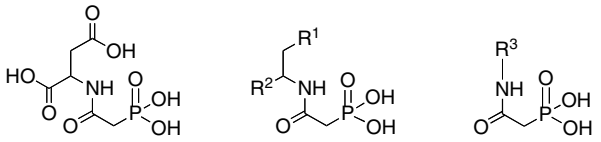
A series of new inhibitors of the D-alanine-D-alanine ligase (Ddl) is presented. Thirteen diazenedicarboxamides were better inhibitors than D-cycloserine, and some of them also possess antibacterial activity, which makes them a promising starting point for further development.



mGluR5 Ki = 0.04 ± 0.01 μM



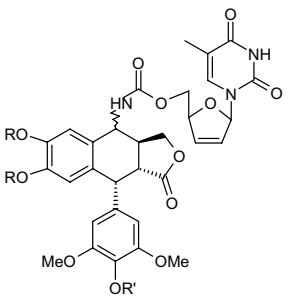
The synthesis and structure–activity relationship of pyrazole derivatives are described as novel low molecular weight LH-receptor agonists.



PALA **1-4** **5-9**

A series of novel PALA derivatives were synthesized and ability to inhibit aspartate transcarbamoylase was determined.

A novel series of derivatives of podophyllotoxin as anti-HIV-1 agents were synthesized, **19d** and **19c** showed the highest anti-HIV activity with EC_{50} values of 0.17 and 0.29 μM and TI values of 466.9 and 354.5, respectively.



OTHER CONTENTS

*Corresponding author
ⓘ+ Supplementary data available via ScienceDirect

COVER

The identification of peptidomimetic inhibitors that selectively disrupt STAT1 or STAT3 homodimerization at low μM concentrations [Gunning, P. T.; Katt, W. P.; Glenn, M.; Siddique, K.; Kim, J. S.; Jove, R.; Sebti, S. M.; Turkson, J.; Hamilton, A. D. *Bioorg. Med. Chem. Lett.* **2007**, *17*, 1875].

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