Perceived Social Support: Its Genetic and Environmental Etiology and Association with Depression

William Luya Coventry

B. Ag. Eco. - University of Sydney
Grad. Dip. Soc. Sci. - University of New England
Hon. Arts (Psychology) – University of Queensland

A thesis submitted for the degree of Doctor of Philosophy of the University of New England

February 2007

Acknowledgments

This PhD has been a wonderful experience, and I have learnt a great deal about so many different aspects of research. Firstly, I would like to thank all the fellow students and staff I worked alongside during this PhD. This list is vast, so I shall not name you (no Mark, you don't get a special mention). It includes those I commenced my candidature with at UNE in Armidale. Since then, I have enjoyed the wonderful company of many at both QIMR and UNE; you have been a great source of support.... through these stressful times! I have enjoyed meeting you all. Further, I have academically learnt so much by interacting with each of you and Matt Keller in particular, it has been an absolute pleasure working with you in this regard. This has made the PhD process hugely rewarding. Sammie Lane, our discussions about writing were fun.

My family also, you have been a valuable source of support, allowing me to maintain an obsession whilst feigning positive regard! Dad, particularly, thankyou for minding the fort whilst I have been away, I am exceedingly grateful. I am not sure this completely conveys my gratitude: I will make it up to you in due course.

To my supervisors, Brian and Nick; I was extremely fortunate to have had you both as supervisors. You are both exceptional and sensible, with fantastic judgment. It has been enjoyable working with each of you. You have both been very good to me over these years, and I am grateful for all you have done.

To all the twins who provided the data, and all the staff at QIMR who were involved in all aspects of preparing the data I used throughout this thesis. From the interviewers, to those entering the data, to the IT and administrative staff involved in compiling and organizing the database, and to the genotyping lab for their incredible amount of work on the serotonin transporter genotype, thankyou.

Abstract

Almost all research on social support has ignored the genetic contribution to individual differences. The current thesis used a sample of 8,000 male and female adult twin pairs. After refining the perceived social support phenotype, this research explored its genetic and environmental etiology using the classical twin design (CTD). In attending to the biases of this design, the thesis compared CTD estimates for a range of phenotypes against estimates from the more comprehensive extended twin-family design. Although estimates of additive (A) and non-additive (NA) genetic variance were over- and under-estimated respectively, common environment (C) and broadsense genetic (G; A + NA) estimates were reasonably accurate with the latter only slightly overestimated. With these biases considered, C and G each explained approximately 1/5th of the variance in perceived support; the balance of variation was unique environment, although approximately 1/3rd of this may be measurement error. The additive genetic contribution differed in males and females.

The next section explored the association between perceived support, stress and depression. A comparison between phenotypic associations and those from a discordant monozygotic (MZ) design showed part of the associations reported in the literature are due to familial effects (G or C). After removing these, stress was still associated with depression while perceived support was not, except as a buffer against depression later on in males who experienced multiple stressors.

The final section extended the association between perceived support, stress and depression to include the serotonin transporter polymorphism (5HTTLPR). A review of the literature on stress × 5HTTLPR in predicting psychopathology suggested publication bias and raised questions about some of the interactions reported. The current analyses improved on the previous research by using 5HTTLPR (S/L) redefined by the SNP rs25531 (a/g) within the L allele. There was no interaction between the polymorphism and (a) stress or (b) both stress and social support in predicting depression. Future research exploring the associations studied in this thesis should use the full panel design in absence of behavior genetic techniques. Further, research on social support that uses quasi-experimental designs of traumatised populations would be valuable.

Certification

I certify that the substance of this thesis has not already been submitted for any degree and is not currently being submitted for any other degree or qualification.

I certify that any help received in preparing this thesis, and all sources used, have been acknowledged in this thesis.



TABLE OF CONTENTS

	Introduction	
1.1 Backgrour	nd	
1.1.1 A brief	history of social support	
1.1.1 A brief history of social support		
1.1.3 Depres	ssion	
1.1.4 Social	ssionsupport – perceived versus received nvironment and individual difference perspectives of perceived support	
1.1.5 The er	ivironment and individual difference perspectives of perceived support	6
1.1.6 Percei	ved support in a genetically sensitive design	
1.2 Outline of	the thesis	8
1.2.1 Chapte	er 2	8
1.2.2 Chapte	er 3	8
1.2.3 Chapte	er 4er 5 (and 6)	8
1.2.4 Chapte	er 5 (and 6)	{
1.2.5 Chapte	er 6	9
1.3 Data oven	view	10
Chapter Two		
ar	nd sex differences	13
Coventry et al. (2	2004)	13
Frratum		21
Statement of orig	ginality and of the contribution by others	27
Chantar Three	Constituted and any iron mantal influences on parasited as	oio!
	Genetic and environmental influences on perceived so	
SL	upport; differences by sex and relationship	
A b =4 =4		0.4
ADSTract		
4 14		J
1. Introduction		
0.85.41.1		31
		31
2.1.1 Partici	pants	31 36
2.1.1 Partici _l 2.1.2 Instrun	pantsnents	36 36 36
2.1.1 Partici _l 2.1.2 Instrun 2.1.3 Data c	pantsnents	36 36 36 37
2.1.1 Partici _l 2.1.2 Instrun 2.1.3 Data c	pantsnents	36 36 36 37
2.1.1 Partici _l 2.1.2 Instrun 2.1.3 Data c 2.1.4 Zygosi 2.1.5 Tests c	pantsnents	36 36 37 38 40 40
2.1.1 Partici 2.1.2 Instrun 2.1.3 Data c 2.1.4 Zygosi 2.1.5 Tests c 2.1.6 Statisti	pants	3636363638384040
2.1.1 Partici 2.1.2 Instrun 2.1.3 Data c 2.1.4 Zygosi 2.1.5 Tests c 2.1.6 Statisti	pants	3636363638404041
2.1.1 Participole 2.1.2 Instrum 2.1.3 Data construction 2.1.4 Zygosi 2.1.5 Tests construction 2.1.6 Statistic 3. Results	pants	36363637383838404041
2.1.1 Particip 2.1.2 Instrum 2.1.3 Data c 2.1.4 Zygosi 2.1.5 Tests c 2.1.6 Statisti 3. Results	pants	36363636373840404147
2.1.1 Particip 2.1.2 Instrum 2.1.3 Data c 2.1.4 Zygosi 2.1.5 Tests c 2.1.6 Statisti 3. Results	pants	3636363738404041474747
2.1.1 Particip 2.1.2 Instrum 2.1.3 Data c 2.1.4 Zygosi 2.1.5 Tests c 2.1.6 Statisti 3. Results	pants	3636363638404041474750
2.1.1 Participe 2.1.2 Instrum 2.1.3 Data con 2.1.4 Zygosi 2.1.5 Tests con 2.1.6 Statisti 2.1.6 Statisti 3. Results	pants	36363636363738404041474750
2.1.1 Particip 2.1.2 Instrum 2.1.3 Data of 2.1.4 Zygosi 2.1.5 Tests of 2.1.6 Statisti 3. Results	pants	31363636373840414147475057
2.1.1 Particip 2.1.2 Instrum 2.1.3 Data of 2.1.4 Zygosi 2.1.5 Tests of 2.1.6 Statisti 3. Results	pants	31363636373840414147475057

		IV
Univariate sex limitation script		65
Cholesky scr	ript of the KPSS sub-scales, females	68
	ginality and of the contribution by others	
	Parameter bias in the Classical Twin Designerceived social support	
1. Overview		76
2. Summary of K	Celler et al. (2005)	76
2.1.1 Parame	eter bias in the classical twin design (CTD)	76
2.1.2 Attendi	ing to parameter bias in the CTD	76
2.1.3 The rar	ing to parameter bias in the CTD nge of unbiased estimates for perceived social support	77
3. Coventry et al. (2005)		78
•	ginality and of the contribution by others	
Chapter Five bu	A prospective co-twin control design identiful infering against depression in males but not femo	
Abstract		92
1. Introduction		93
	of methodologies used in the literature	
	iation between social support and depression	
1.2.1 Longitu	udinal studies.	96
1.2.2 Behavio	or genetic studies.	97
1.2.3 Sex diff	ferences	98
	iation detween stress and depression	
	iation between social support, stress and depression	
1.4.1 Longitu	udinal studiesor genetic studies	99
1.4.2 Behavio	or genetic studies.	100
1.5 Summary		103
	's	
2.2.1 Kessler	r Perceived Social Support (KPSS).	106
2.2.2 Stressf	ful Life Events (SLE).	107
	sion.	
	oric correlations	
2.3.2 Filenot	typic analysesdant analyses	113
	ective design	
2.3.5 Statistic	cs used	115
	tions for multiple testing	115
	ring the phenotypic and discordant analyses	116
2.3.8 Alterna	tive analytic techniques	116
3.1 Polychoric	correlations	117
3.2 Phenotypic	analyses	120

	V
3.2.1 Sex	120
3.2.2 Age	
3.2.2 Age	121
3.2.4 SLE	121
3.2.5 The SLE × KPSS interaction	121
3.2.6 Addressing skewness	122
3.2.6 Addressing skewness 3.2.7 Addressing non-independence between twin pairs	127
3.2.8 Summary	127
3.3 Analyses of MZ pairs discordant for depression	128
3.3.1 Fit of the full regression models	128
3.3.2 KPSS	128
3.3.3 SLE	129
3.3.4 The SLE × KPSS interaction	132
4. Discussion	133
4.1.1 Limitations.	
5. Conclusion	137
Acknowledgments	120
References	140
Statement of originality and of the contribution by others	146
•	
review and an analysis with 5HTTLPR re-defined by rs2555	31 and
	31 and
review and an analysis with 5HTTLPR re-defined by rs2553	31 and 147
review and an analysis with 5HTTLPR re-defined by rs2555 perceived social support as a moderator	31 and 147 149
review and an analysis with 5HTTLPR re-defined by rs2555 perceived social support as a moderator	31 and 147 149 150
review and an analysis with 5HTTLPR re-defined by rs2555 perceived social support as a moderator	31 and147149150
review and an analysis with 5HTTLPR re-defined by rs2555 perceived social support as a moderator	31 and147149150151
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and147149150151
review and an analysis with 5HTTLPR re-defined by rs2555 perceived social support as a moderator	31 and147149150151151151
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and147
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and147
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and147150
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255; perceived social support as a moderator	31 and
Abstract 1. Introduction 2. Literature review 2.1 Overview of studies exploring the interaction between stressful life events and 5HTTLPR genotypes on psychopathology. 2.1.1 Interactions in a direction consistent with Caspi et al. (2003) 2.1.2 Findings inconsistent with those of Caspi et al. (2003) 2.1.3 Papers not exploring an interaction 2.2 Issues common across the papers reviewed 2.2.1 Additive v's dominant genotypes 2.2.2 Graphical considerations 2.2.3 Publication bias	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and
review and an analysis with 5HTTLPR re-defined by rs255: perceived social support as a moderator	31 and

	VI
3.1.7 Genotype totals	174
3.2 Measures	
3.2.1 Genotype	175
3.2.2 Kessler Perceived Social Support (KPSS) a	nd Stressful Life Events (SLE) 175
3.2.3 Depression	176
3.3 Analysis	
	177
3.3.2 Ordinal regressions	178
2 2 2 Chatiatian	178
3.3.4 Additive or dominant genotypes	179
3.3.5 Corrections for multiple testing	179
3.3.6 Addressing the interdependence between tw	vin pairs 181
4. Results	182
	182
4.1.2 Main effects	183
4.1.3 Two-way interactions	184
4.1.4 Three-way interactions	187
5. Discussion	
5.1.1 Additive v dominant modes of action	
5.1.2 Does redefining 5HTTLPR by rs25531 make	e a difference? 189
5.1.3 Interactions between SI F and the genotype	s190
5 1 4 KPSS as a moderator	191
5.2 Limitations	191
6. Conclusion	
Acknowledgments	
•	
References	195
Statement of originality and of the contribution	by others198
Chapter Seven Discussion and conclusion	ons 199
1. Summaries and future directions	
1.1.1 Chapter 2	
1.1.2 Chapter 3	200
1.1.3 Chapter 4	
1.1.4 Chapter 5	203
1.1.5 Chapter 6	
2. Novelty	208
3. Limitations	209
4. Conclusions	210
4.1.1 The bias in the CTD estimates	210
4.1.1 The bias in the CTD estimates4.1.2 Genetic and environmental etiology of socia	support 210
4.1.3 The association between social support and	depression 210

References 212

Appendix 228