

# AN EXAMINATION OF THE NATURE OF SLEEP FRAGMENTATION IN CHILDREN WITH UPPER AIRWAY OBSTRUCTION

---

**Scott Wade Coussens**

**Bachelor of Science, Bachelor of Science (Honours)**

**A thesis submitted for the degree of Doctor of Philosophy in Physiology by a mixed  
portfolio of publications, submitted manuscripts, manuscripts and traditional chapters**

**Discipline of Physiology**

**Faculty of Health Sciences**

**University of Adelaide**

**September 2015**



**THE UNIVERSITY**  
*of* **ADELAIDE**



# TABLE OF CONTENTS

---

TABLE OF CONTENTS .....	i
TABLE OF CONTENTS (Tables) .....	ix
TABLE OF CONTENTS (Figures) .....	xi
<hr/>	
<i>i. Abstract</i> .....	13
<i>ii. Signed Declaration</i> .....	15
<i>iii. Acknowledgements</i> .....	15
<i>iv. Publications</i> .....	18
<i>v. Style and Referencing Format of Jointly Authored Papers, Documents and Unpublished Manuscripts</i> .....	20
<i>vi. Abbreviations</i> .....	21
<b>CHAPTER 1</b> .....	<b>24</b>
<hr/>	
<i>Introduction to Sleep and Sleep Disorders in Children</i> .....	24
1.1. SLEEP .....	25
<b>Sleep Architecture</b> .....	25
<b>The Transition to Sleep</b> .....	25
<b>NREM Sleep Stage 1</b> .....	26
<b>NREM Sleep Stage 2</b> .....	27
<b>Slow Wave Sleep</b> .....	27
<b>Rapid Eye Movement Sleep (REM Sleep)</b> .....	27
1.2. Polysomnography.....	29
1.3. Paediatric Sleep Analysis .....	31
1.4. SLEEP Function.....	31
1.5. Sleep DISORDERS.....	32
<b>Upper Airway Obstruction (UAO) in children</b> .....	32
<b>Prevalence of Upper Airway Obstruction in children</b> .....	33
<b>The Incomplete Story of Hypoxia as the Pathophysiological Mechanism of Upper Airway Obstruction</b> .....	35
1.6. References.....	37
<b>CHAPTER 2</b> .....	<b>43</b>
<hr/>	

<i>A Review of Clinical Sleep Fragmentation Indices in Children with Upper Airway Obstruction</i> .....	43
2.1. Introduction.....	44
2.1.1. Sleep Fragmentation IN CHildren .....	44
<b>Limitations on the use of sleep fragmentation measures in children</b> .....	49
2.2. AIMS OF THE REVIEW .....	51
2.3. Sleep Fragmentation Indices Derived from Measures of Cortical Neural Activation ...	52
2.3.1. EEG Synchrony.....	52
2.3.2. Awakenings .....	52
<b>Awakenings are easy to measure</b> .....	53
<b>Awakenings may underestimate fragmentation</b> .....	53
<b>Awakenings Number and Awakenings Index</b> .....	53
2.3.3. ASDA Standard Arousals.....	54
<b>Arousal Length</b> .....	55
<b>Total Arousals</b> .....	55
<b>Spontaneous Arousals</b> .....	56
<b>Respiratory Arousals</b> .....	56
2.3.4. The Sleep Pressure Score .....	57
2.3.5. Microarousals.....	58
<b>Inter-Scorer Variability</b> .....	59
<b>Score Reliability</b> .....	59
<b>Time and Cost</b> .....	59
<b>Microarousal Index</b> .....	60
2.3.6. Spectral Analysis .....	60
2.3.7. Sleep Continuity .....	62
<b>Survival Curve Analysis</b> .....	62
2.3.8. EEG and Snoring.....	63
2.4. Sleep Fragmentation Indices Derived from Measures of Sub-Cortical Activation .....	64
2.4.1. Movement.....	64
<b>Standard Movement Measures</b> .....	64
<b>Movement Arousals</b> .....	65
2.4.2. Heart Rate changes .....	65
2.4.3. Blood Pressure changes .....	66
2.4.4. Pulse Transit Time changes.....	67
2.5. DISCUSSION .....	69
2.5.1. Summary .....	69

2.5.2. Future directions for performance and analysis of PSGs in children .....	69
<b>Standard Scoring Methods .....</b>	<b>69</b>
<b>Standardized Daytime Functional Comparisons .....</b>	<b>70</b>
Total Sleep Time.....	71
2.5.3. future directions for investigating underlying mechanisms of sleep fragmentation in children with uao.....	72
<b>Autonomic Measures.....</b>	<b>72</b>
<b>EEG Measures.....</b>	<b>72</b>
2.5.4. AIMS of this study.....	73
<b>Identification of indirect sleep fragmentation measures .....</b>	<b>74</b>
<b>Adaptation of Adult Indices.....</b>	<b>74</b>
2.6. References.....	76
<b>CHAPTER 3.....</b>	<b>88</b>
<b><i>General Methods.....</i></b>	<b>88</b>
<hr/>	
3.1. Subjects .....	89
3.2. Overnight Polysomnography .....	90
3.3. Data Analysis.....	92
3.3.1. PSG Data Analysis.....	92
3.4. Standard Research Scoring Rules for Paediatric Sleep Studies .....	93
3.4.1. Sleep Stage Scoring .....	93
<b>NREM Stage 1 – (Set Rules) .....</b>	<b>93</b>
<b>NREM Stage 1 – (Supporting Evidence).....</b>	<b>93</b>
<b>NREM Stage 2 – (Set Rules) .....</b>	<b>94</b>
<b>NREM Stage 2 – (Supporting Evidence).....</b>	<b>94</b>
NREM Stage 3 – (Set Rules).....	94
<b>NREM Stage 3 – (Supporting Evidence).....</b>	<b>94</b>
<b>NREM Stage 4 – (Set Rules).....</b>	<b>95</b>
<b>NREM Stage 4 – (Supporting Evidence).....</b>	<b>95</b>
<b>REM – (Set Rules) .....</b>	<b>95</b>
<b>REM– (Supporting Evidence) .....</b>	<b>95</b>
<b>Movement.....</b>	<b>96</b>
<b>Awake – (Set Rules) .....</b>	<b>96</b>
<b>Awake – (Supporting Evidence) .....</b>	<b>97</b>
<b>Unknown.....</b>	<b>97</b>

<b>Artefact</b> .....	97
<b>Dealing with “Long Arousals”</b> .....	97
3.4.2. Arousal Scoring.....	97
<b>Movement Time</b> .....	97
<b>Cortical Spontaneous Arousal (Spontaneous)</b> .....	98
<b>Sub-Cortical Spontaneous Arousal (Sub-Cortical)</b> .....	98
<b>Respiratory Arousal (Respiratory)</b> .....	99
<b>Periodic Leg Movement Arousal (PLMA)</b> .....	99
<b>Technician-Caused Arousal (Scored as Resp. Event – TECH)</b> .....	100
<b>Sub-Cortical Respiratory Arousal (SCRA)</b> .....	100
<b>Spindles in Arousals</b> .....	101
3.4.3. Respiratory Event Scoring .....	101
<b>Central Apnoea – (Set Rules)</b> .....	101
<b>Central Hypopnea – (Set Rules)</b> .....	102
<b>Central Hypopnea – (Supporting Evidence)</b> .....	103
<b>Obstructive Apnoea – (Set Rules)</b> .....	103
Obstructive Apnoea – (Supporting Evidence) .....	104
Obstructive Hypopnea – (Set Rules) .....	104
<b>Obstructive Hypopnea – (Supporting Evidence)</b> .....	105
<b>Mixed Apnoea – (Set Rules)</b> .....	105
<b>Inspiratory Flow Reduction Event (IFRE) – (Set Rules)</b> .....	106
<b>Post Movement Central Apnoea – (Set Rules)</b> .....	107
<b>Respiratory Paradox – (Set Rules)</b> .....	108
<b>Post Sigh Central Apnoea – (Set Rules)</b> .....	108
3.4.4. Periodic Leg Movement Scoring .....	109
3.4.5. Signal Artefact Scoring.....	110
<b>TcCO<sub>2</sub> Artefact</b> .....	110
<b>SpO<sub>2</sub> Artefact</b> .....	110
3.5. References.....	111
3.6. Appendix 1 .....	114
3.6.1. Neurocognitive assessment.....	115
Stanford Binet Intelligence Scale .....	115
A Developmental NEuroPSYchological Assessment.....	116
3.6.2. Statistical Analysis.....	117
3.7. References.....	119

<b>CHAPTER 4.....</b>	<b>122</b>
<hr/>	
<i>Sleep Spindle Activity and Cognitive Performance in Healthy Children.....</i>	122
4.1. ABSTRACT.....	127
4.2. Introduction.....	128
4.3. Method.....	130
Design.....	132
Materials.....	132
Statistical Analysis.....	135
4.4. Results.....	136
4.5. Discussion.....	143
4.6. References.....	148
<b>CHAPTER 5.....</b>	<b>153</b>
<hr/>	
<i>EEG Changes Accompanying Spontaneous Arousals during Sleep Correlate with Neurocognitive Performance in Children with Upper Airway Obstruction.....</i>	153
5.1. Introduction.....	159
5.2. Methods.....	163
5.2.1. Subjects & Procedure.....	163
5.2.2. EEG Signal Processing.....	163
5.2.3. Neurocognitive testing.....	165
5.2.4. Statistics.....	166
5.3. RESULTS.....	167
5.3.1. Subjects.....	167
5.3.2. Polysomnography.....	167
5.3.3. neuropsychological Testing.....	168
5.3.4. Spontaneous Arousal Duration.....	173
5.3.5. Mean Spectral Power associated with Spontaneous Arousals.....	173
5.3.6. relative Change in Spectral EEG Power with Spontaneous Arousal Onset.....	180
5.3.7. Correlations between Neuropsychological performance and Spectral Power.....	182
5.3.8. Correlations between neurocognitive performance and EEG power with spontaneous arousals for the Control group.....	182
5.3.9. Correlations between neurocognitive performance and EEG power with spontaneous arousals for the Primary Snoring group.....	183
5.3.10. Correlations between neurocognitive performance and EEG power with spontaneous arousals for the OSAS group.....	184
5.4. DISCUSSION.....	185

5.4.1. Spontaneous arousal alteration in children with UAO as measured by common polysomnographic metrics.....	185
5.4.2. Spontaneous arousal alteration in children with UAO as determined by spectral analysis.....	186
5.4.3. Spontaneous arousal alteration in children with UAO and neurocognitive performance .....	188
5.5. Conclusion.....	191
5.6. References.....	199
<b>CHAPTER 6.....</b>	<b>204</b>

***Movement in Sleep*..... 204**

---

Acknowledgements.....	208
Disclosure statement .....	209
6.1. ABSTRACT.....	210
6.2. Introduction.....	212
6.3. Methods.....	215
6.3.1. Participants .....	215
6.3.2. Overnight Polysomnography .....	216
6.3.3. Movement Measures .....	218
6.3.4. Measures of Sleep Continuity .....	219
6.3.5. Survival Curve Analysis.....	219
6.3.6. Statistics.....	220
6.4. RESULTS.....	221
6.4.1. Subjects .....	221
6.4.2. Polysomnography.....	221
6.4.3. Movement Measures .....	222
6.4.4. Survival Curve Analysis.....	223
6.4.5. Stage 1 NREM sleep and wake as markers of sleep fragmentation .....	223
6.4.6. Movement Time as a marker of sleep fragmentation.....	223
6.4.7. Movement events as a marker of sleep fragmentation.....	224
6.4.8. Sleep fragmentation and AHI.....	225
6.5. DISCUSSION.....	226
6.6. Conclusion.....	230
6.7. Tables and Figures.....	231
6.8. References.....	242
<b>CHAPTER 7.....</b>	<b>250</b>

---

*Towards a Composite Index of Upper Airway Obstruction Severity in Children* ..... 250

7.1.	Abstract.....	255
7.2.	INTRODUCTION.....	256
	7.2.1. Limitations of Sleep Fragmentation as an explanantion of Neurocognitive and Behavioural Deficits in children with UAO .....	256
	7.2.2. Indices of upper airway obstruction Disease Severity in children .....	257
	7.2.3. Composite Indices .....	257
	The Physical Severity Index.....	258
	The Functional Severity Index.....	259
	The Polysomnographic Severity Index .....	260
	Measures of Hypoxemia .....	260
	Measures of Sleep Fragmentation.....	260
	Respiratory Measures .....	261
	7.2.4. outcome measures of disease burden.....	262
	7.2.5. Hypotheses.....	262
7.3.	Methods.....	264
	7.3.1. Subjects.....	264
	7.3.2. Overnight Polysomnography.....	265
	7.3.3. outcome measures of disease burden.....	267
	7.3.4. Movement Analysis.....	267
	7.3.5. Polysomnographic severity index – The TOTAL Sleep disturbance index .....	268
	7.3.6. Functional Severity index with Sleep Continuity Analysis.....	269
	7.3.7. Physical severity index.....	271
	7.3.8. composite Upper (airway obstruction) disease severity index (for) children (CUDSIC).....	271
	7.3.9. Statistics .....	272
7.4.	RESULTS .....	273
	7.4.1. Subjects.....	273
	7.4.2. Polysomnography.....	273
	7.4.3. Neuropsychological Testing.....	273
	7.4.4. Movement Measures .....	273
	7.4.5. Total sleep disturbance index .....	274
	7.4.6. CUdsic.....	274
7.5.	DISCUSSION.....	275
	Limitations.....	277
	Conclusion.....	277
7.6.	Figures and Tables.....	279
7.7.	References.....	288

**CHAPTER 8..... 299**

***Conclusion and Future work ..... 299***

---

8.1.	Overview.....	300
8.2.	summary of findings.....	302
8.3.	LIMITATIONS.....	304
8.4.	Future work.....	305
	8.4.1. NOVEL COMBINATIONS OF KNOWN FACTORS.....	305
	Physiological Factors.....	305
	Severity Factors.....	306
	Psychological Factors.....	306
	8.4.2. NOVEL MEASURES.....	307
	Sleep Spindles.....	307
	Disease Burden and Resiliency.....	307
	Esophageal Pressure Monitoring.....	308
	8.4.3. AUTOMATION.....	308
	8.4.4. CONCLUSION.....	309
8.5.	REFERENCES.....	310
	Inclusion Criteria For Candidate Spindles.....	315
	Exclusion Criteria For Candidate Spindles.....	315
	Staging Rules.....	316
	Scoring Rules.....	316
	Protocol.....	316
	<b>Quality of Life Survey.....</b>	<b>318</b>

---

## TABLE OF CONTENTS (Tables)

---

### **CHAPTER 4.....Error! Bookmark not defined.**

#### *Sleep Spindle Activity and Cognitive Performance in Healthy Children*

<b>Table 4.1</b> Demographic details.....	131
<b>Table 4.2</b> Sleep Parameters .....	137
<b>Table 4.3</b> Stanford Binet results .....	138
<b>Table 4.4</b> NEPSY results .....	139
<b>Table 4.5</b> Sleep spindle data.....	140

### **CHAPTER 5.....Error! Bookmark not defined.**

#### *EEG Changes Accompanying Spontaneous Arousals during Sleep Correlate with Neurocognitive Performance in Children with Upper Airway Obstruction*

<b>Table 5.1</b> Subject demographic data for control (C) , primary snoring (PS) and obstructive sleep apnoea syndrome (OSAS) children. ....	169
<b>Table 5.2</b> Polysomnography results for control (C), primary snoring (PS) and obstructive sleep apnoea syndrome (OSAS) children. ....	170
<b>Table 5.3</b> Mean neurocognitive test results and between group comparisons for control (C), primary snoring (PS) and obstructive sleep apnoea syndrome (OSAS) children. ....	172
<b>Table 5.4</b> Mean duration (in seconds) of spontaneous arousals and comparisons between control (C), primary snoring (PS) and obstructive sleep apnoea syndrome (OSAS) groups.....	174
<b>Table 5.5</b> Mean relative change (difference) in spectral power with spontaneous arousal onset observed between the different disease groups. PS = primary snorers; OSAS = Obstructive sleep apnoea syndrome. RP = Relative Power. ....	181
<b>Table 5.6</b> Correlations between neurocognitive testing results and Mean Relative Power Pre Post and During Spontaneous Arousals in NREM Stage 2 in the primary snorers (PS) group.....	192
<b>Table 5.7</b> Correlations between neurocognitive testing results and Mean Relative Power Pre, Post and During Spontaneous Arousals in REM in Primary Snorers (PS).....	193
<b>Table 5.8</b> Correlations between neurocognitive testing results and Mean Relative Power in Frequency Bands Around and During Spontaneous Arousals in SWS in the Primary Snorers (PS) group. ....	195
<b>Table 5.9</b> Correlations between neurocognitive testing results and Mean Relative Power Pre, Post and During Spontaneous Arousals in Stage 2 NREM in the OSAS group. ....	196
<b>Table 5.10</b> Correlations between neurocognitive testing results and Mean Relative Power Pre, Post and During Spontaneous Arousals in REM in the OSAS group.....	197

<b>Table 5.11</b> Correlations between neurocognitive testing results and Mean Relative Power in Frequency Bands pre, post and During Spontaneous Arousals in SWS in the OSAS group.....	198
--	-----

**CHAPTER 6.....**Error! Bookmark not defined.

*Movement in Sleep*

<b>Table 6.1</b> Demographic characteristics for control (C) primary snoring (PS) and obstructive sleep apnoea syndrome (OSAS) children at baseline (study 1) and follow up polysomnography (study 2).....	231
<b>Table 6.2</b> Baseline polysomnography results for control (C), primary snoring (PS) and obstructive sleep apnoea syndrome (OSAS) children. The grey shaded rows indicate traditional measures of sleep fragmentation.....	232
<b>Table 6.3</b> Follow-up polysomnography results for control (C), primary snoring (PS) and obstructive sleep apnoea syndrome (OSAS) children. The grey shaded rows indicate traditional measures of sleep fragmentation.....	234
<b>Table 6.4</b> Summary of movement measures for controls (C), primary snorers (PS) and children with clinically defined obstructive sleep apnoea syndrome (OSAS) at baseline (1) and follow up (2) polysomnography (PSG). .....	236

**CHAPTER 7.....**Error! Bookmark not defined.

*Towards a Composite Index of Upper Airway Obstruction Severity in Children*

<b>Table 7.1</b> Quality of Life Scale (OSA-18) with 20 original conceptual categories and scale items (adapted from Franco 2000).....	283
<b>Table 7.2</b> Mean subject biophysical characteristics for UAO children at polysomnography. ....	284
<b>Table 7.3</b> Polysomnography results for a single night polysomnography in children with UAO .....	285
<b>Table 7.4</b> Mean Values for Total Sleep Disturbance Index (TSDI) and component indices and correlations with Subjective Disease Severity (Quality of Life (OQOL)) and Objective Disease Severity (O2 nadir during sleep). .....	286
<b>Table 7.5</b> Mean Values for Composite Upper airway disease Disturbance of Sleep Index for Children (CUDSIC) and component indices and correlations with Objective Disease Severity (Apnoea/Hypopnoea Index and O2 desaturation index during sleep) and Subjective Disease Severity (Quality of Life – QOL) .....	287

## TABLE OF CONTENTS (Figures)

---

### **CHAPTER 1**.....Error! Bookmark not defined.

*Introduction to Sleep and Sleep Disorders in Children*

**Figure 1.1** Spectrum of Upper Airway Obstruction (UAO).....34

### **CHAPTER 4**.....Error! Bookmark not defined.

*Sleep Spindle Activity and Cognitive Performance in Healthy Children*

**Figure 4.1** Scatterplot of spindle mean central frequency against hand positioning (fine motor performance). ..... 152

### **CHAPTER 5**.....Error! Bookmark not defined.

*EEG Changes Accompanying Spontaneous Arousals during Sleep Correlate with Neurocognitive Performance in Children with Upper Airway Obstruction*

**Figure 5.1:** Group mean relative Alpha power in NREM stage 2 for prior to, following or during spontaneous arousals..... 175

**Figure 5.2:** Group mean relative Sigma power in NREM stage 2 for prior to, following or during spontaneous arousals..... 176

**Figure 5.3:** Group mean relative Beta power in NREM stage 2 for prior to, following or during spontaneous arousals..... 177

**Figure 5.4:** Group mean relative Delta power in NREM stage 2 for prior to, following or during spontaneous arousals..... 178

**Figure 5.5:** Group Mean Relative Theta Power in NREM stage 2 for prior to, following or during spontaneous arousals ..... 179

### **CHAPTER 6**.....Error! Bookmark not defined.

*Movement in Sleep*

**Figure 6.1** The exponential coefficient (theta) of the distribution of sleep runs terminated by wake or NREM stage 1 sleep for the baseline PSG in the control (C), primary snoring (PS) and obstructive sleep apnoea (OSA) groups..... 237

**Figure 6.2** The exponential coefficient (theta) of the distribution of sleep runs terminated by movement epochs for the baseline PSG in the control (C), primary snoring (PS) and obstructive sleep apnoea (OSA) groups..... 238

**Figure 6.3** The exponential coefficient (theta) of the distribution of sleep runs terminated by movement events (ME) for the baseline PSG in the control (C), primary snoring (PS) and obstructive sleep apnoea (OSA) groups. .... 239

**Figure 6.4** Correlation between the exponential coefficient (theta) of movement events (ME) and log transformed apnoea-hypopnoea index for the children with obstructive sleep apnoea at baseline PSG..... 240

**Figure 6.5** Correlation between changes in movement event exponential coefficient (Theta) between studies and changes in AHI between studies in the OSAS group. .... 241

**CHAPTER 7**.....Error! Bookmark not defined.

*Towards a Composite Index of Upper Airway Obstruction Severity in Children*

**Figure 7.1:** The Sleep Apnoea Severity Index..... 279

**Figure 7.2** Quality of Life (OSA-18) Survey Overall Quality of Life score (OQOL) Results ..... 280

**Figure 7.3** The Composite Upper Airway Obstruction Disease Severity Index for Children (CUDSIC). .... 281

**Figure 7.4** The Composite Upper Airway Obstruction Disease Severity Index for Children (CUDSIC). .... 282

## ***i. Abstract***

An examination of the nature of sleep fragmentation in children with upper airway obstruction.

**Introduction** – Sleep related upper airway obstruction (UAO) in children disrupts breathing in sleep, resulting in sleep fragmentation and subsequent neurocognitive and behavioural deficits. Unfortunately the nature of this fragmentation in children is poorly understood and a universally accepted, clinically valid, measure of sleep fragmentation has been elusive. This limits our ability to accurately determine and measure the consequences of sleep fragmentation on a child's development due to UAO, as well as the success of any treatment administered.

**General Aims** - The aim of the current study was to (i) examine the nature of sleep fragmentation in children with upper airway obstruction and (ii) to develop a new sleep fragmentation index for use in paediatric clinical populations with upper airway obstruction. When this study began no such index existed that was widely accepted and utilized. A range of sleep fragmentation measures already trialed in children with upper airway obstruction were reviewed to identify problems and limitations with current and previous methods of measuring sleep fragmentation in these children. An attempt was also made to identify other possible additional factors that mediate sleep fragmentation so as to develop a workable and generally applicable sleep fragmentation index for children with upper airway obstruction.

**Methods** – We performed a series of analyses on sleep and neurocognitive data from children with upper airway obstruction to identify and quantify neural activity associated with sleep fragmentation. We then used these measures and other mediating factors to create a composite measure of sleep fragmentation in children.

**Results** – We found that children with upper airway obstruction had characteristically altered neural activity as measured by electroencephalogram (e.g. changes in sleep spindle density,

decreased alpha and sigma power around spontaneous arousals from sleep). They also had an altered movement distribution in sleep (increased exponential distribution coefficient when sleep runs between movements are modeled on a survival curve), when compared to normal controls. The studies also demonstrated the potential ability of a composite measure of such sleep fragmentation markers and mediating vulnerability factors to more accurately and usefully quantify the negative impacts of upper airway obstruction.

**Conclusions** - Sleep fragmentation is a significant consequence of UAO in children, however the current measure of UAO severity is insufficient for determining the overall impact on a child's development. As this study demonstrates, the impact of sleep fragmentation is dependent on a complicated set of variables including: age, health factors (e.g. BMI), exposure time, disease severity (e.g. AHI), genetics, trait-like factors, social factors (e.g. SES) and family history. The arousals, or disruptions to sleep, are also altered in children with UAO compared to normal controls. We therefore propose a composite measure of these important factors as a more accurate tool for determining the impact of sleep fragmentation and overall severity of UAO in children.

## ***ii. Signed Declaration***

This work contains no material which has been accepted for the award of any other degree or diploma in any university or other tertiary institution and, to the best of my knowledge and belief, contains no material previously published or written by another person, except where due reference has been made in the text. I give consent to this copy of my thesis when deposited in the University Library, being made available for loan and photocopying, subject to the provisions of the Copyright Act 1968. The author acknowledges that copyright of published works contained within this thesis (as listed below) resides with the copyright holders of those works.

Chatburn, A., S. Coussens, K. Lushington, D. Kennedy, M. Baumert, and M. Kohler. (2013) "**Sleep spindle activity and cognitive performance in healthy children.**" *Sleep*, 36(2): pp. 237-243.

Coussens, S., M. Kohler, M. Baumert, J. Martin, D. Kennedy, K. Lushington, D. Saint, and Y. Pamula. "**EEG spectral changes associated with spontaneous arousals in children with upper airway obstruction.**" In *The Journal of Sleep Research*, vol. 20, (2008): pp. 52-52.

Coussens, S., Baumert, M., Kohler, M., Martin, J., Kennedy, D., Lushington, K., & Pamula, Y. (2014). "**Movement distribution: a new measure of sleep fragmentation in children with upper airway obstruction.**" *Sleep*, 37(12), pp. 2025-2034.

Scott Coussens

Date 29<sup>th</sup> January 2014

### ***iii. Acknowledgements***

First and foremost I would like to thank my supervisors Dr. David Saint and Dr. Yvonne Pamula for the opportunity. I am extremely grateful for your patience and guidance over the last several years. I especially would like to thank Yvonne for the constant motivation and support. You taught me the skills that have been vital for my growth from an undergraduate student to an independent scientist. I would also like to thank David Parsons for giving me a chance and beginning my research career. Your initial and continued support has helped me to become the enthusiastic researcher I am today.

I extend my gratitude to Department of Respiratory and Sleep Medicine Team, particularly Dr. Declan Kennedy and Dr. James Martin; The Women's and Children's Hospital including Kurt Lushington; and to the Department of Electrical and Electronic Engineering at Adelaide University including Matthew Berryman, Derek Abbott, Cosma Shalizi, and Andrew Allison. It has been a valuable experience working and learning from you all, without which this project would not be possible.

I would also like to acknowledge Dr. Mark Kohler for his invaluable knowledge, encouragement and selfless dedication to helping myself and others. Thank you for taking the time to read through, assist and provide feedback on my thesis, your sage council and the various studies and projects you have generously included me in. There is no way that this thesis would have been finished without you and I look forward to many years of working in research together.

Finally I would like to thank my colleagues, both past and present, for accompanying me on this journey. Your help, advice and encouragement has been greatly appreciated during the course of this research. In particular I would like to thank Alex Chatburn for sharing my passion for research.

I look forward to many collaborations in the future, and many more insightful discussions. And to Melissa Cava, my 'editor in chief'. Thank you for your personal and professional support over the last 5+ years.

This research was generously funded by The Women's and Children's Hospital, The University of Adelaide, Australian Research Council, NHMRC and Australasian Sleep Association.

#### ***iv. Publications***

Baraglia, D. P., M. J. Berryman, S. W. Coussens, Y. Pamula, D. Kennedy, A. J. Martin, and D. Abbott. "**Automated sleep scoring and sleep apnea detection in children [6039-35].**" In *PROCEEDINGS-SPIE THE INTERNATIONAL SOCIETY FOR OPTICAL ENGINEERING*, vol. 6039, p. 60390T. International Society for Optical Engineering; 1999, 2006.

Baraglia, David P., Matthew J. Berryman, Scott W. Coussens, Yvonne Pamula, Declan Kennedy, A. James Martin, and Derek Abbott. "**Automated sleep scoring and sleep apnea detection in children.**" In *Microelectronics, MEMS, and Nanotechnology*, pp. 60390T-60390T. International Society for Optics and Photonics, 2005.

Blunden, Sarah, Emily Watson, Gabby Rigney, Siobhan Banks, Scott Coussens, Gilly Hendrie, and Mark Kohler. "**TO-136 the effects of a high sugar diet on sleep quality and attentional capacity in prepubescent girls: a preliminary study.**" *Sleep Medicine* 12 (2011): S94.

Chatburn, A., S. Coussens, K. Lushington, D. Kennedy, M. Baumert, and M. Kohler. "**Sleep spindle activity and cognitive performance in healthy children.**" *Sleep* 36, no. 2 (2013): 237-243.

Coussens, S., M. Kohler, M. Baumert, J. Martin, D. Kennedy, K. Lushington, D. Saint, and Y. Pamula. "**EEG spectral changes associated with spontaneous arousals in children with upper airway obstruction.**" In *JOURNAL OF SLEEP RESEARCH*, vol. 20, pp. 52-52. COMMERCE PLACE, 350 MAIN ST, MALDEN 02148, MA USA: WILEY-BLACKWELL, 2011

Matthew Berryman, Scott Coussens, Martin, James A., David A. Saint, Derek Abbott, Yvonne Pamula, Declan Kennedy, Kurt Lushington, Cosma Shalizi, and Andrew Allison. "**Nonlinear aspects of the EEG during sleep in children.**" In *PROCEEDINGS-SPIE THE 3<sup>rd</sup> INTERNATIONAL SYMPOSIUM ON FLUCTUATIONS AND NOISE*, pp40-48, vol. 6039, p. 60390T. International Society for Optics and Photonics. 2005.

Pamula, Y., A. Campbell, S. Coussens, M. Davey, M. Griffiths, J. Martin, J. Maul *et al.* "**ASTA/ASA addendum to the AASM guidelines for the recording and scoring of paediatric sleep.**" In Journal of Sleep Research, vol. 20, no. Supp. 1, pp. 4-4. Wiley-Blackwell Publishing, 2011.

Coussens, S., Baumert, M., Kohler, M., Martin, J., Kennedy, D., Lushington, K., & Pamula, Y. "**Movement distribution: a new measure of sleep fragmentation in children with upper airway obstruction**". Sleep, 37(12), 2025-2034.2014.

**v. *Style and Referencing Format of Jointly Authored Papers, Documents and Unpublished Manuscripts***

The manuscript style and referencing format for chapters 1,2,3,5,7 and 8 are that prescribed by the journal Sleep Medicine.

The manuscript style and referencing format for Chapters 4 and 6 are that prescribed by the journal Sleep.

## ***vi. Abbreviations***

---

$\alpha$	<i>Alpha Wave Form In EEG (~ 9 – 12 Hz)</i>
ACPT	Auditory Continuous Performance Test
Adolescents	12-18 Years Old
AHI	Apnoea-Hypopnea Index
ANOVA	One Way Analysis Of Variance
Artotl	Total Arousal Index
BMI	Body Mass Index
C	Control
CA	Central Apnoea
CBCCL	Child Behaviour Check-List
Children	4-11 Years Old
CPRS	Conner's Parents Rating Scale
CRS-R	Conner's Rating Scale – Revised (Conners 1997)
DAS	Differential Abilities Scale
DBP	Diastolic Blood Pressure;
DS	Daytime Sleepiness
DTVMI	The Developmental Test Of Visual-Motor Integration
EMG (SM)	Electromyogram (Submental) – Chin Electrode Measuring Local Muscle Activity
ESS	Epworth Sleepiness Scales. A Subjective Measure Of Daytime Sleepiness
GDSVD - s	The Gordon Diagnostic System Vigilance And Distractibility Subtests (Gordon 1983)
Infants	1-3 Year Old
MAP	Mean Arterial Pressure
MDI	Mental Development Index

---

---

ME	Movement Events
MSLT	Multiple Sleep Latency Test. An Objective Measure Of Daytime Sleepiness
nCPAP	Nasal Continuous Positive Airway Pressure
NEPSY	Neuropsychological Developmental Assessment (Korkman 2001)
NIPPV	Nocturnal Non-Invasive Positive Pressure Ventilation
NN	SFI Calculated Using A Neural Network Approach To Complicated To Be Summarized In This Paper
ns	Not Significant
nt	Night Of Study Including Clinical PSG
OA	Obstructive Apnoea
OAI	Obstructive Apnoea Index
OSAS	Obstructive Sleep Apnea Syndrome
OSLER	Oxford Sleep Resistance Test. An Objective Measure Of Daytime Sleepiness
PeS	Esophageal Pressure Monitoring
PLMI	Periodic Limb Movement Index
PS	Primary Snoring
R/K	Remember/Know (R/K) Procedure (Tulving 1985)
RAI	Respiratory Arousal Index
REM	Sleep Stage REM (Rapid Eye Movement)
S1, S2, S3, S4	Sleep Stage 1, 2, 3 And 4 Respectively
SAI	Spontaneous Arousal Index
SaO <sub>2</sub>	Arterial Oxyhaemoglobin Saturation
SBP	Systolic Blood Pressure
SPT	Sleep Period Total
TAC	Thoracoabdominal Asynchrony
TCAoSAS	Subjects From The Tucson Children's Assessment Of Sleep Apnoea Study (Mulvaney 2005)

---

---

TST	Total Sleep Time (In Hours)
TWT	Total Wake Time In Hours
Ua	Unattended In Home PSG
WASI	The Wechsler Abbreviated Scales Of Intelligence
WCST- 64	The Wisconsin Card Sorting Test-64 Card Version
WISC-III	Wechsler Intelligence Scale For Children-Third Edition
wk	Weeks
WPPSI-R	Wechsler Pre-School And Primary Scale Of Intelligence–Revised
WRAML	Wide Range Assessment Of Memory And Learning

---