

# Molecular design and synthesis of histone deacetylase inhibitors as antineoplastic agents

Thesis

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## List of abbreviations

2D: 2-Dimensional

3D: 3-Dimensional

3D QSAR: 3-Dimensional Quantitative structure activity relationship

4-DMAP: 4-dimethylaminopyridine

Å: Angstrom

Adj: Adjusted

AML: Acute myeloid leukemia

Asn: Asparagine

Asp: Aspartate

Bcl-2: B-cell lymphoma 2

BSA: Bovine serum albumin

CDK: Cyclin dependent kinase

CHAPs: Cyclic hydroxamic acid-containing peptides

CLL: Chronic lymphocytic leukemia

CML: Chronic myeloid leukemia

CTCL: Cutaneous T-cell lymphoma

DMF: Dimethylformamide

DMSO: Dimethyl sulphoxide

DNA: Deoxyribonucleic acid

EDTA: Ethylenediaminetetraacetic acid

ELISA: Enzyme linked immunosorbent assay

FBS: Fetal bovine serum

FDA: Food and Drug Administration

FT-IR: Fourier transform-Infrared

HAT: Histone acetyl transferase  
HBA: Hydrogen bond acceptor  
HBD: Hydrogen bond donor  
HCC: Hepatocellular carcinoma  
HDA1: Histone deacetylase 1  
HDAC: Histone deacetylase  
HDACIs: Histone deacetylase inhibitors  
HDAH: Histone deacetylase like amidohydrolase  
HDLP: Histone deacetylase-like protein  
HIF: Hypoxia inducible factor  
His: Histidine  
HL: Hodgkin's Lymphoma  
H: Hours  
HSP90: Heatshock protein 90  
HYP: Hydrophobic  
IC<sub>50</sub>: 50% Inhibitory concentration  
MHz: Mega Hertz  
Mp: Melting point  
MS: Mass spectroscopy  
NCI: National Cancer Institute  
NMP: N-methyl-2-pyrrolidone  
NMR: Nuclear magnetic resonance  
PDB: Protein data bank  
Phe: Phenylalanine  
POC: phosphorous oxy chloride

ppm: part per million

Pred: predicted

RA: Ring aromatic

Ras: Rat sarcoma

RNA: Ribonucleic acid

ROS: Reactive oxygen species

Rpd3: Reduced potassium dependency 3

RPMI: Roswell Park Memorial Institute medium

RUNX: Runt-related transcription factor

SAHA: Suberoylanilide hydroxamic acid

SIRT: Sirtuins

SNAr: Nucleophilic Aromatic Substitution reactions

SUMO: Small Ubiquitin-like Modifier

SRB: Sulforhodamine B

SRM: Surface recognition moiety

TCA: Trichloroacetic acid

THF: Tetrahydrofuran

TPX: Trapoxins

TSA: Trichostatin A

TSG: Tumor suppressor gene

VEGF: Vascular endothelial growth factor

VEGFR: Vascular endothelial growth factor receptor

ZBG: Zinc binding group

## **Abstract**

### *Title of Thesis*

# **Molecular design and synthesis of histone deacetylase inhibitors as antineoplastic agents**

### *Name of candidate*

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