การสังเคราะห์และฤทธิ์กำจัดวัชพืชของกรคซินนามิกและสารประกอบที่เกี่ยวข้อง



นางสาว สุจิตตรา คีเสมอ

วิทยานิพนธ์นี้เป็นส่วนหนึ่งของการศึกษาตามหลักสูตรปริญญาวิทยาศาสตรมหาบัณฑิต สาขาวิชาเคมี ภาควิชาเคมี คณะวิทยาศาสตร์ จุฬาลงกรณ์มหาวิทยาลัย ปีการศึกษา 2543 ISBN 974-347-270-3 ลิขสิทธิ์ของจุฬาลงกรณ์มหาวิทยาลัย

SYNTHESIS AND HERBICIDAL ACTIVITY OF CINNAMIC ACID AND RELATED COMPOUNDS

Miss Sujittra Deesamer

A Thesis Submitted in Partial Fulfillment of the Requirements
for the Degree of Master of Science in Chemistry

Department of Chemistry

Faculty of Science

Chulalongkorn University

Academic Year 2000

ISBN 947-347-270-3

Thesis Title	Synthesis and Herbicidal Activity of Cinnamic Acid and Related		
	Compounds		
By	Miss Sujittra Deesamer		
Field of Study	Chemistry		
Thesis Advisor Assistant Professor Warinthorn Chavasiri, Ph.D.			
Accepted 1	by the Faculty of Science, Chulalongkorn University in Partial		
Fulfillment of the F	Requirements for the Master's Degree.		
	Wand Pattle Dean of Faculty of Science		
	(Associate Professor Wanchai Phothiphichitr, Ph.D.)		
Thesis committee			
	Oldom Rotopal Chairman		
	(Professor Udom Kokpol, Ph.D.)		
	Warmhan Chwashi Thesis Advisor		
	(Assistant Professor Warinthorn Chavasiri, Ph.D.)		
	Poder S. Ls - Member		
	(Professor Padet Sidisunthorn, Ph.D.)		
	Siripon Zy Member		
	(Dr. Siriporn Zungsonthiporn, Ph.D.)		

สุจิตตรา ดีเสมอ : การสังเคราะห์และฤทธิ์กำจัดวัชพีชของกรดชินนามิกและสารประกอบที่
เกี่ยวข้อง (SYNTHESIS AND HERBICIDAL ACTIVITY OF CINNAMIC ACID
AND RELATED COMPOUNDS), อ. ที่ปรึกษา : ผส.ดร. วรินทร ชวศิริ; 133 หน้า;
ISBN 947-347-270-3.

สังเคราะห์สารในกลุ่มกรคซินนามิกสี่สิบเจ็ดชนิดโดยอาศัยปฏิกิริยาคอนเดนเซชันระหว่างกรด มาโลนิกกับแอลดีไฮด์ นำสารที่ได้ไปทดสอบฤทธิ์ยับยั้งการเจริญเติบโตของไมยราบยักษ์ (Mimosa pigra Linn.) และหญ้าข้าวนก (Echinochloa crus-galli Beauv.) พบว่ากรคซินนามิกแปดชนิคที่มีหมู่ แทนที่เป็น 3-ฟลูออโร, 3-คลอโร, 2,6-ใคคลอโร, 2-คลอโร-6-ฟลูออโร, 3-เมทอกซี, 4-เมทอกซี, 3,4-เมทิลลีนไคออกซี และ 3-ในโตร สามารถยับยั้งการเจริญเติบโตของไมยราบยักษ์ได้ดี และกรด ซินนามิกสิบเอ็คชนิดที่มีหมู่แทนที่เป็น 4-ฟลูออโร, 2-คลอโร, 4-คลอโร, 2-, 3- และ 4-โบรโม, 2,4-, 2,6- และ 3,4-ไคคลอโร, 4-เมทอกซี และ 4-เทอชีรีบิวทิล แสดงฤทธิ์การยับยั้งการเจริญเติบโตของ หญ้าข้าวนกได้ดี นอกจากนี้ ได้สังเคราะห์ซินนามามีคสิบสองชนิด, ซินนาเมดเอสเทอร์สิบห้าชนิด, เกลือโซเดียมซินนาเมดหกชนิด และเกลือแคลเซียมซินนาเมดสามชนิด แล้วนำไปทดสอบฤทธิ์การ ยับยั้งการเจริญเติบโตของไมยราบยักษ์ พบว่า เอ็น-2,4-ไคคลอโรฟีนิล 3,4-เมทิลลีนไดออกซีซินนามีค, 2.4-ใคคลอโรฟีนิล 3-ในโตรซินนาเมค, โซเคียม 3-ฟลออโรซินนาเมค, โซเคียม 3-คลอโรซินนาเมค, โซเดียม 3-เมทอกซีซินนาเมค, โซเดียม 3-ในโตรซินนาเมค, โซเดียม 3,4-เมทิลลีนใดออกซินนาเมค, แคลเซียม 3-ฟลออโรซินนาเมด และ แคลเซียม 3,4-เมทิลลีน ไดออกซีซินนาเมด แสดงฤทธิ์การยับยั้งที่ดี นอกจากนี้พบว่า กรด 3,4-เมทิลลีนไดออกซีซินนามิก และ 3-ในโตรซินนามิก ไม่แสดงฤทธิ์การยับยั้ง สารหลายชนิดแสดงฤทธิ์เทียบเคียงได้กับสารกำจัดวัชพืชที่ใช้ในทางการ การงอกของพืชทดสอบ เกษตรในปัจจุบัน สารดังกล่าวนี้มีศักยภาพที่จะพัฒนาเป็นสารกำจัดวัชพืชชนิดใหม่ต่อไป

ภาควิชา	เคมี	ลายมือชื่อนิสิตคากาก	คิเลนอ
สาขาวิชา	เคมี	ลายมือชื่ออาจารย์ที่ปรึกษา.	Johns Volts
ปีการศึกษา	2543		

4172492123 : MAJOR CHEMISTRY

KEY WORD: CINNAMIC ACID / HERBICIDAL

SUJITTRA DEESAMER: SYNTHESIS AND HERBICIDAL ACTIVITY
OF CINNAMIC ACID AND RELATED COMPOUNDS.
THESIS ADVISOR: ASSISTANT PROFESSOR WARINTHORN
CHAVASIRI, Ph.D. 133 pp. ISBN 947-347-270-3.

Forty-seven substituted trans-cinnamic acids were synthesized by condensation reaction between malonic acid and selected aldehydes. The inhibitory effect of these well-characterized compounds were tested with Mimosa pigra Linn. and Echinochloa crus-galli (L.) Beauv. The eight trans-cinnamic acids bearing, 3-F, 3-Cl, 2,6-diCl, 2-Cl-6-F, 3-OMe, 4-OMe, 3,4-OCH₂O- and 3-NO₂ showed high inhibitory effect on growth of M. pigra. While the eleven trans-cinnamic acids containing 4-F, 2-Cl, 4-Cl, 2-, 3- and 4-Br, 2,4-, 2,6- and 3,4-diCl, 4-OMe and 4-t-C₄H₉ showed high inhibitory effect on growth of E. crus-galli. In addition, twelve cinnamamides, fifteen cinnamate esters, six sodium cinnamate and three calcium cinnamate derivatives were prepared. These well-characterized compounds were tested for antigrowth activity of M. pigra. N-(2,4-dichlorophenyl)-3,4-methylenedioxycinnamamide, 2,4-dichlorophenyl 3-nitrocinnamate, sodium 3-fluorocinnamate, sodium 3-chlorocinnamate, sodium 3-methoxycinnamate, sodium 3-nitrocinnamate, sodium 3,4-methylenedioxycinnamate, calcium 3-fluorocinnamate and calcium 3,4-methylenedioxycinnamate exhibited high activities. Besides, 3,4-methylenedioxycinnamic and 3-nitrocinnamic acids did not show germination inhibition activity against selected weeds. Several compounds displayed the activity comparable to various commercial herbicides in terms of growth inhibition activity test. Some of them revealed promising tendency for the development as new herbicides.

DepartmentChemistryStudent's signatu	ire Suiittra	Deesamer	
Field of studyChemistryAdvisor's signatu	ne Naturan	enovoohi	
Academic year2000			

ACKNOWLEDGEMENT

Firstly, the author would like to express her sincere gratitude to Assistant Professor Dr. Warinthorn Chavasiri, her major advisor, for his kindly helpful suggestions, valuable assistance and encouraging throughout the entire period of this research. Sincere thanks are also extended to Professor Dr. Udom Kokpol, Professor Dr. Padet Sidisunthorn and Dr. Siriporn Zungsonthiporn, serving as the chairman and members of her theses committee, respectively, for their valuable comments and suggestion.

Gratitude is also expressed to the staff of Natural Products Research Unit, Department of Chemistry, Chulalongkorn University for their helpful discussion and support. In addition, the author greatly appreciated the Faculty of Science, Chulalongkorn University for granting her a teaching assistantship during 1998-2000 and partial financial support of this research work.

The author also greatest thanks to her bioassay supports: Dr. Siriporn Zungsonthiporn (Weed Science Sub-division, Botany and Weed Science Division, Department of Agriculture, Ministry of Agriculture and Cooperatives) for providing and giving an advice on bioassay of herbicidal activity.

Special thanks are extended to Department of Chemistry, Chulalongkorn University for supporting materials, chemicals and everything else.

Moreover, the author would like to express her sincere gratitude to her parents and family members for love, understanding, encouragement, limitless sacrifice and advice throughout the entire study. Ultimately, she never forgets to profoundly thank someone special who is a nice person for continuously cheering up. Without them, the author would never have been able to achieve this goal.

CONTENTS

	Pages
Abstract in Thai	iv
Abstract in English	v
Acknowledgement	vi
List of Figures	x
List of Tables	xiii
List of Abbreviations	xiv
CHAPTER I : INTRODUCTION.	1
1.1 Literature Review	5
1.2 Synthesis of trans-Cinnamic acid	8
1.3 Knowledge about Studied Weeds	9
1.3.1 Mimosa pigra Linn	9
1.3.2 Echinochloa crus-galli (L.) Beauv	10
1.4 Goal of Research	12
CHAPTER II : EXPERIMENTAL	13
2.1 Instruments and Equipment	13
2.2 Chemicals.	13
2.3 Synthesis of Substituted trans-Cinnamic Acids	14
2.4 Synthesis of trans-Cinnamic Acid Derivatives	27
2.4.1 Synthesis of Cinnamamides	27
2.4.2 Synthesis of Cinnamate Esters	32
2.4.3 Preparation of Sodium Cinnamate Derivatives	37
2.4.4 Preparation of Calcium Cinnamate Derivatives	38
2.5 Bioassay on Herbicidal Activity	39
2.5.1 General Procedure for Weed Growth Inhibition Test	39
2.5.2 General Procedure for Weed Germination Inhibition Test	40

	Pages
2.6 Herbicidal Activity of Commercially Available Herbicides	41
CHAPTER III : RESULTS AND DISCUSSION	42
3.1 Synthesis of Substituted trans-Cinnamic Acids	42
3.2 Spectroscopic Data of Substituted trans-Cinnamic Acids	45
3.3 Synthesis and Spectroscopic Data of trans-Cinnamic Acid Derivatives.	. 50
3.4 Herbicidal Potential	. 51
3.4.1 Weed Growth Inhibition against Mimosa pigra Linn	51
3.4.2 Weed Growth Inhibition against Echinochloa crus-galli (L.)	
Beauv	70
3.5 Herbicidal Potential of trans-Cinnamic Acid Derivatives	. 82
3.5.1 Cinnamamides (D1-D12)	82
3.5.2 Cinnamate Esters (E1-E15)	86
3.5.3 Sodium Cinnamate Derivatives (S1-S6)	91
3.5.4 Calcium Cinnamate Derivatives (CS1-CS3)	95
3.6 Weed Germination Inhibition	99
3.7 Commercially Available Herbicides Against Mimosa pigra Linn	102
CHAPTER IV : CONCLUSION.	104
REFERENCES	108
APPENDICES	112
Appendices A	112
A.1 Synthesis of Starting Materials	112
A.1.1 Synthesis of 4-Alkoxybenzaldehyde	112
A.1.2 Synthesis of 4-Alkoxy-3-methoxybenzaldehyde	114
Appendices B	116
Appendices C	124
C.1 Herbicidal Activity of Substituted trans-Cinnamic Acids	124
C.2 Herbicidal Activity of Substituted trans-Cinnamic Acids	
Derivatives	128
C.2.1 Cinnamamides (D1-D12)	128
C.2.2 Cinnamate Esters (E1-E15)	129

		Pages
	C.2.3 Sodium Cinnamate Derivatives (S1-S6)	130
	C.2.4 Calcium Cinnamate Derivatives (CS1-CS3)	130
	C.3 Germination and Root Growth Inhibition	131
	C.4 Herbicidal Activity of Commercial Available Herbicides	132
VITA		133

List of Figures

Figs		Pages
1.1	Mimosa pigra Linn. (Giant mimosa)	10
1.2	Echinochloa crus-galli (L.) Beauv. (Barnyard grass)	11
2.1	Structures of synthesized substituted trans-cinnamic acids	14
2.2	Structures of synthesized cinnamamides	27
2.3	Structures of synthesized cinnamate esters	32
2.4	Structures of sodium cinnamate derivatives	37
2.5	Structures of calcium cinnamates derivatives	38
3.1	The area of Mimosa pigra distribution.	51
3.2	Inhibitory effect of substituted trans-cinnamic acids on root and shoot	
	growth of <i>M. pigra</i> (%)	52
3.3	Inhibitory effect of substituents at various positions on a benzene ring	
	of trans-cinnamic acids on root growth of M. pigra at 300 ppm (%)	64
3.4	Inhibitory effect of various substituents on a benzene ring of trans-	
	cinnamic acids on root growth of M. pigra at 300 ppm (%)	65
3.5	Inhibitory effect of one, two and three methoxy group on a benzene ring	
	of trans-cinnamic acids on root growth of M. pigra at 300 ppm (%)	69
3.6	Inhibitory effect of substituted trans-cinnamic acids on root growth of	
	M. pigra at 300 ppm (%)	69
3.7	Inhibitory effect of substituted trans-cinnamic acids on root growth	
	of E. crus-galli (%)	70
3.8	Inhibitory effect of substituted trans-cinnamic acids on root growth of	
	E. crus-galli at 300 ppm (%)	81
3.9	Inhibitory effect of cinnamamides on root and shoot growth of M. pigra (%).	82
3.10	Inhibitory effect of cinnamamides and their parent compounds on root	
	and shoot growth of M. pigra (%)	85

Figs		Pages
3.11	Inhibitory effect of cinnamate esters on root and shoot growth of	
	M. pigra (%)	86
3.12	Inhibitory effect of cinnamate ester and their parent compounds on root	
	growth of M. pigra (%)	90
3.13	Inhibitory effect of sodium cinnamate derivatives on root and shoot growth	
	of <i>M. pigra</i> (%)	91
3.14	Inhibitory effect of the various 3-position substituent on a benzene ring of	
	sodium cinnamate derivatives on root growth of M. pigra (%)	92
3.15	Inhibitory effect of sodium cinnamate derivatives on root growth of	
	M. pigra at 300 ppm (%)	93
3.16	Inhibitory effect of sodium cinnamate derivatives and their parent	
	compounds on root growth of <i>M. pigra</i> (%)	94
3.17	Inhibitory effect of sodium cinnamate derivatives and their parent	
	compounds on root growth of <i>M. pigra</i> at 300 ppm (%)	95
3.18	Inhibitory effect of calcium cinnamate derivatives on root and shoot	
	growth of <i>M. pigra</i> (%)	95
3.19	Inhibitory effect of calcium cinnamate derivatives and their parent	
	compounds on root growth of M . $pigra$ (%)	96
3.20	Inhibitory effect of sodium and calcium cinnamate derivatives and	
	their parent compounds on root growth of M. pigra at 300 ppm (%)	97
3.21	Inhibitory effect of 3,4-methylenedioxycinnamic acid and its derivative	
	on root growth of <i>M. pigra</i> at 300 ppm (%)	98
3.22	Inhibitory effect of 3-nitrocinnamic acid and its derivative on root	
	growth of <i>M. pigra</i> at 300 ppm (%)	99
3.23	Germination and root growth inhibition of 3,4-methylenedioxycinnamic	
	(C33) and 3-nitrocinnamic (C35) acids against M. pigra	100
3.24	Germination and root growth inhibition of 3,4-methylenedioxycinnamic	
	(C33) and 3-nitrocinnamic (C35) acids against E. crus-galli	100
3.25	Germination and root growth inhibition of 3,4-methylenedioxycinnamic	
	(C33) and 3-nitrocinnamic (C35) acids against E. geniculata	101

Figs		Pages
3.26	Germination and root growth inhibition of 3,4-methylenedioxycinnamic	
	(C33) and 3-nitrocinnamic (C35) acids against A. americana	101
3.27	Inhibitory effect of eight potent trans-cinnamic acids (C1, C4, C10, C12,	
	C14, C15, C33 and C35) and commercially herbicides (H1 and H2) on	
	root growth of M. pigra (%)	102
3.28	Inhibitory effect of eight potent trans-cinnamic acids and commercially	
	herbicides (H1 and H2) on root growth of M. pigra at 300 ppm (%)	103
A . 1	Structures of synthesized 4-alkoxybenzaldehydes	112
A.2	Structures of synthesized 4-alkoxy-3-methoxybenzaldehyde	114
B.1	The FT-IR spectrum of C24	116
B.2	The ¹ H-NMR spectrum of C24	117
B.3	The ¹³ C-NMR spectrum of C24	118
B .4	The mass spectrum of C24.	119
B.5	The FT-IR spectrum of C25.	120
B .6	The ¹ H-NMR spectrum of C25.	121
B.7	The ¹³ C-NMR spectrum of C25	122
B.8	The mass spectrum of C25	123

0.

List of Tables

Tab	les	Pages
3.1	Physical properties and % yield of synthesized substituted <i>trans</i> -cinnamic acids	43
3.2	FT-IR absorption band assignment of new synthetic trans-cinnamic acids	
	(C24 and C25)	45
3.3	¹ H-NMR spectral assignments of new synthetic trans-cinnamic acids	
	(C24 and C25)	47
3.4	¹³ C-NMR spectral assignments of new synthetic trans-cinnamic acids	
	(C24 and C25)	49
3.5	Inhibitory effect of halocinnamic acids at 300 ppm	61
C .1	The results of weed growth inhibition of substituted trans-cinnamic acids	
	against M. pigra.	124
C .2	The results of weed growth inhibition of substituted trans-cinnamic acids	
	against E. crus-galli	126
C.3	The results of weed growth inhibition of cinnamamides against M. pigra	128
C.4	The results of weed growth inhibition of cinnamate esters against M. pigra	129
C.5	The results of weed growth inhibition of sodium cinnamate derivatives	
	against M. pigra	130
C .6	The results of weed growth inhibition of calcium cinnamate derivatives	
	against M. pigra.	130
C .7	The results of germination and root growth inhibition of 3,4-methylene-	
	dioxycinnamic acid (C33) against various weeds	131
C.8	The results of germination and root growth inhibition of 3-nitrocinnamic	
	acid (C35) against various weeds	131
C .9	The results of weed growth inhibition of commercially available herbicides	
	against M. pigra.	132

List of Abbreviations

br	broad	mm	millimeter
°C	degree Celsius	m.p.	melting point
cm ⁻¹	unit of wavenumber	MS	mass spectroscopy
Cpd	compound	m/z	mass per charge
d	doublet (NMR)	NMR	nuclear magnetic resonance
dd	doublet of doublet (NMR)	ppm	part per million
DMSO	dimethylsulfoxide	q	quartet (NMR)
g	gram (s)	rel. int.	relative intensity
h	heptet (NMR)	$R_{\rm f}$	retardation factor
Hz	hertz	S	strong (IR)
IR	infrared	S	singlet (NMR)
J	coupling constant	t	triplet (NMR)
lit	literature	w	weak (IR)
m	multiplet (NMR)	wt	weight
m	medium (IR)	δ	chemical shift
mL	milliliter (s)	%	percent