

بسم الله الرحمن الرحيم



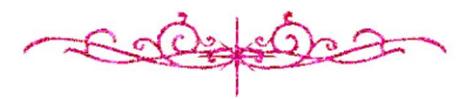
-Call 1600-2

COEFOR COEGOROS





شبكة المعلومات الجامعية التوثيق الالكتروني والميكروفيلم



CORRECT CORRECTOR



جامعة عين شمس

التوثيق الإلكتروني والميكروفيلم قسم

نقسم بالله العظيم أن المادة التي تم توثيقها وتسجيلها علي هذه الأقراص المدمجة قد أعدت دون أية تغيرات



يجب أن

تحفظ هذه الأقراص المدمجة بعيدا عن الغبار

-Caro-

COEFERS CARGORNOR





بعض الوثائق

الأصلية تالفة



CORRECT CORRECTION





بالرسالة صفحات

لم ترد بالأصل



COEFECT CARGOSTON

SYNTHESIS OF CERTAIN TETRAHYDRONAPHTHYL THIAZOLES OF POSSIBLE BIOLOGICAL ACTIVITY

THESIS

PRESENTED BY

MANAL MOHAMED ANWAR HASAN

(B. Pharm. Sci. 1986)

Submitted in Partial Fulfilment

For Master Degree in

PHARMACEUTICAL SCIENCES

(PHARMACEUTICAL CHEMISTRY)

UNDER THE SUPERVISION OF

Prof. Dr. Mohamed Yassin Ebeid
Prof. of Pharmaceutical Chemistry,
Faculty of Pharmacy,

Cairo University.

Prof. Dr. Mohsen Mohamed Kamel
Prof. of Medicinal Chemistry
National Research Centre.

Dr. Magdy Ibrahim El Zahar

Associate Prof. of Medicinal Chemistry,

National Research Centre.

Faculty of Pharmacy-Cairo University
1995



APPROVAL SHEET

This thesis has been approved by the examination committee on /

/1995

Prof. Dr. Ahmad Mahmoud Farghaly

Professor of Pharmaceutical Chemistry,
Faculty of Pharmacy,
Alexandria University.

A Laplas

Prof. Dr. Mohamed Yassin Ebeid

Professor of Pharmaceutical Chemistry,
Faculty of Pharmacy,
Cairo University.

M. Shand

Prof. Dr. Mohsen Mohamed Kamel
Professor of Medicinal Chemistry,
National Research Centre.

M. M. Karul

Prof. Dr. Kamelia Mahmoud Amin

Professor of Pharmaceutical Chemistry,

Faculty of Pharmacy,

Cairo University.

XQ:

APPROVAL SHEET

This thesis has been approved by the examination committee on / /1995

Prof. Dr. Ahmad Mahmoud Farghaly

Professor of Pharmaceutical Chemistry,
Faculty of Pharmacy,
Alexandria University.

Prof. Dr. Mohamed Yassin Ebeid

Professor of Pharmaceutical Chemistry,
Faculty of Pharmacy,
Cairo University.

Prof. Dr. Mohsen Mohamed Kamel
Professor of Medicinal Chemistry,
National Research Centre.

Prof. Dr. Kamelia Mahmoud Amin

Professor of Pharmaceutical Chemistry,

Faculty of Pharmacy,

Cairo University.

ACKNOWLEDGEMENT

First of all, I thank God for giving me the potential to achieve this work.

I would like to express my deepest appreciation to **Prof. Dr. Mohamed Yassin Ebeid**, Professor of Pharmaceutical Chemistry, Faculty of Pharmacy,

Cairo University, for his active supervision, encouragement, kindness and generous help during this work.

Mohamed Kamel, Professor of Medicinal Chemistry, Medicinal Chemistry Department, National Research Centre, for his supervision and planning the course of investigation. Helpful co-operation and sincere advices offered by him were the principal factors which brought this work to the present standard.

I am also deeply grateful to **Dr. Magdy Ibrahim El Zahar**, Assistant Professor of Medicinal Chemistry, Medicinal Chemistry Department, National Research Centre, for his supervision, encouragement and help during this work.

Also, I would like to express my deep thanks to **Dr. Omar Abdel**Fatah, Assistant Professor of Medicinal Chemistry, Medicinal Chemistry

Department, National Research Centre, for his kind help during this work.

Many Thanks to Dr. Abdel Hamid Zaki Abdel Hamid Associate Professor of Biochemistry, Medicinal Chemistry Department, National Research Centre, for his help during performing the biological part, and to Prof. Dr. Mostafa Abdalla, Head of Biology Department, Faculty of Education-Ain Shams University, for his help in antimicrobial screening.

Deep thanks to every member in National Research Centre who helped and supported me during this work.

CONTENTS

· *

ì

	Page
- Abstract	
	1
- Introduction	1
- Aim of the present investigation	34
- Theoretical discussion	39
Section 1: Synthesis of certain tetrahydronaphthyl-	
thiazoles.	44
Section 2: Synthesis of some new thiazolidinone	
and rhodanine-Mannich bases.	52
Section 3: Synthesis of new 1,2,3,4-tetrahydro-	
naphthylthiazolidin-4-one derivatives	
and their related products.	59
- Biological part.	67
- Experimental part	79
- References.	100
- A robic cummary	

Ø

Synthesis of certain tetrahydronaphthyl thiazoles of possible biological activity

Abstract

In the present work, a variety of tetrahydronaphthyl thiazole derivatives was prepared for the purpose of evaluation as antiparasitic or as antimicrobial agents. These compounds include 1-alkyl(or aryl)-3-[4-[6-(1,2,3,4-tetrahydronaphthyl) thiazol-2-yl] ureas and thioureas. In addition, several reactions were carried out on the heterocyclic compounds to obtain a series of possible biologically active N-3-[6-(1,2,3,4-tetrahydronaphthyl)]-2-arylthiazolidin-4-ones and their Mannich bases. Synthesis were achieved through different routes of chemical reactions and transformations starting with 6-amino tetralin and 2-amino-4-[6-(1,2,3,4-tetrahydronaphthyl)]thiazole. All of the new compounds were confirmed with concordant microanalytical data, IR, ¹HNMR and Mass spectra.

Some of the newly synthesized compounds were found to exhibit various inhibitory effects on egg masses production of <u>Biomphalaria</u> alexandrina snails, the intermediate host of <u>Schistosoma mansoni</u>. Also some other compounds showed antimetabolic effect in experimental animals livers, and antimicrobial effects.

Moreover, one of the newly synthesized compounds (Compound IVc) showed moderate activity against human immuno deficiency virus type-1 (HIV-1) (Anti-Aids) and anticancer against malignant tumors (anticancer)*

^{*(}Department of Health and Human Services, National Institute of Health, National Cancer Institute, Bethesda, Maryland, U.S.A.) (cf. the enclosed letter of the said institute). Biological part page 78.

7

The thesis comprises the following parts:

A- Introduction:

This part includes a survey of the different chemical, biological and pharmacological activities of tetrahydronaphthalene (tetralin) derivatives and those incorporated into heterocyclic moieties.

B- Aim of the present investigation:

This part includes the chemical and pharmacological basis on which the synthesized groups of the new compounds were chosen.

C- Theoretical discusion:

This part deals with the main commonly applied methods for the preparation of the intermediates and the different pathways leading to the final compounds.

furthermore, it contains the investigation of some reactions and the results.

D- Experimental part:

This part includes the methods of the synthesis of the starting materials, the intermediates and the new products.