

UNIVERSITI TEKNOLOGI MARA

**CHEMICAL STUDIES OF
3-PYRROLIN-2-ONE
FOR THE SYNTHESIS OF
 γ -LACTAM γ -LACTONE**

AIMI SUHAILY BINTI SAAIDIN

Thesis submitted in fulfilment
of the requirements for the degree of
Master of Science

Faculty of Applied Sciences

March 2015

CONFIRMATION BY PANEL OF EXAMINERS

I certify that a Panel of Examiners has met on 14th January 2015 to conduct the final examination of Aimi Suhaily binti Saaidin on her Master of Science thesis entitled “Chemical studies of 3-pyrrolin-2-one for the synthesis of γ -lactam γ -lactone” in accordance with Universiti Teknologi MARA Act 1976 (Akta 173). The Panel of Examiners recommends that the student be awarded the relevant degree. The Panel of Examiners was as follows:

Nor’ashikin Saim, PhD

Professor
Faculty of Applied Sciences
Universiti Teknologi MARA
(Chairman)

Nor Hadiani Ismail, PhD

Professor
Faculty of Applied Sciences
Universiti Teknologi MARA
(Internal Examiner)

Zanariah Abdullah, PhD

Professor
Faculty of Science
Universiti Malaya
(External Examiner)

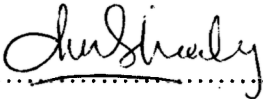
SITI HALIJJAH SHARIFF, PhD

Associate Professor
Dean
Institute of Graduate Studies
Universiti Teknologi MARA
Date : 24th March, 2015

AUTHOR'S DECLARATION

I declare that the work in this thesis/dissertation was carried out in accordance with the regulations of Universiti Teknologi MARA. It is original and is the results of my own work, unless otherwise indicated or acknowledged as referenced work. This thesis has not been submitted to any other academic institution or non-academic institution for any degree or qualification.

I, hereby, acknowledge that I have been supplied with the Academic Rules and regulations for Post Graduate, Universiti Teknologi MARA, regulating the conduct of my study and research.

Name of student	:	Aimi Suhaily binti Saaidin
Student I.D. No.	:	2011974393
Programme	:	Master of Science (Chemistry)
Faculty	:	Applied Sciences
Thesis Tittle	:	Chemical Studies of 3-Pyrrolin-2-one For The Synthesis of γ -Lactam γ -Lactone
Signature	:	
Date	:	March 2015

ABSTRACT

Unsaturated pyrrolidinone has been extensively studied due to its unique structure and its presence in various biologically active natural compounds. In this study, 3-pyrrolin-2-one, **162** has been constructed *via* five steps which included condensation of glycine methyl ester and methyl malonyl chloride, Dieckmann cyclization, decarbomethoxylation, reduction and elimination with an overall yield of 33%. The synthesized compound **162** was then used as the intermediate for a few chemical transformations such as Michael addition and epoxidation reaction. Synthesis of fused bicyclic 3,4- γ -lactone- γ -lactam towards mescaline isocitrimide was also attempted. The synthetic approach was divided into two different routes. The first route began by insertion of dimethyl malonate onto the synthesized intermediate of 3-pyrrolin-2-one *via* Michael addition reaction, followed by methylation of the β -diester and bromination at C3 position. However, the bromination reaction failed to give us the desired product which then diverted us to the second route. In this route, the γ -lactone- γ -lactam ring system was successfully synthesized by coupling readily available benzylated glycine methyl ester and methyl malonyl chloride followed by Dieckmann cyclization, alkylation at C3 position and Krapcho decarboxylation. The fused bicyclic ring was afforded through stereoselective reduction which accompanied concomitant lactonization with an overall yield of 0.8%. In brief, we have developed an operationally simple procedure towards the synthesis of 3-pyrrolin-2-one and fused bicyclic 3,4- γ -lactone- γ -lactam ring skeleton. The results extracted from this study thus far may be used to develop new scientific knowledge and remarkable findings.

ACKNOWLEDGEMENTS

First of all, I would like to express my deepest appreciations and sincere gratitude to my supervisor, Prof. Dr. Ahmad Sazali Hamzah for his valuable advice and continuous assistance through useful comments, suggestions and supports.

I would also like to register my heartfelt gratitude to my co-supervisor, Prof. Madya Dr. Zurina Haji Shaameri for her comments, guidance and a very helpful critical reading of the manuscript, without which it would not have been possible for me to shape the thesis in the present form.

I would very much like to thank all the staff in the Organic Synthesis and Natural Product laboratory, Puan Zaleha, Encik Sharizan and Puan Irmazatul for their invaluable assistance and unfailing support. I would also like to thank all my friends and colleagues of the block G especially Haff, Hanim, Huda, Ain, Wani, Syafiqah, Asmah, Haslini, Nisa', Khadijah, Hakimah dan Norli for their assistance whenever it was needed.

Last but not least, I would like to express my special thanks to my parents, Saaidin Abu Bakar and Nor Fuzaina Ismail for their constant prayers, patience and untiring support in every way during my study and not forgetting my siblings for their endless support and confidence in me. Most of all, I am very grateful to Allah for blessing me with good health and a lot of patience, without which this work will not be completed.