# Analytical studies on clomipramine and paroxetine hydrochloride drugs

## Presented by

Mohamed Abdel-Razik Mohamed (B.Sc.Degree, Chemistry, Cairo University; 1999)

A Thesis Submitted For the Partial Fulfillment of The Degree of M.Sc. in Science (Analytical Chemistry)

To

**Chemistry Department Faculty of Science Cairo University** (2010)

#### **ABSTRACT**

Name: Mohamed Abdel-Razik Mohamed.

Title of thesis: Analytical studies on clomipramine and paroxetine hydrochloride drugs.

Degree: (M.Sc.) Un Published Master of Science, Faculty of Science, Cairo

University, 2009/2010.

This work has been carried out to investigate the spectrophotometric determination of Paroxetine (PRX) and clomipramine (CLO) antidepressant drugs in pure and pharmaceutical preparation so the spectrophotometric methods involve the reaction of these drugs through ion pair reaction with Mo(V)-thiocyanate binary complex and inorganic complexes like iron(III)-thiocyanate followed by their extraction in the organic phase (1,2–dichloroethane or methylene chloride) and determination by absorption spectrometry. The sensitivity of these procedures for the spectrophotometric determination of these drugs has been improved and the optimum practical conditions for the analysis of these antidepressant drugs are selected. In additions the second part of this research is the formation of ion selective electrode for PRX and CLO drugs. Ion selective electrodes are a technology that gives direct measurement of these drugs. Time consuming steps such as filtrations, weightings and distillation are not required in most cases. Concentrations can be read out directly on a pH/MV/ION meter, or can be read from a constructed calibration curve. Electrochemical methods save time, and since electrodes are portable.

Keywords: Paroxetine, Clomipramine, Mo(V)-thiocyanate binary complex, iron thiocyanate, ion-pair formation reaction, ion selective electrodes.

#### **Supervisors:**

### signature:

1. Prof. Dr: Gehad G. Mohamed

2. Dr: Huda Mustafa Mahmud

3. Dr: Eman Yossiry Zaki.

Prof. Dr. Mohamed A. Badawy

Chairman of Chemistry Department Faculty of Science- Cairo University

### **APPROVAL SHEET FOR SUBMISSION**

Title of [M.Sc] Thesis: Analytical studies on some antidepressant drugs.

Name of the candidate: Mohamed Abdel-Razik Mohamed

This has been approved for submission by the supervisors:

1. Prof. Dr: GEHAD GENIDY MOHAMED

Professor of Inorganic and Analytical Chemistry, Chemistry Department, Faculty of Science, Cairo University.

Signature:

2- Dr: Huda Mustafa Mahmud.

Lecturer of Analytical Chemistry, National Organization for Drug Control and Research.

Signature:

Prof. Dr. Mohamed A. Badawy

Chairman of Chemistry Department

Faculty of Science- Cairo University

## Preliminary courses for M.Sc.

Beside the work carried out in this thesis, the candidate had attended and successfully

passed a final examination of M.SC. Courses (2006) in non	-organic chemistry covering the
Following topics:	
1. Inorganic reaction mechanism.	
2. Advanced analytical chemistry.	
3. Compleximetry.	
4. Statistical thermodynamics.	
5. Quantum chemistry.	
6. Molecular structure.	
7. Molecular spectroscopy.	
8. X-ray analysis.	
9. Surface chemistry.	
10. Electrochemistry of molten salts and physical metallurgy.	
11. Voltammetry.	
12. Nuclear chemistry.	
13. Electrochemistry.	
14. Electro kinetic's phenomena.	
15. Thermal method of analysis.	
16. Inorganic chemistry.	
17. Solar energy.	
18. Polymer.	
19. German language.	Prof. Dr. Mohamed A. Badawy

Chairman of Chemistry Department

Faculty of Science-Cairo University

#### **AKNOWLEGMENT**

I am deeply indebted to Prof. Dr. Gehad G. Mohamed, professor of Inorganic and Analytical Chemistry, Chemistry Department, Faculty of Science, Cairo University, for his standing help, excellent scientific guidance, suggesting the plane of research and his valuable contribution to this work to come to light.

Many thanks to Dr. Huda Mustafa, Lecturer of Analytical Chemistry, National Organization for Drug Control and Research (NODCR) for providing us with standard drug materials, suggesting the plane of research, revising the work, her valuable contribution and encouragements throughout the whole work.

Finally, I would like to thank all my colleague in the department of chemistry, who helped me in the preparation and completion of this work.

Mohamed Abdel-Razik Mohamed

#### **Abbreviation**

PRX Paroxetine

CLO Clomipramine

SSRIs Selective serotonin-reuptake inhibitors

5-HT 5-Hydroxytryptamine

PVC Polyvinylchloride

TCAs Tricyclic antidepressants

SNRIs Serotonin-norepinephrine reuptake inhibitors

HPLC High performance chromatography

UV Ultra violet

LC-MS/MS Liquid chromatographic – tandem mass spectrometric method

QC Quality control

BDD Boron-doped diamond

TLC Thin layer chromatographic method

ESI Electrospray ionization

SBSE/LC Stir bar sorptive extraction and liquid chromatography

NSAIDs Non-steroidal anti-inflammatories

LOD Limit of detection

LOQ Limit of quantification

CV Coefficient of variation

WW Waste waters

PPY Poly (pyrrole)

SWAdSV Square-wave adsorptive-stripping voltammetry

FIA-SWAdSV Flow-injection analysis with SWAdSV detection.

HMDE Hanging mercury drop electrode

FIA Flow-injection analysis-

OCD Obsessive-compulsive disorder

Rf Retardation factor

TCNQ Tetra-cyanoquinodimethane

cvm Continuous variation method

mrm Molar ratio method

MRM Multiple Reactions Monitoring



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#### Aim of the present work

In the recent years, many scientists have interested on account of the rapid valuation in the field of drug research industry. These include research work applying modern techniques and new methodologies of analysis. They tried to develop more selective and sensitive methods for antidepressant drug analysis. It is important in this field of research to develop new analytical techniques, and / or modify the already existing procedures, in order to follow up the test of purity of pharmaceutical compounds. Studying the properties of pharmaceutical compounds, via interaction with organic compounds, leads to results actually of analytical importance. Paroxetine (PRX) and clomipramine (CLO) antidepressants are the active ingredients of many pharmaceutical preparations concerned with the treatment of the obsessive compulsive perturbation states of phobia and panic as well as in depression and other emotional disturbances and major depressive, obsessive compulsive, generalized anxiety, social anxiety, post traumatic stress disorders. The medical importance of the antidepressant drugs under investigation, encourage us to suggest new methodology for the determination of these drugs. In the present work, a search for new spectrophotometric procedures is carried out on the various reactions of those drugs with Mo(V)-thiocyanate binary complex and inorganic complex Fe(SCN)<sub>6</sub><sup>3</sup>- to improve the sensitivity of these procedures for the spectrophotometric determination of these drugs. Many factors such as time, temperature and reagent concentrations were thoroughly investigated. The effect of these factors on the above mentioned reactions was studied, and the principal underlying the determination of PRX and CLO were investigated carefully in order to select the optimum practical conditions for the analysis of these antidepressant drugs.

The other part involves fabrication of ion selective electrode for PRX and CLO drugs determination. Ion selective electrodes are a technology that gives direct measurement of these drugs. These electrodes are rigid, inexpensive and characterized by its small size. ISEs presented here are based on polyvinylchloride (PVC) membrane supported with electro-active ion pair complexes. Time consuming steps such as filtrations, weightings and distillation are not required in most cases. Electrode methods save time, and since electrodes are portable, measurements can be made on a laboratory bench, the bank of a river or pond, the floor of a manufacturing plant, or the tray of a truck.

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