

**The Influence of Mother Care on the relationship
between Self-esteem and Neural Substrates in
young men and women: A Neuroimaging Study.**

**Mehereen Wadiwalla
Department of Neurology and Neuroscience,
McGill University, Montreal, QC**

Sept 2007

A thesis submitted to McGill University in fulfillment of
the requirements of the degree of M. Sc. in the department
of Neurology and Neurosurgery.

Mehereen Wadiwalla © 2007



Library and
Archives Canada

Published Heritage
Branch

395 Wellington Street
Ottawa ON K1A 0N4
Canada

Bibliothèque et
Archives Canada

Direction du
Patrimoine de l'édition

395, rue Wellington
Ottawa ON K1A 0N4
Canada

Your file Votre référence
ISBN: 978-0-494-38439-8
Our file Notre référence
ISBN: 978-0-494-38439-8

NOTICE:

The author has granted a non-exclusive license allowing Library and Archives Canada to reproduce, publish, archive, preserve, conserve, communicate to the public by telecommunication or on the Internet, loan, distribute and sell theses worldwide, for commercial or non-commercial purposes, in microform, paper, electronic and/or any other formats.

The author retains copyright ownership and moral rights in this thesis. Neither the thesis nor substantial extracts from it may be printed or otherwise reproduced without the author's permission.

AVIS:

L'auteur a accordé une licence non exclusive permettant à la Bibliothèque et Archives Canada de reproduire, publier, archiver, sauvegarder, conserver, transmettre au public par télécommunication ou par l'Internet, prêter, distribuer et vendre des thèses partout dans le monde, à des fins commerciales ou autres, sur support microforme, papier, électronique et/ou autres formats.

L'auteur conserve la propriété du droit d'auteur et des droits moraux qui protègent cette thèse. Ni la thèse ni des extraits substantiels de celle-ci ne doivent être imprimés ou autrement reproduits sans son autorisation.

In compliance with the Canadian Privacy Act some supporting forms may have been removed from this thesis.

Conformément à la loi canadienne sur la protection de la vie privée, quelques formulaires secondaires ont été enlevés de cette thèse.

While these forms may be included in the document page count, their removal does not represent any loss of content from the thesis.

Bien que ces formulaires aient inclus dans la pagination, il n'y aura aucun contenu manquant.


Canada

ACKNOWLEDGEMENTS	III
COMMONLY USED ABBREVIATIONS	VIII
ABSTRACT	1
RESUME	2
INTRODUCTION	3
<i>MATERNAL CARE AND SELF-ESTEEM.</i>	<i>5</i>
<i>What is Self-esteem?</i>	<i>5</i>
<i>Why we do we need self-esteem?.....</i>	<i>7</i>
<i>Self-esteem Development and Mother Care</i>	<i>9</i>
<i>Self-esteem, Mother Care and Hippocampus (HC).....</i>	<i>10</i>
<i>A case for an association between self-esteem, Mother care and the Prefrontal cortex.....</i>	<i>12</i>
<i>Prefrontal cortex and self-esteem have convergent ontogeny.....</i>	<i>14</i>
<i>Prefrontal cortex is also sensitive to maternal programming.....</i>	<i>15</i>
<i>HYPOTHESIS AND AIMS OF STUDY.</i>	<i>16</i>
2. METHODS AND MATERIALS	18
<i>PARTICIPANTS.....</i>	<i>18</i>
<i>NEUROPSYCHOLOGICAL ASSESSMENT.</i>	<i>18</i>
<i>The Parental Bonding Inventory (PBI) (Parker et al 1979).....</i>	<i>18</i>
<i>Rosenberg Self-esteem Scale (RSE) (Rosenberg, 1965):</i>	<i>19</i>
<i>Questionnaire for Competency and Control (QCC) (Krampen, 2001)</i>	<i>19</i>
<i>MRI ACQUISITION AND ANALYSIS.....</i>	<i>20</i>
<i>The Medial Prefrontal Cortex (MPFC) and the Dorsolateral Prefrontal Cortex (DLPFC) template. ...</i>	<i>22</i>
<i>Automated and semi automated segmentation.....</i>	<i>24</i>
<i>STATISTICAL ANALYSIS USING SEMI AUTOMATED DLPFC/MPFC VOLUME.....</i>	<i>25</i>
<i>ADDITIONAL AUTOMATED PROTOCOLS:</i>	<i>26</i>
<i>Voxel Based Morphometry (VBM):.....</i>	<i>26</i>
<i>Probabilistic maps</i>	<i>27</i>
3. RESULTS.....	29
<i>PARTICIPANT'S DEMOGRAPHICS.</i>	<i>29</i>
<i>CORRELATION ANALYSIS.</i>	<i>29</i>
<i>VOLUMETRIC DIFFERENCES BETWEEN MEN AND WOMEN IN LOW AND HIGH MOCA GROUPS.</i>	<i>31</i>
<i>Correlation Analysis</i>	<i>32</i>
<i>MEDIATIONAL ANALYSIS.</i>	<i>33</i>
<i>VBM RESULTS</i>	<i>34</i>
<i>PROBABILISTIC MAPS (PB MAPS) RESULTS.....</i>	<i>34</i>
4.DISCUSSION.....	36
<i>THE BI-REGIONAL ASSOCIATION BETWEEN MEN AND WOMEN.</i>	<i>37</i>
<i>Self esteem different for men and women.....</i>	<i>37</i>
<i>Mother care contributes to sex difference in self-esteem</i>	<i>38</i>
<i>NEGATIVE ASSOCIATION BETWEEN SELF-ESTEEM AND DLPFC VOLUME IN WOMEN: A POSSIBLE EXPLANATION.</i>	<i>40</i>
<i>THE SHAPE OF THINGS.</i>	<i>41</i>
<i>LACK OF MPFC ASSOCIATION.....</i>	<i>43</i>
5.LIMITATIONS AND FUTURE DIRECTIONS.....	45
<i>CONCLUSIONS</i>	<i>48</i>
6. BIBLIOGRAPHY.....	50
7. FIGURES	58
<i>FIGURE 1: THE MEDIAL PREFRONTAL CORTEX (MPFC) TEMPLATE</i>	<i>58</i>

FIGURE 2: THE DORSOLATERAL PREFRONTAL CORTEX (DLPFC) TEMPLATE	59
FIGURE 3: VOLUMETRIC GROUP DIFFERENCE BETWEEN LOW AND HIGH MOTHER CARE.....	60
FIGURE 4: ASSOCIATIONS BETWEEN DLPFC AND SELF ESTEEM IN HIGH AND LOW MOCA GROUPS.	61
FIGURE 5: ASSOCIATION BETWEEN HC VOLUME AND SELF ESTEEM IN HIGH AND LOW MOCA GROUPS.....	62
FIGURE 6: BAR GRAPH FOR DLPFC VOLUMES AS A FUNCTION OF SELF ESTEEM AND SEX IN HIGH MOCA GROUP	63
FIGURE 7: BAR GRAPH FOR HC VOLUMES AS A FUNCTION OF SELF ESTEEM AND SEX IN HIGH MOCA GROUP .	64
FIGURE 8: ASSOCIATION BETWEEN SELF ESTEEM AND DLPFC IN MEN AND WOMEN ON LOW AND HIGH MOCA GROUPS.....	65
FIGURE 9: ASSOCIATION BETWEEN SELF ESTEEM AND HC IN MEN AND WOMEN ON LOW AND HIGH MOCA GROUPS.....	66
FIGURE 10: VBM RESULTS.....	67
FIGURE 11: PROBABILISTIC MAPS.....	68
Figure 11a. Probabilistic maps in the Low MOCA female groups.	68
Figure 11b: Probabilistic maps for just Mother care in women.	69
Figure 11c. A comparison of high self esteem women in both high and low mother care groups.	70
8. TABLES	71
TABLE 1: DEMOGRAPHICS FOR THE WHOLE GROUP SPLIT BY MOTHER CARE.....	71
TABLE 2: PBI SCORES ON ALL FOUR QUADRANTS.....	72
TABLE 3: GROUP DEMOGRAPHICS FOR MEN AND WOMEN IN HIGH AND LOW MOCA GROUP	73
1.COMPLIANCE CERTIFICATE	74
2.QUESTIONNAIRES USED.	74

Acknowledgements

Kant claimed that science is organized knowledge, and wisdom is organized life, and I leave this lab with knowledge that I can apply, and the wisdom to realize what I want from it. A lot of the credit goes to my supervisor Dr. Jens Pruessner who very bravely, took on a philosophy major with dream in her eyes and questions about consciousness and the self. It was through his infinite patience and countless mistakes on my part that we came to a project that both of us could be proud of. He had enough confidence in my abilities to let me explore my potential while making all the tools available at my disposal. I was like a kid in a candy store sampling new techniques, creating templates and generally having a blast. The level of expertise that I have gained and the avenues it has opened up for me, are squarely due to Dr. Pruessner's supervision and guidance. So thank you Jens for everything I will always be grateful for everything that you have taught me. It was an honor and privilege to be your student and I will carry you and the lab with me wherever I go.

There is a woman who I consider my idol; Dr. Sonia Lupien. From the fateful moment when I entered her undergraduate course, and ever since, I continue to be inspired by this remarkable woman. Your passion for science, your creativity and your ability to inspire every one to give their best, is a joy to witness and learn from. If I can be half as good as you, I think I will be a damn good scientist. I guess over the years you may have realized that I am a bonafide member of the Sonia Lupien fan club and I still get a little star struck every time I meet you or see you. I hope one day, I can work with as a peer and earn your respect as woman, a mother and a scientist.

I have been fortunate to be part of a lab that thrives on corporation and camaraderie. I have met wonderful people like Alexandra Fiocco, Vincent Corbo, Carole Scherling, Benjamin Lai and all the rest who have on more than one occasion shared my joys, my sorrows and have made my life all the more richer and sweeter. So I would love this opportunity to thank every body in the Pruessner and Lupien labs from the bottom of my heart for sharing these precious moments with me.

There are a few exceptional people who started of as peers but have now become an integral part of heart. Being a prefrontal geek I am quite proud of my brain analogy when I refer to them as the “lobes” of my life, my sisters, my girls; Catherine Lord, Katarina Dedovic, Veronika Engert, Najmeh Khalili-Mahani and Claudia Buss. You girls have taught me important life lessons that neither TLC nor animal planet can ever teach me and I am thankful to each of you for your continuing presence.

My sweet Catou, I still remember the day I met you. I was the gawky undergraduate and I was doing a project as your summer student. You read my first attempt at a paper and made changes that were not only relevant, but also made me think on tangents I wouldn't necessarily have gone on. You still do that, but now, you do it as my friend. I love you for the long meaningful talks, the mind-blowing understanding and capacity for love you exhibit, the dinners and who can forget “Lost!” You've always been my wise go-to friend who listens to the crazy brown girl yammer away for hours and I am thrilled that you and I will get to spend some more time in close proximity. So here's to the new adventures of Catou and Me in Toronto!

What can I say for you my wonderful Kat? We have been joined to the hips pretty much since we met. I don't even remember when we crossed the line from just colleagues to friends. I guess when I barged into your room and claimed the empty desk, and I probably floored you with my charm! You have been a constant in my life for so many things. We have experienced so much together (BAW! Volunteering, teaching) and you have taught me so much over these years. Everything I know is in part due to your patient tutoring at some point or the other. I am so used to seeing you everyday, that it's going to be an adjustment to not have you there. After all, on average I do see you more than my boyfriend and parents combined! I will miss our never ending coffee money sagas, our rides together, and our gossip sessions. I love you so much and even though you and I may not be in the same city soon, I still intend to have "Mehereen time"... so kindly make the necessary arrangements!

My darling Vero you have always listened to me whine and shown an incredible thick skin to all my musings and rants. I can't remember the number of times you have been my sounding board for the crazy wacky ideas that ferment in my brain and the patience you exhibit has always helped me in more ways than one. There have been so many times when I have been, lets just say in a "hyper state of emotional discontent" and just looking at you across the table, or the room and you smiling at me knowingly, telling me silently "its going to be ok sweetie" has just made all the difference. I will miss that so much and of course whom else will I share the "perfect quiet moment" with.

My old, lovable Naj, you just get me. You get my inner struggles and demons that plague my soul. You understand the fundamental values I espouse and you know that there is a method to my madness. Very few people understand that behind my relatively simply façade lives a dreamer living on a temporally different plane. You get that and you don't judge me on it. You have often encouraged me to elaborate on my aspirations and have helped me to reacquaint myself with the wide-eyed philosophy major, who I thought I had lost along the way. Our general dissatisfaction with the world, our shared meaningful talks, I will lament the loss of that so dearly.

Oh Claudia, we started off in a rather unconventional way don't you think?? You choking on a pizza was I guess, the start of a long a fruitful association. The data we gathered has sustained my academic career so far and has provided the basis on which I can now fly solo. You taught me so much. Your drive, your creativity, your thirst to succeed is something I would love to emulate and espouse. I'm glad we got to meet in San Diego though we couldn't spend as much time as I like together, but it was so great to see you and see how well you are doing. Of course being close to the beach always helps but I'm glad that we are in the same continent and I look forward to working with you again.

My love Aamir, you are my "jaan". There is nothing without you. You are my biggest strength, and paradoxically, my biggest weakness. Your love and support over the last seven years have made this journey so much easier. Every insurmountable obstacle that I have faced, the horrible days of uncertainty seem just a little bit more bearable because you are with me. Your hugs, your unquestionable love, your fierce pride in me have nourished me and given me the strength to fight on. We are about to embark on a new journey and

with you by my side, holding my hand I know I can face anything. Your presence in my life has grounded me, given me the stability and comfort that comes from knowing that no matter how bad it gets, there is one person who thinks the world of you. I love you so much, my darling and when I look into your eyes I see not only my other half but the future father of my very cute kids (of course they will be cute! They will have their mother's genes!). So thank you for the unasked but timely caffeine fixes, the food, the quiet reassuring kisses, for every thing, my love.

Last but not the least, I can't forget to thank my parents, Aspi and Gulshan Wadiwalla. My parents have always wanted nothing but the best for me. The person I am today is in large part due to the love and care my parents gave me. They have been my biggest cheerleaders in the struggle of life. They may not like the decisions I make, but they have always supported me and have been stalwarts in my corner ready to fight with me, even though half the time they don't even know what the battle is. The love and support that my parents have given me, means the world to me. I love the both of you so much and the older I get, I appreciate and respect the sacrifices you have made for me over the years. My parents have always put me before themselves and I hope that they are as proud of me, as their daughter as I am of them being my parents. I love you Mom and Dad.

Commonly used Abbreviations

- I. ANOVA: Analysis of Variance
- II. BA: Brodmann Areas
- III. DLPFC: Dorsolateral Prefrontal Cortex
- IV. HC: Hippocampus
- V. MPFC: Medial Prefrontal Cortex
- VI. MOCA: Mother care
- VII. PBI: Parental Bonding Index
- VIII. PB Maps: Probabilistic Maps
- IX. PFC: Prefrontal Cortex.
- X. QCC: Questionnaire for Competency and Control
- XI. RSE: Rosenberg Self esteem
- XII. SE: Self esteem
- XIII. VBM: Voxel Based Morphometry

Abstract

Introduction: Numerous studies have suggested that maternal care can influence the development and expansion of an individual's self-esteem. Yet the neural mechanisms of this relationship remain unexplored. Incidentally, it has already been demonstrated that a brain region, namely the Hippocampus (HC) is associated with both self-esteem and maternal care. Thus suggesting that there may be a three-way relationship. This also provided the impetus to speculate that a similar interaction could be observed in other brain regions like for example, the Dorsolateral Prefrontal cortex (DLPFC) and the Medial Prefrontal Cortex (MPFC). Therefore, the aim of this study is to scrutinize the possible relationship between mother care, self-esteem and neural correlates including the DLPFC, MPFC and HC, with emphasis on how normal variations in mother care could have consequences for the relationship between self-esteem and particularly the prefrontal cortices. **Methods:** Fifty-one subjects were recruited on the basis of their maternal scores, as assessed by the Parental Bonding Index and were consequently assigned to either a high mother care (MOCA) or low MOCA group. Their self-esteem was measured through various self-esteem scales including the Rosenberg self-esteem scale. The structural integrity of the regions was ascertained through the use of both manual and semi automated segmenting procedures. **Results:** Initial Bivariate correlations reported a negative association between DLPFC volumes and self-esteem in the high MOCA group while HC was positively associated with self-esteem in both high and low MOCA groups. There were no associations to report for the MPFC. Additional analysis revealed that the Biregional association was sex specific. **Discussion:** For the first time, we were successful in associating DLPFC volume with self-esteem. In addition, we successfully replicated the association between self-esteem and HC volume. This study could provide an indication how of maternal care could have a sex specific affect not only on the evolution of self-esteem, but also on the regions they may be targeting.

Resume

Plusieurs études ont suggéré que le soin maternel peut influencer le développement et l'accroissement de l'estime de soi. Pourtant, les mécanismes neuronaux sous-jacents demeurent encore inconnus. Par ailleurs, il a été démontré qu'une région du cerveau en particulier, l'hippocampe (HC), est associée à l'estime de soi et au soin maternel, suggérant ainsi une relation à trois voies. Ceci a également fait émerger l'hypothèse qu'on pourrait observer une interaction semblable dans d'autres régions du cerveau par exemple, le cortex préfrontal dorsolatéral (DLPFC) et le cortex préfrontal médian (MPFC). Ainsi, le but de cette étude est d'examiner la relation entre soin maternel, estime de soi et structures cérébrales comprenant le DLPFC, le MPFC et l'HC en mettant l'accent sur comment les variations normen du soin maternel pourraient influencer la relation entre l'estime de soi et le cortex préfrontal. Méthodes: Cinquante et un participants ont été recrutés selon leurs scores au questionnaire sur les soins maternels, soit le «Parental Bonding Index». Les sujets ont ensuite été attribués aux groupes soin maternel (MOCA) faible ou élevé. L'estime de soi a été mesurée par diverses échelles, comprenant, entre autres, le questionnaire d'estime de soi de Rosenberg. L'intégrité structurale des régions cérébrales a été mesurée par des protocoles de segmentations manuelles et semi-automatisées. Résultats : Les corrélations bi-variées ont révélé une association négative entre les volumes du DLPFC et l'estime de soi dans le groupe MOCA élevé tandis qu'une association positive entre le volume de l'HC et l'estime de soi a été observée dans les 2 groupes (MOCA faible et élevé). Aucune association n'a été remarquée pour le MPFC. Des analyses additionnelles ont aussi indiqué que cette association bi-régionale était dépendante du sexe. Discussion : C'est la première fois que le volume du DLPFC est associé à l'estime de soi. De plus, nous avons répliqué l'association entre l'estime de soi et le volume de l'HC. Cette étude pourrait fournir une indication sur comment le soin maternel peut affecter un individu selon le sexe, non seulement dans l'évolution de l'estime de soi mais également sur les régions cérébrales impliquées dans l'estime de soi.

Introduction

Many studies have investigated the influence of Mother care on a child's social, emotional and cognitive development (Beers & De Bellis, 2002; Cicchetti & Rogosch, 2001; Cicchetti & Toth, 1995, 2000; De Bellis, 2005; Glaser, 2000; Hildyard & Wolfe, 2002; Keiley, Howe, Dodge, Bates, & Petti, 2001; Pelcovitz, Kaplan, DeRosa, Mandel, & Salzinger, 2000; Salzinger, Feldman, Ng-Mak, Mojica, & Stockhammer, 2001; Teicher et al., 2003; Teicher, Tomoda, & Andersen, 2006; Weber & Reynolds, 2004). In humans, there are critical periods in social development where the quality of mother care could result in different consequences (Cole et al., 2001). More specifically, Hildyard et al (2002) identified three key developmental periods during which variances in maternal care could have debilitating short-term and long-term effects on the child's socio-emotional, cognitive and behavioral development: infancy, young adolescence and older adolescence (Hildyard & Wolfe, 2002; Keiley et al., 2001).

A seminal study by Bowlby (1979) was the first to demonstrate the impact of early maternal-infant bonding, which sets the template for the child's future adult relationships. Interestingly, a 10-year follow-up on children whose initial level of care at infancy was classified as neglectful, revealed that they were more vulnerable to pathological life events in comparison to children with more adaptive maternal styles (Hildyard & Wolfe, 2002). In addition, during adolescence, they often exhibited compromised social adaptation and this trend continued all the way into adulthood where they were more distrustful of others, and often had negative self-representations (Cicchetti & Rogosch, 2002; Cicchetti & Toth,

2000; Hartner, 1999; Hildyard & Wolfe, 2002; Teicher, Andersen, Polcari, Anderson, & Navalta, 2002). Furthermore, neglected children are also more prone to delinquent behavior, running away from home and are four times at risk for elevated symptoms of anxiety and affective psychopathologies like depression, substance abuse and alcoholism (Clark, De Bellis, Lynch, Cornelius, & Martin, 2003; De Bellis et al., 2005; Hildyard & Wolfe, 2002). Besides the impact on the social and cognitive development of child, maternal neglect is also believed to affect the general health and mortality of the individual. These children are also at a greater risk for Type 2 diabetes and cardiovascular infractions (Barker, 2004).

Perhaps one of the best examples of how parental and mother care in particular, can have a significant impact on a child's social development is the Harvard Mastery of Stress Study. There, the investigators asked normal, healthy, college students to rate their parental care by completing the Harvard parental care scale. Thirty-five years later, those students that used a higher degree of negative words to describe their mothers were more likely to suffer from physiological infractions like hypertension and coronal complications (Russek & Schwartz, 1996, 1997a, 1997b), as well as psychological pathologies like depression and alcohol abuse (Canetti, Bachar, Galili-Weisstub, De-Nour, & Shalev, 1997; Parker, 1990; Sato, Uehara, Narita, Sakado, & Fujii, 2000). It is important to keep in mind that these individuals were not severely neglected or abused as children. The only difference in these children was the parenting style that was securing and loving in some, and aloof and disciplinarian in others. Yet those with inadequate maternal care styles were considerably disadvantaged in the long-term follow-up. On a positive note, this study also validated the

fact that good parenting can serve as a source of resilience and a buffer, providing better coping skills and quality of life (J. Smith & Prior, 1995) .

Maternal care and Self-esteem.

A key developmental milestone in socio-cognitive development is the management and expansion of self-esteem. Since it has been reliably established that mother care in fact, does influence the general social development trajectory, therefore, it stands to reason that it would similarly influence self-esteem development as well.

What is Self-esteem?

More than a century ago, William James (1890) was the first to formally define self-esteem as a value an individual places on themselves (James, 1890). Since then, many theories have attempted to explain the concept of self-esteem. However all of them ultimately differ on two major themes. The former is a teleological argument about the nature of self-esteem. More fundamentally, the question is if it is a personality trait or a psychological state. Most researchers tend to consider it as a relatively stable aspect of an individual, which one brings to a variety of situations, rather than as a dynamic state in response to them. On the other hand, many have conceptualized it as a state, which varies across situations and fluctuates in response to particular events.

The latter theme, in contrast seeks to investigate the influence of self-esteem in terms of whether it is a global all encompassing multimodal system or if it is restricted to a

particular facets of one's life i.e. it seeks its conception and expansion from limited and self-identified valuable domains of one's life. As a global measure, self-esteem adopts a top-down approach (Crocker et al 2001). The self-worth that an individual perceives for him or her self is not based on success in any particular domain but rather success in the course of life is attributed to the proper functioning of self-esteem in the first place. Others have conversely suggested that self-esteem could be a bottom-up process where it is predominantly contingent on continuing success in few particular domains. This consequently translates into a general feeling of wellness and increase in positive self-evaluation, which is then generalized to every other facet of one's life (Crocker et al, 2001).

The general consensus in any case for the first theme seems to be in favor of self-esteem being a stable personality trait that remains constant in most situations. However, there is no such agreement with respect to the second theme, and to date the issue remains contentious.

As mentioned earlier, many studies over the years have tried to conceptualize self-esteem in novel and innovative ways, but ultimately all of them end up being variants of the original two fundamental issues. Some notable examples would be theories by Crocker and Wolfe (2001) where they make a case for self-esteem being more domain-specific or "contingent vs. non-contingent (Crocker & Wolfe, 2001). Or Pelham's (Pelham & Swann, 1989) stable vs. unstable self-esteem, which is starkly reminiscent of the state vs. trait argument. Nevertheless, there is an underlying assumption that all the theories subscribe to; that is, the need for self-esteem is universal and indeed people are motivated to

maintain high levels of it and defend it when under threat (Pyszczynski, Greenberg, Solomon, Arndt, & Schimel, 2004).

Why we do we need self-esteem?

There has been an increasing interest in understanding the psychological need for self-esteem and its role in daily human interaction and growth. A possible explanation has been provided by the Terror management theory of self-esteem (Pyszczynski et al., 2004), which contends that self-esteem seeks to fundamentally mitigate the deep-rooted anxiety about an individual's mortality. It accomplishes this by enhancing an individual's personal value, which in turn, alleviates the apprehension one feels about their eventual demise. The sense of personal worth is obtained from believing in the validity and truth of one's own cultural views as well as the sense of living up to those cultural contingencies (Crocker & Wolfe, 2001). However while the cultural values are derived from the cultural milieu, they are largely integrated in individuals in purely subjective and thus create a uniquely individualized worldview. Hence self-esteem becomes an extremely personal experience where a person extracts what he or she considers valuable from the environment and that ultimately governs the course of the individual's life.

Another very influential theory is posited by Leary and Baumeister and is known as the "sociometer theory" (Leary, 2003). They speculate that self-esteem serves five fundamental needs; the maintenance of well being and positive effect, feedback for own coping and adequacy, reflection of individual status in social group, facilitation of self

determination and perhaps most importantly as providing vital information about individual's eligibility for social inclusion and exclusion.

The sociometer theory concludes that self-esteem “serves as a subjective monitor of ones own relational valuation” or the degree to which other individuals value their relationship with the person in question (R. F. Baumeister & Leary, 1995). It proposes that self-esteem acts an internal barometer that is sensitive to changes in one's relational value. While high self-esteem signifies an acceptance in the group, low self-esteem is associated with feelings of rejection and not feeling valuable to others (R. F. Baumeister & Leary, 1995).

Self-esteem therefore is a highly valued and imperative cognitive milestone whose proper functioning is required for the emotional and psychological wellbeing of an individual. Keeping in line with this, is the observation that high levels of self-esteem have subsequently been associated with both positive and negative life consequences. For example, positive self-esteem has been correlated with life affirming characteristics such as resilient coping styles, longevity, general happiness and satisfaction with oneself (R. F. Baumeister, Campbell, J.K., Krueger, J.I., Vohs, K.D., 2003), while low self-esteem has been causally implicated in substance abuse and affective disorders including depression, anxiety, antisocial behavior and eating disorders (Crocker & Wolfe, 2001; Hoyle, 1999; Leary, 2003).

Self-esteem Development and Mother Care.

Self-esteem development can be construed as an “experience dependent phenomenon” (Greenough, Black, & Wallace, 1987). The earliest environment a child experiences is contingent on the quality of parental care he or she is exposed to. While paternal care is important in mitigating a child’s social development (Hall, Peden, Rayens, & Beebe, 2004; Heider, Matschinger, Bernert, Alonso, & Angermeyer, 2006), maternal influence is, more often than not, implicated in the management and expansion of the child’s social self (Heinonen, 2003). Therefore, the initial environment can set the stage for future emotional, cognitive, and behavioral outcomes (Cole et al., 2001). Indeed, there is evidence that self-esteem ontogenesis is greatly dependent on the quality of maternal attachment the child receives (Bowlby, 1969). Interestingly, children of anxious/ambivalent and avoidant mothers often exhibit behavioral responses that are strangely reminiscent to responses that one expects from low self-esteem men and women (Bost et al., 2006; Domingo, Keppley, & Chambliss, 1997; Heider et al., 2006; Johnson, Gilbert, & Herdt, 1979; LeCuyer-Maus, 2000; Leifer, Kilbane, & Skolnick, 2002; McGarvey, Kryzhanovskaya, Koopman, Waite, & Canterbury, 1999; Uehara, Sato, Sakado, & Someya, 1998).

Furthermore, a child’s ability to internalize feedback obtained during the course of their maturation plays a significant role in the construction of their self-esteem and self-defining characteristics (Block, 1993; Hartner, 1999; Mead, 1934). Additionally, adolescence is a

time where a child creates, develops and internalizes a worldview, which then assists in establishing their self worth in the scheme of things. Maternal influence at this point, serves as a catalyst in form of support and nurturing and also, as a source for internalizing world views (Pyszczynski et al., 2004). That is, a child looks to the maternal figure as a model for the creation of his or her own value system. Therefore maternal care can directly and indirectly impact the development of self-esteem.

Self-esteem, mother care and neural substrates.

In short, it is apparent that mother care directly contributes to the development of self-esteem. There is an existing body of literature, which has studied the behavioral ramifications of how mother care can influence self-esteem. More recently, however, the focus has shifted to the arena of neural and biological substrates and as a result, there has been an increasing interest in studying this relationship through the context of neurobiological substrates. Two main regions of interest include the Hippocampus (HC) in the Medial temporal lobe, and the Prefrontal cortex (PFC) in the frontal lobes.

Self-esteem, Mother Care and Hippocampus (HC).

To date, there is some evidence that both mother care and self-esteem are linked to the morphology of the hippocampus (HC). The first paper to disentangle the influence of mother care on the hippocampal volume was by Buss et al (2007). They posited that higher

levels of postnatal maternal care could mitigate the influence of prenatal adversity on the hippocampus. More specifically, higher mother care resulted in higher HC volume in a group of participants who were designated as a high prenatal stress group. It is important to keep in mind that this group reported higher levels of prenatal stress that included emotional and financial distress during pregnancy as well as birth complications. Despite the disadvantaged beginnings, those individuals who had high levels of post natal maternal care, were considerably better off than their counterparts whose levels of maternal care was less than stellar. These individuals reported lower than average hippocampal volumes. Thus suggesting that good postnatal maternal care could in fact, serve as a buffer against the debilitating effects of early prenatal insults. Furthermore, animal studies have provided evidence of a similar phenomenon in rodents where higher levels of maternal care has been known to affect the neurogenesis of the HC (Bredy, Grant, Champagne, & Meaney, 2003; Liu, Diorio, Day, Francis, & Meaney, 2000; Liu et al., 1997; Weaver, Meaney, & Szyf, 2006).

With respect to the link between self-esteem and hippocampal volumes, Pruessner et al (2005) were first to discover this association. They demonstrated that higher levels of self-esteem were significantly associated with higher hippocampal volumes and more interestingly, this association was consistent across the lifespan. This paper was the first to show a link between self-esteem and the structural integrity of a brain region and thus raised the possibility that this relationship could also be realized in other regions of the brain as well.

A case for an association between self-esteem, Mother care and the Prefrontal cortex

This being said, another region that should also be investigated is the Prefrontal cortex (PFC). The PFC has a well established roles in the domain of the self (Eslinger, Flaherty-Craig, & Benton, 2004; Eslinger, Grattan, Damasio, & Damasio, 1992; C. D. Frith & Frith, 1999; Gallagher, 2000) as well has been known to be vulnerable to mother care (Teicher et al., 2002). Evidence below will be provided to make a case for investigating the Prefrontal cortex in relation to self-esteem and mother care.

The Prefrontal Cortex.

The PFC is a heterogeneous area, which is most developed in primates and it is an area that has afferents to cortical sensory, motor and sub-cortical structures (Miller & Cohen, 2001). It is situated in the frontal lobe and can be divided into four subdivisions: Medial, Orbital, Dorsal and Dorsolateral. In humans and non-human primates, the PFC includes areas rostral to the premotor cortices. It also includes part of the anterior Cingulate cortex including Brodmann (BA) areas 24a, 24b, 23a and 23b. Each of these areas have their own vigorous connections with the amygdala, HC, entorhinal and parahippocampal cortices (Carmichael & Price, 1995). It also has direct connections with the inferior parietal lobe, the amygdala and the occipital cortex (Fossati et al., 2004). For the purposes of our study the two areas of special interest within the PFC, are the Dorsolateral Prefrontal Cortex (DLPFC) and the Medial Prefrontal Cortex (MPFC).

The Dorsolateral Prefrontal cortex (DLPFC)

The DLPFC is the most lateral portion of the prefrontal cortex and has been featured prominently in the working memory literature (Mitelman, Buchsbaum, Brickman, & Shihabuddin, 2005; Petrides, 2000a, 2000b), as well as schizophrenia (Mitelman et al., 2005; Prasad, Sahni, Rohm, & Keshavan, 2005). Its primary role is the cognitive assessment and decision-making using visceral emotional and environment cues for the process of executive functioning. It accomplishes this by initiating a triangular circuitry between HC, thalamus and itself (Barbas, 1995; Carmichael & Price, 1995; Petrides & Pandya, 1999). Thus making this area integral to the functioning of emotional and cognitive decision-making.

The Medial Prefrontal cortex (MPFC).

The MPFC, in concert with the DLPFC is involved in the integration and processing of sensory information from the external world, (Barbas, 1995; Carmichael & Price, 1995). It is colloquially referred to as the “area of the self” (C. Frith & Frith, 2005; U. Frith & Frith, 2003) and one of the areas that shows high neural activity at rest that is, it is functionally active at baseline (Gusnard, 2005; Gusnard, Akbudak, Shulman, & Raichle, 2001; Gusnard, Raichle, & Raichle, 2001). In addition, it also has the highest amount of activity when the task at hand is self-relevant (Raichle & Gusnard, 2005; J. R. Simpson, Jr., Drevets, Snyder, Gusnard, & Raichle, 2001; J. R. Simpson, Jr., Snyder, Gusnard, & Raichle, 2001).

Together these two subdivisions of the prefrontal cortex are responsible for the monitoring of self-relevant information and higher order cognitive processes such as decision-making, perception and emotional regulation (D'Argembeau et al., 2005; DeYoung, Peterson, & Higgins, 2005; C. D. Frith & Frith, 1999; Gusnard, Akbudak et al., 2001; Northoff & Bermpohl, 2004; Ochsner et al., 2005; Petrides, 2005; Schmitz, Kawahara-Baccus, & Johnson, 2004; J. R. Simpson, Jr., Snyder et al., 2001) and therefore, are perfectly poised to contribute to the evolution and maintenance of self-esteem.

Prefrontal cortex and self-esteem have convergent ontogeny

Perhaps the most significant factor in considering a relationship between self-esteem and DLPFC and MPFC in particular, is that these regions experience exponential growth during early adolescence, which coincides with the maturing of social cognition in children (Andersen, 2003; Teicher et al., 2003; Teicher et al., 1997). Early puberty is a time where a child develops a sense of social order and an understanding of societal rules (Nelson et al 2005). Incidentally, right around this time, the young adolescents also learn to actively and quite expertly engage in theory of mind processes (Gallagher et al 2003) that is, the ability to perceive an other's state of mind. Furthermore, during this period there is an eventual cognitive shift, which enables them to regulate their emotions and engage in necessary inhibitory responses (Nelson et al 2005). The social evolution continues right unto the second decade of one's life where these early social experiences assist in molding and consolidating the individual's basic self worth (Hildyard 2002). So far the neural underpinnings of these burgeoning social networks have not been studied. This is quite

surprising considering that along with this dramatic period of social growth, there are concurrent changes occurring in the prefrontal cortex, as a result of extensive neurogenesis and pruning (Teicher et al., 2002). The mirroring of these two processes, precisely at the same time is intriguing to say the least. In short, the convergent ontogeny of these two processes suggests a possible link that cannot simply be attributed to chance.

Prefrontal cortex is also sensitive to maternal programming

In addition to having theoretical reasons for being associated with self-esteem, there are also waves of studies, which have demonstrated that a gross lack of maternal care at least, can affect the morphology and integrity of these neuro-anatomical structures as well. Abused children tend to have smaller volumes in different sub regions of the prefrontal cortex (Bremner, 2002; Glaser, 2000; Teicher, 2002; Teicher et al., 2002; Teicher et al., 2003).

In brief it is possible that key limbic structures, including HC and PFC, are vulnerable to variations in mother care. While it is certainly true that in most cases, the effect on the integrity of neural structures can be attributed to a lack of maternal care, it cannot be ignored that normal deviations in mother care have also been known to exert a tremendous influence on the social and cognitive development of an individual and therefore, by extension can be equally important for the development and integrity of key morphological structures.

Hypothesis and aims of study.

The literature reviewed above suggests that mother care can be linked to self-esteem. In addition, evidence has been put forth showing a link between quality of mother care and neurodevelopment of brain regions. Furthermore, results from our lab indicate that there is a relationship between integrity of brain structures (HC in particular) and self-esteem. Interestingly, ample background literature suggests that such an interaction could also exist in regions of the prefrontal cortex. However, this relationship remains unexplored, particularly, in the PFC regions of the DLPFC and MPFC.

Considering that mother care is the one variable that seems to influence the development and maintenance of both self-esteem and neural substrates, a case can be made for suggesting that variation in both self-esteem and neural substrates are a consequence of variations in maternal care. We hypothesize that higher levels of maternal care would be more likely to contribute to a positive association between self esteem and Hippocampal volume and given the precedence in the hippocampus, we expect to see a similar positive association in the prefrontal cortex as well. The main reason for investigating the volumes of these structures is that it can be taken as a proxy measure for cellular processing. In fact, recent studies have suggested that volumes might reflect differential neuronal and glial packing density, as well as differences in neuronal soma sizes which could be indicative of proper functioning (Miguel-Hidalgo et al., 2002; Stockmeier, 2004).

Nevertheless, the one thing that is beyond the scope of this study is to establish directionality in the association between self-esteem and neural substrates. This is a considerably problematic venture, especially in the case of the Prefrontal cortex. Given that both self-esteem and prefrontal development share convergent growth periods, it is therefore unclear which one is the causal variable. On the other hand, there is controversy about the directionality of the association between self-esteem and hippocampus. There has been speculation that a pre-existing abnormality in its morphology could serve as a vulnerability factor for future affective and cognitive disturbances (Gilbertson et al., 2002). Furthermore, the early maturity of the HC makes it principally sensitive to early maternal programming (Akers et al., 2006; E. Gould & Tanapat, 1999; McEwen, 2003). So it is unclear if a personality variable could be contributing to the morphological variances or if pre-existing abnormalities may cause the individual to embrace certain personality characteristics. Thus, our question is simply to investigate if maternal care can influence the association between self-esteem and neural substrates.

Therefore, the aim of this study is to scrutinize the possible relationship between mother care, self-esteem and neural correlates including the DLPFC, MPFC and HC, with emphasis on how normal variations in mother care could have consequences for the relationship between self-esteem and particularly the prefrontal cortices. Furthermore, since previous studies have already documented a link between self-esteem and HC, we would also like to replicate that finding in our own sample.

2. Methods and materials

Participants

Fifty-one (26 women) healthy university students between the ages of 20-32 (mean age =23.14 years \pm 3.04) were recruited on and around McGill university campus. Written consent was obtained from each subject prior to entering the study. The Douglas Hospital Research Centre Ethics Board and the Montreal Neurological Institute Ethics Board approved the study. Each respondent was specially selected based on the Mother care score (G. Parker, Tupling, H., Brown, L.D., 1979) and were consequently assigned to either a high or low Mother care (MOCA) group.

Neuropsychological assessment.

Psychological assessment included the following questionnaires:

The Parental Bonding Inventory (PBI) (Parker et al 1979)

The PBI is a standard instrument to retrospectively measure parental style (Parker 1979). The 48-items questionnaire assesses care and overprotection children received independently from mother and father during their first 16 years of life, resulting in four subscales: mother care, mother overprotection, father care, and father overprotection. We used the Mother care variable as our discerning variable of choice. We employed the mean

split method to create two groups: Low Maternal care (Low MOCA) and High Maternal care (high MOCA) groups.

Rosenberg Self-esteem Scale (RSE) (Rosenberg, 1965):

RSE is a 10 scale self report that assesses global self-esteem levels. It was designed to be a Guttman scale, which means that its items were to represent a continuum of self-worth statements. Rosenberg (1965) scored his 10-question scale that was presented with four response choices, ranging from strongly agree to strongly disagree. The 10 items RSE has been translated into many languages and has been used worldwide as measure of global self-esteem.

Questionnaire for Competency and Control (QCC) (Krampen, 2001)

The QCC is a 32 item questionnaire that uses the four primary scales: global self-concept, chance, control of others and internality. It combines aspects of locus of control, self-concept and self-esteem (Krampen 2001). The internality sub scale was used as a mean to validate our results as previous findings from our lab have shown that Rosenberg self-esteem is highly correlated with internal locus of control as well as the global self-concept (Pruessner et al 2005).

Additional battery of questionnaires included NEO Personality Inventory (Costa, Busch, Zonderman, & McCrae, 1986) , Mastery (Pearlin & Schooler, 1978) and Ways of coping (Vitaliano, 1985).

MRI Acquisition and analysis

Structural scans for all subjects were acquired using the protocol of the international consortium of brain mapping (ICBM) initiative to create a statistical atlas of the normal adult brain (Mazziotta, Toga, Evans, Fox, & Lancaster, 1995). This protocol generates T1, T2 and PD weighted images. For the purpose of this analysis, only T1 weighted images were used. Several algorithms were applied in order to prepare the images for manual segmentation. This included non uniformity corrections (Sled, Zijdenbos, & Evans, 1998). They correct for MRI intensity inhomogeneities due to individual anatomical variability, or poor Radio Frequency (RF) coil uniformity that generates signal drift.

The second process is linear stereotaxic transformation (Collins, Neelin, Peters, & Evans, 1994) into coordinates based on the Talairach atlas (Talairach, 1988). This process allows for comparison across groups and large subjects by insuring that the results are as coordinates in a standardized space. This is accomplished through affine linear registration using the ICBM 152 average model. The affine registration protocol transforms the file by rotating, translating, scaling and skewing the file in all three x, y and z dimensions, in order to ensure that the brain is as standardized as possible.

After the registration the file undergoes non-intensity normalization, which makes the signal range comparable between subjects. In addition, the files were subjected to an automated tissue classifying protocol created by Zijdenbos (1994) which specifically generates individual subject grey matter (GM), white matter (WM) and Cerebro-spinal fluid (CSF) maps. It accomplishes this by classifying every voxel in the file as either grey

white or CSF. The process can be split into two parts. The first is a fully automated “prior probability method” which based on an existing template deduces the probability of a voxel belonging to any of the three categories. The second step of this process is more intuitive as it recalculates the probability of a voxel being a grey matter voxel, for example by conducting a k-neighbor clustering interpolation where it assesses the likelihood of a particular voxel based on the category its neighbor belongs to (Cocosco, Zijdenbos, & Evans, 2003).

The Volumetric analysis was performed with the interactive software package DISPLAY developed at the Brain Imaging Centre of the Montreal Neurological Institute in Montreal, QC, Canada.

Hippocampal volume was assessed using a highly validated and widely used protocol developed in our laboratory (For more details please refer to Pruessner et al (2000)). Structural segmenting of the hippocampus was conducted by Dr. Claudia Buss and was validated by Dr. Jens Pruessner.

Since there were no pre-existing protocols on the assessment of volumetric difference in the PFC, we developed a new segmentation protocol to specifically evaluate the MPFC and the DLPFC regions.

The Medial Prefrontal Cortex (MPFC) and the Dorsolateral Prefrontal Cortex (DLPFC) template.

The labels were created on an average brain map of all the subjects to ensure the best fit possible. The author undertook the creation of the two labels manually.

MPFC label

The MPFC label includes Brodmann Area (BA) 9m, 10m, 24 and 32. BA 9 has a lateral and medial component and 9M encompasses the most rostral portion of the medial longitudinal fissure. BA 10 refers to the frontal pole and 24 and 32 are part of the anterior cingulate cortex.

When segmenting the MPFC regions, a precise way to orient your self is to paint primarily in the coronal view, which proves to be the most reliable indicator of boundaries. Therefore, in this view in particular, the most rostral part of the frontal pole (10) acts as its anterior boundary. The posterior boundary is evident on the most caudal slice in which the anterior commissure is visible. The ventral border is the most caudal slice in which the inner curvature of the corpus collusum is visible. In the sagittal view the anterior border was defined as the first deep sulcus measured from the genu of CC and is usually the cingulate sulcus (See figure 1).

DLPFC label

The DLPFC cortex is the most lateral portion of the prefrontal cortex and includes BA 8,9 and 46. These regions are encapsulated in the superior and middle frontal gyri.

BA 8, which is also known as intermediate frontal region, and 9, which is known as the granular frontal, has a lateral and medial component. BA 46 is located primarily in the middle frontal gyrus.

With respect to the borders, the best option is to segment in the coronal and sagittal views as they allow the most optimal view for segmenting. Therefore, in the coronal and sagittal views, the ventral lateral border is the inferior frontal sulcus. The Posterior Border is the precentral sulcus and the anterior border is demarcated by the frontal pole region which Tisserand et al (2002) designated as all anterior slices before Y=44. Anatomically after this slice the inferior frontal sulcus starts to become visible in the coronal view. The rostral medial border is the most rostral part of the medial longitudinal sulcus. In the sagittal view the anterior border was the paracingulate sulcus, which is dorsal to the cingulate cortex. In case the paracingulate was not discernible, the most caudal slice was designated as the slice in which the curvature of the corpus collusum was visible (See Figure 2).

Automated and semi automated segmentation

With the prefrontal labels created the next step was superimposing the labels unto the individual subject grey matter map files (Cocosco et al., 2003). The files are in MINC (medical imaging net CDF) format, which is a medical imaging file format. The MINC format was created by Peter Neelin in 1993 and is based on the UNIX platform. In order to work with the files a series of MINC tools were employed to initiate the necessary changes.

The first was ‘minclookup- discrete which performs lookup table operations on each voxel in the classified files. The most common use of the lookup table is to convert intensity values into RGB colors and where all input values range from 0 to 1(therefore instead of 3 separate labels for GM, WM and CSF there are only two labels 0 and 1). This method achieves this by employing discrete look up tables, where the classified files are remapped and each input value, (in this case the grey matter, white matter and CSF classified voxels) is compared to a look up table and if the input value is found, a corresponding output value is prescribed. If it is not found then a null value is indicated for that particular voxel. The algorithm is specifically designed to select the GM by assigning the integer value of 1 to it, while the white matter and CSF are consequently discarded as they are automatically assigned a null value.

Furthermore, the dimensions of the DLPFC and MPFC labels were resized with the intention of making them more compatible with classified files. The MINC command that is used is “Mincreshape”. This command initiates a series of conversions including

changing type, range and normalization of the pixel values, expanding or contracting images to give a specified image size. It achieves this by normalizing the image to a real value in this case the minimum being 0 and the maximum being 255.

The next step of this procedure involves resampling the individual files to be in the same spatial dimensions as the label template file therefore ensuring that the individual classified file is resampled in the same voxel positions as the template. After correcting for file dimensions, label template was superimposed on the individual on individual's files to ascertain an approximate estimate of volume (for both MPFC and DLPFC), for each subject. However, in order to account for the interindividual variability found in these regions, manual correction of each individual file was subsequently undertaken.

The volume difference before and after the manual correction was significant for both the DLPFC ($t_{(45)} = 14.08$ $p < .001$) and MPFC ($t_{(46)} = 26.75$ $p < .001$). All statistical analyses were therefore conducted only on the manually corrected volumes.

Statistical analysis using Semi Automated DLPFC/MPFC Volume

The subjects were divided into low and high Mother care (MOCA) groups using the mean split method. The Mean score (Mean = 26 ± 8) was used as a cut off point in order to facilitate the creation of the groups in this sample as it represents a better estimate of the population. Since our sample was normally distributed, the mean, median and mode were

not significantly different thus the mean was deemed to be a good estimate for our sample in general. Analysis of variance (ANOVA) and student t tests were employed in order to assess mean differences between our groups, where suitable.

Bivariate Pearson Correlational Analysis was employed to assess an association between self-esteem and regions of interests in our groups. In course of our analysis we also conducted Mediation analysis using hierarchical regression analysis approach in order to see if mother care mediated the relationship between self-esteem, hippocampus and the PFC. The dependent variable was either PFC volume or HC and the independent variable was entered as Rosenberg self-esteem score and mother care. All statistical analysis was conducted using the Statistical Package for Social Sciences (SPSS) 11.0.

Additional automated protocols:

Voxel Based Morphometry (VBM):

As an initial exploration, Voxel Based Morphometry (Ashburner & Friston, 2000) was used in order to confirm if there was an association present between self-esteem and the prefrontal cortex quite early in the project and this provided the basis for our future explorations. It is a computational neuroanatomy method to assess anatomical differences throughout the brain. It is usually used for a voxel wise comparison of local content of grey matter between two groups. It essentially looks for differences in local composition of brain tissue after macroscopic shape differences have been accounted for. VBM analysis

was conducted after spatially normalizing the native brains in standard stereotaxic space and partitioning grey matter, white matter and cerebrospinal fluid. This process, which is known as segmentation, is primarily used so that grey matter and white matter regions can be correctly identified for future analysis. The next step of preprocessing involves smoothing the regions with a Gaussian kernel called full width half maximum (FWHM). Smoothing allows VBM to be comparable to region of interest analysis as each voxel in the smoothed image contains an average concentration of grey matter around the voxel (Ashburner, Andersson, & Friston, 2000; Ashburner & Friston, 1999, 2005; Friston & Ashburner, 2004). Statistical analysis involved the use of general linear modeling (GLM)(Worsley, Taylor, Tomaiuolo, & Lerch, 2004) to identify regions for grey matter concentration that were related to the specific parameters of the study.

Probabilistic maps

Probabilistic maps (PB maps) were computed from the manually corrected individual structural files in order to check for the probability that the difference may be attributed to the homogeneity of the structure. PB maps indicate the probability for each voxel to be included in the volume of the structure in one subject across all subjects of a prescribed group. For each voxel the probability is indicated ranging from 0 to 1. A high probability indicates that for many subjects in the group that particular voxel is a part of the structure examined. This implies that if there are any shape differences the map will show voxels with lower probability scores. Consequently, if the shape is homogenous the map will show less voxels but each with higher probability (Corbo, Clement, Armony, Pruessner, &

Brunet, 2005). Three separate sets of analysis were conducted with our respondents being grouped according to their self-esteem and mother care affiliation.

3. Results

Participant's demographics.

There were 27 participants in the High MOCA group and 24 in the Low MOCA groups. There was a significant difference between low and high MOCA groups in their Mother care scores ($t_{(49)}=11.11$ $p < .001$). There was also a difference in Rosenberg self esteem scores ($t_{(49)}= 2.572$ $p < .01$). Neither age nor QCC internality scores were significantly different between both groups. (See table 1). Additionally there were no significant volumetric differences observed between low or high MOCA groups for either the left ($t_{(45)}= 1.2$, $p > .05$) or right ($t_{(45)}= .774$ $p > .05$) DLPFC, left ($t_{(46)}= 1.7$, $p > .05$) or right ($t_{(46)}= 1.1$, $p > .05$) MPFC or left ($t_{(47)}= .81$, $p > .05$) or right ($t_{(47)}= .25$, $p > .05$) Hippocampal volume (See figure 3).

Correlation Analysis.

In order to assess if there was an association between the morphological integrity of brain regions and self-esteem, we conducted Pearson correlation analysis in low and high MOCA groups separately. It is important to note here that, in our sample, higher mother care coincided with higher levels of father care as well as, lower levels on maternal and

paternal overprotection; therefore, higher mother care can also be construed as a proxy for optimal parenting styles (See table 2).

In the high MOCA group, our initial analysis revealed a negative association between self-esteem and left ($r = -.409$, $p = .05$) and right ($r = -.399$, $p = .05$) DLPFC (See figure 4). There was no significant association with MPFC ($p > .05$) or HC volume ($p > .05$) in both hemispheres. With respect to the internality subscale of the QCC, we observed a similar pattern of association with both left ($r = -.494$, $p = .02$) and right ($r = -.439$, $p = .03$) DLPFC. There were no significant associations to report for either MPFC or HC volumes.

In the low MOCA group, there was no association observed between self-esteem and neither DLPFC ($p > .05$), nor MPFC ($p > .05$) volumes. Interestingly, there was a strong positive association observed between right HC volume and self-esteem ($r = .486$, $p = .02$) (see figure 5). There were no significant associations reported in the left HC ($p > .05$). Furthermore, there were no significant associations evident for the Internality subscale and all three volumetric measures. In addition there was no association between total gray matter volume and self esteem in either groups ($p > .05$) as well.

To summarize, while we observed a negative association between left and right DLPFC volume, Rosenberg self-esteem (RSE) and the internality subscale of the QCC in the high MOCA group, the only significant association in the low MOCA group was found between right Hippocampal volume and Rosenberg self-esteem.

The next question that we wanted to investigate was if sex differences could be a contributing factor in the associations described above; especially since there is evidence that there are sex differences found in the prefrontal cortex. (Gur et al., 2000; Tisserand et al., 2002). With respect to the hippocampus, we also had reason to believe that we could be seeing a similar sex difference especially since the association between self-esteem and Hippocampal volume has so far only been evident only in men (Pruessner et al 2005). Therefore lending credence to the possibility that sex differences could be an important variable to consider for further analysis.

As a result, we decided to perform all subsequent analysis with sex as a grouping variable in addition to mother care. We consequently had 13 men and 11 women who reported high MOCA scores and 11 men and 12 women who reported low MOCA scores. (See table 3). With respect to personality measures, there were significant differences in PBI Mother care, as expected, between low and high MOCA men $t_{(23)} = 7.373$ $p = .001$ and women $t_{(24)} = 8.398$ $p = < .001$.

Volumetric differences between Men and Women in Low and High MOCA groups.

Additional analysis revealed that there was an 11% difference between right DLPFC volume and a 12% difference in Left DLPFC volume between low and high self-esteem groups in the high mother care group only in women but not in men ($f_{(1,20)} = 4.485$ $p = .047$ (DLPFC X RSE X SEX), mixed design ANOVA) (see figure 6) which revealed that

women with higher self-esteem has lower DLPFC volumes in the high MOCA group. There were no significant effects to report in the Low MOCA group.

Intriguingly in the hippocampus, we discovered that there was a volumetric difference between low and high self-esteem men. That is, there was 12% difference in the right, and a 13% difference in the left Hippocampal volumes between men with low and high self-esteem in the high MOCA group ($f_{(1, 19)}=3.049$ $p=.05$, Mixed design ANOVA (RSE x HC) (Figure 7). There were no significant effects to report in the Low MOCA group. There were no significant volumetric differences in MPFC volumes in either the High or Low MOCA groups.

Correlation Analysis

After rerunning the correlational analysis, we noticed that the association between DLPFC volume and Rosenberg self-esteem was still significant. However it was present only in women for both the right ($r=-.64$, $p=.04$) and partially for the left ($r=-.544$ $p=.07$) DLPFC. (See figure 8). That is, this association was observed only in women and was subsequently absent in the men. Therefore indicating that the women in our sample could have driven the initial association. This was accordingly substantiated in the case of the Internality subscale as well, where once again left ($r=-.626$, $p=.04$) and right ($r=-.613$, $p=.05$) DLPFC were associated with internality in women, but not in men ($p>.05$).

There were no significant associations to report for men or women with respect to MPFC volumes for both Rosenberg self-esteem scale and the internality subscale of the QCC.

For the hippocampus, we observed a robust association between right HC and Rosenberg self-esteem in the high MOCA group, which was present only in men ($r = .58$, $p = .03$) (see figure 9) but was completely absent in women ($p > .10$). With the internality subscale, the earlier association between right HC and internality was once again evident, but only in men ($r = .566$, $p = .06$). In the low MOCA group, the only significant association was between RSE and right HC ($r = .711$, $p = .02$) only in men. There was no significant relationship between hippocampal volume and self esteem in women in both high and Low MOCA groups. There was no significant relationship between total gray matter volume and self esteem, in neither men nor women for both MOCA groups.

To summarize once again, we found an association between both measures of self-esteem and DLPFC in women, only in the High MOCA group, and HC volumes and self esteem in men in both Maternal care groups.

Mediational analysis.

Since our initial statistical analysis revealed a strong association between DLPFC regions and self-esteem, we chose to include only the DLPFC volume for the subsequent Mediational analysis. In our first set of analysis using DLPFC as the dependent variable, and RSE and MOCA subsequently entered as independent variables, we found that there were no significant associations in either men or women ($p > .05$)

Using hippocampal volume as the dependent variable and RSE and MOCA subsequently entered as the independent variables, it was revealed that mother care contributed considerably to the relationship between self-esteem and HC ($F_{(1,22)} = 4.166$ $p = .03$) only in men, but not in women ($p > .05$)

VBM results

Preliminary VBM results indicated a trend towards lower self-esteem, higher grey matter density in the area of BA 10 or the frontal pole region ($t = -3.43$, $x = 27$, $y = 60$, $z = 18$) (see figure 10) in women. There were no significant grey matter density differences evident neither in the frontal lobes nor in the hippocampus for men.

Probabilistic maps (PB Maps) results

We wanted to investigate the reason for the lack of any significant associations between Prefrontal volume and self-esteem in the Low MOCA group. Hence the decision was made to examine if there were any possible shape discrepancies instead of volumetric scale measurements, which could be affected. Standardized segmenting protocols do not allow us to assess variability in shape and therefore we employed automated probabilistic maps method to identify any discrepancies. In the low MOCA group, specifically for women, we observed that the high self-esteem group had a more homogenous shape relative to the low self-esteem group (see figure 11a). There were no differences between high and low mother care women per se (see figure 11b).

Furthermore, we also chose to compare the high mother care, high self-esteem women with low mother care, high self-esteem women and we observed that once again the optimal case of high mother care and high self-esteem resulted in a shape that was much more homogenous as indicated by the presence of voxels with higher probability in this group (See figure 11c).

4. Discussion

Our aim was to investigate if variances in mother care could influence the relationship between self-esteem and three specific regions: the hippocampus in the medial temporal lobe, and the Dorsolateral and Medial prefrontal cortex in the frontal lobes. This of course assumes that a relationship exists between self-esteem and neural substrates and indeed in the case of the hippocampus this was shown to be the case (Pruessner et al 2005). Furthermore, a review of literature proved to reliably indicate that a similar relationship would be plausible in the frontal lobes as well. However, given the implicit association between mother care and self-esteem, an important question on whether mother care could influence this relationship was raised.

Our results revealed that indeed, mother care did contribute to the relationship between self-esteem and structural correlates. Most interestingly, this relationship was sex specific.

While these results are extremely exciting, they raise more questions; the most pertinent one is incontestably the presence of a sex difference in the first place. More specifically, that the relationship between self esteem and neural substrates was found for the DLPFC in women, but not in men; while for men but not women, that relationship was predominant in the HC. Another interesting question involves the negative association between self-esteem and DLPFC in high MOCA women, which implies that bigger is not necessarily better.

Lastly, the lack of any associations between the Medial prefrontal cortex and self esteem was certainly surprising, given the role that medial prefrontal cortex plays in self referencing and self monitoring of social information (Fossati et al., 2003; C. D. Frith & Frith, 1999; U. Frith & Frith, 2003; Ochsner et al., 2005) .

The Bi-regional association between men and women.

With reference to the first question about self esteem's bi-regional association in men and women, it has been observed that self esteem in general is different for men and women and this in itself could provide some clues about why different regions could be targeted.

Self esteem different for men and women

Men in general seem to base their levels of self-esteem on more achievement oriented goals, whereas women are more inclined to modulate their levels of self-esteem through more socially relevant means (Vohs & Heatherton, 2003).

Vohs et al (2003) contend that when men are in a socially threatened environment they are more likely to appear competitive and antagonistic. Incidentally, the authors also noticed that men with higher levels of self-esteem generally chose to highlight their unique competencies, while low self-esteem men were more inclined to emphasize their positive social qualities as a defense mechanism against the perceived insults to their self.

Conversely, high self-esteem women when threatened, focused more on qualities, which signified their value to a group whereas low self-esteem women chose to rely on their social affiliations as a validation of their worth. Additional evidence in women points to the fact that women are more likely than men to focus on social acceptance and therefore are more preoccupied by body image. The need to belong seems to be stronger in women relative to men, and thus ties into the concept that different “selves” may be relevant in self-esteem (Biro, Striegel-Moore, Franko, Padgett, & Bean, 2006).

Mother care contributes to sex difference in self-esteem

Interestingly enough, it has also been found that maternal control often directly contributes to the sex differences in self-evaluative development (Pomerantz, 1998). Mothers are more likely to inculcate the concept of socialization and social responsibility in their daughters and conversely give greater autonomy to their sons while mitigating societal constraints. This could be one of the mechanisms by which women develop a self-esteem that is more socially defined while men chose to define it through achievement-based goals (Vohs et al, 2003).

Moreover, since their primary avenue to gain maternal approval is through means where they get rewarded, this is what defines their value. This may drive them to adopt coping mechanisms, which facilitate this achievement-driven reward system. Intriguingly, the coping style of men is usually more sensory focused coping (Keogh & Herdendorf, 2002) which seeks to contextualize the problem with the least amount of abstraction. This form

of direct contextualization and information processing is the hallmark of the hippocampus and therefore the increasing dependence on the hippocampus seems more plausible (D. M. Smith & Mizumori, 2006; Somerville, Wig, Whalen, & Kelley, 2006).

On the other hand female self-esteem is much more multifaceted and seeks approval for a number of sources beyond the mother. It caters to societal expectations (Stroink, 2004) and depending on their levels of self-esteem will either invoke their belonging to their social group or will highlight facts that signal their valued worth in the group (Vohs et al 2003). This complicated form of informational processing complete with the requisite monitoring of social cues and emotional regulation (Schmitz et al., 2004), is indeed the domain of the DLPFC and therefore the impact of self-esteem in this context is once again not surprising.

Interestingly, one of the major consequences of the sex differences in self-esteem is that they also seem to have an impact on how men and women respond to stress. It has been demonstrated that men seem to respond more to stressors which the challenges are more achievement based in comparison to women who would report higher levels of stress when faced with paradigms in which they encountered social rejection (Stroud, Salovey, & Epel, 2002). Thus it highlights the need to examine the role of sex differences in greater detail, as it is apparent their influence is not limited to just differences in self-esteem, but can also have a significant impact on how men and women conduct their lives and cope in stressful situations.

Negative association between self-esteem and DLPFC volume in women: a possible explanation.

The negative association that we observed seems to suggest that that in women, lower levels of self-esteem may correspond with higher DLPFC volume. This could imply a number of things. Firstly, it could be indicative of abnormal or inefficient pruning. As mentioned earlier, prefrontal development undergoes exponential synaptic remodeling during puberty. Right before puberty there is an increase in grey matter volume, which then precipitates a post adolescence decline (Sowell et al., 2001). Grey Matter decline in the prefrontal cortex is exclusively achieved through the process of pruning that results in synaptic remodeling of existing networks. The maturing neural networks are extremely sensitive to environmental manipulations (Andersen, 2003; Teicher et al., 2003), so at this critical point, the trajectory of normal development can easily be compromised by the lack of a nourishing environment. Maternal care, especially during puberty could have a tremendous influence by ensuring that the child is exposed to an environment that encourages the proper trajectory of development. To give a drastic example of how important maternal care could be in dictating the environment, is a study by Gould et al (2005) which revealed that one of the reasons for adolescence suicide were problems with maternal figure of authority. If discrepancies in maternal care can have an influence on the mortality of an individual it is thus entirely possible that it could influence general brain development as well.

An alternate explanation may be that the individuals with lower self esteem are more prone to higher levels of self reflection and rumination and given the role of PFC in self-relevant information, it maybe a developmental safeguard that allows for allocation of more resources for vigilance required to assess a situation. For example, a study by Magnini et al (2004) demonstrated that the frontal and occipital lobes experienced an incremental change in neuronal density in parallel with the progression of hypertension in rats. Thus, along the same lines, a bigger DLPFC may be a necessary adaptation in order to better process the information to assign socially relevant contexts.

The shape of things.

However in case of women, this still does not explain why we only saw the association in the high MOCA group. Since we were relatively assured that there were no volumetric differences in our Low MOCA group, we chose to explore the possibility that perhaps along with the size, the shape too may be similarly affected. In addition, our exploratory VBM analysis had previously indicated that there was a gray matter discrepancy in all women with respect to their Rosenberg self esteem scores. Since VBM does not provide any further information with respect to the nature of the gray matter discrepancies, (Bookstein, 2001; Corbo et al., 2005) we wanted to investigate the possibility that we may encounter shape discrepancies which may perhaps be especially prominent in the LOW MOCA group.

Shape differences are not necessarily visible through contemporary volumetric assessment measures, thus we decided to adopt the fully automated probabilistic maps. What we discovered was an extremely interesting observation that in the low MOCA group those women who indicated higher levels of self-esteem had a relatively more homogenous structure than those whose levels of self-esteem was less than optimal.

At this point one can speculate that perhaps self-esteem may very well have a protective influence on the shape of the DLPFC. While proper maternal care may be instrumental in contributing to the robust development of gross size, having a high self-esteem level can at least insure that there is less variability in the shape. This is not surprising considering that puberty is a time where there is shift from parental authority to peer group (Nelson, Leibenluft, McClure, & Pine, 2005). Women are more sensitive to social pressures (Pine, 2002) and seem to be better equipped to negotiate in the changing social environment (Pomerantz, 1998). Therefore one can additionally surmise that those who manage to successfully make the cognitive set shift to friends develop a higher self-esteem are consequently better off than those whose inadequate mother care impedes on their ability to succeed in the social domain. Thus making it difficult for them to develop stable social networks, which could have a mitigating effect.

Nevertheless, we were still extremely keen to understand why the volumetric DLPFC association was only evident in the in high MOCA group, especially considering that the HC association was apparent in both groups. The answer perhaps lies once again in the fact that self-esteem it self could be profoundly affecting this relationship. Especially since we already know that it seems to have a different definition depending on the sex of the

individual. Self-esteem in men is primarily maintained through achievement based success (Vohs et al 2003) and the earliest source of validation for any success in most cases is the maternal figure (Pomerantz et al 1998). This suggests that perhaps, the link between self esteem and maternal care may be considerably more relevant in men than in women (Matsuoka et al., 2006). Our own Mediation analysis confirmed that maternal care contributed significantly in the variance in HC volume only in men. Thus perhaps providing a possible explanation as to why HC could be principally sensitive to both maternal care and self-esteem.

Women's, self-esteem on the other hand, is more socially oriented which provides them with more avenues to develop it independent of the mother's input. Plus in the case of the DLPFC, one can additionally surmise that perhaps the DLPFC is not as sensitive to maternal care as it is to self-esteem. The only way this could truly be validated is through a longitudinal study and that is unfortunately beyond the scope of this study. However it is urged to investigate how gender could significantly affect the relationship between mother care and self esteem.

Lack of MPFC association

The last question that we wanted to examine in detail was the unexpected absence of an effect in the MPFC. However, this could be an indication that the difference may be functional. Gusnard and Raichle (2001) constantly found that the MPFC often showed

increases and decreases specifically in areas BA 8, 9 and 10 (Gusnard, Akbudak et al., 2001). However, most of the increases have often been observed in studies where the experimental condition demanded monitoring or reporting one's own mental state or recollection of personal life events (Castelli, Happe, Frith, & Frith, 2000). The decreases were once again seen when the experimental task relied on cognitively challenging tasks. This prompted Frith et al (2003) to consider the medial prefrontal to be concerned primarily with "explicit representation of states of self".

This has been validated by various studies which have consistently found increases especially when the attention is specifically directed toward self-referential material (Fossati et al., 2004; Gusnard, Akbudak et al., 2001; Gusnard & Raichle, 2001; J. R. Simpson et al., 2000) . Moreover, it is also active during the so-called "theory of mind" i.e. the reading of the mental states of others. This inferring ability can also be translated into being aware of what the other person thinks about the self (C. Frith & Frith, 2005).

This gives an added confirmation that the MPFC may be linked to subtle functional differences rather than structural abnormalities and in our future studies we will definitely re-examine our initial hypothesis that the MPFC may be intrinsically associated with self-esteem.

5.Limitations and Future Directions

There are certain limitations in this study the primary being that we did not have any measure for depression. Given the higher incidence of depression in women (Piccinelli & Wilkinson, 2000) we suspect that particularly, in the low mother care group, and especially those individuals with low self-esteem, there could be additional confounds that could be contributing to the effects we observed.

Another aspect limitation is that our study hinges on the strength of two questionnaires; the PBI and the Rosenberg Self esteem scale. In case of the PBI specially, there have been many questions that have been raised about its validity and consistency. However, it has proven to be quite resistant to these allegations and in fact, it has shown robust test- retest reliability for extended periods up to 10 years (Parker et al 1990). It has also been quite resistant to assertions that it may be susceptible to individual mood states and personality variables. Indeed, Parker (1990) failed to find any significant impact of a response bias on the answers when they re-administered the test to the same set of individuals after a time lag. Perhaps the best example in the PBI immunity from current mood states would be that when it was administered to depressed individuals prior to, and after treatment, their response frequency remained unchanged (G. Parker, 1979). Thus implying that parental care recollections remained unaffected by mood or emotional regulation. Furthermore,

there is also evidence that it reflects true parenting styles rather than subjective assessments. When independently conducted in siblings, monozygotic and di-zygotic twins, there was a remarkable consistency of answers between siblings as well (Parker et al 1990).

The Rosenberg self-esteem scale is widely considered to be an extremely reliable measure of global self-esteem. In fact, when it administered to over 52 nations it showed a consistent cross cultural and internal validity (Schmitt & Allik, 2005). The theoretical midpoint of the scale is designated at $M=25.00$ (Rosenberg et al 1965). This being said, in our samples we seemed to have an unusually high mean of ($M = 32 \pm 5$) and this could potentially be a problem and our inflated scores could perhaps be artificially contributing to results. Fortunately, a review by Schmitt and Allik (2005) confirmed that in several countries including England, Switzerland and the United States of America the mean score for self-esteem tends be higher, especially in the 18-35 age bracket. For instance, the mean value for the United States is higher than 30 (Schmitt and Allik 2005). Therefore our mean seems to fit well within this schema.

With respect to having two dichotomous groups for mother care, we acknowledge that it would have perhaps afforded us a greater degree of precision if we had chosen to treat mother care as a continuous variable. However, our basic premise was to examine how mother care could influence the relationship between self-esteem and neural substrates, we believed that the best course of action would be to first examine the impact in two extreme groups.

Nonetheless, while we were very particular in terms of selecting people who fit within our to discrete mother care groups, we did not deem it necessary to also select them on the basis of their self-esteem scores as well. Given the variability of self-esteem scores in a normal population, we were relatively certain that we would get a normal distribution. In the future, however, more precaution will be taken to ensure that along with Mother care; self-esteem measures are also included in the specific recruitment criteria.

Perhaps another shortcoming of our study is that it is only a structural study and to complete the picture there needs to be a functional component through which structure and form can be seen in the context of functional neuronal changes. While we can speculate about the gross impact that mother care and consequently self-esteem can have, the story remain incomplete unless we can see behavioral manifestations than can be correlated with the prerequisite morphological aberrations. Therefore the next step would be a functional study, which examines the neural differences along with the underlying mechanistic changes.

An interesting direction would be to observe the circuitry changes rather than changes in individual structures. The hippocampus and prefrontal cortex are part of the emotional circuit (Gusnard et al., 2003; Posner & Rothbart, 1998; Vermetten & Bremner, 2002) and therefore changes in one region would undoubtedly influence the functioning of the other as well. Two possible avenues through which this could be plausible would be to utilize Diffusion Tensor Imaging (DTI) for white matter tracts and functional connectivity

analysis for investigating the functional involvement of the entire circuit. It stands to reason that changes in one could influence changes in other. Therefore, it is imperative that future studies see the system in its entirety.

While we are extremely excited about the unique nature of our result and the fact that this would be the first time an association between self esteem and the DLPFC will be reported, it also gives us reason to proceed with caution. The lack of any prior scientific evidence in the burgeoning field of social neuroscience makes it necessary for us to state that at this point our explanation should be treated as speculation and not fact.

Conclusions

To our knowledge, this is the first study to examine how mother care influences the relationship between Prefrontal volume and self-esteem in a sample of healthy young adults. In addition it is also the first study to successfully demonstrate a specific association between self-esteem and DLPFC volume. In addition, these associations are new in nature, as they seem to suggest that larger DLPFC volumes are associated with lower scores on the Rosenberg self-esteem scale, which is in contrast to what one would normally observe. This data is particularly intriguing considering that the correlations between DLPFC volume and low self-esteem were only found in women. In addition to the theoretical contribution this study also provided us with the opportunity to create two new templates, especially for the MPFC, which has as yet, never been attempted. Future fMRI data will examine functional significance of the associations found between self-esteem the

prefrontal cortex. This project, which has thus far embraced the structural complexities of self-esteem, could shed new light on the widespread impact of self-esteem on neural correlates and thus help us better understand a multifaceted trait which is a vulnerability factor for most, if not all chronic diseases and state of being.

6. Bibliography

- Akers, K. G., Nakazawa, M., Romeo, R. D., Connor, J. A., McEwen, B. S., & Tang, A. C. (2006). Early life modulators and predictors of adult synaptic plasticity. *Eur J Neurosci*, 24(2), 547-554.
- Andersen, S. L. (2003). Trajectories of brain development: point of vulnerability or window of opportunity? *Neurosci Biobehav Rev*, 27(1-2), 3-18.
- Ashburner, J., Andersson, J. L., & Friston, K. J. (2000). Image registration using a symmetric prior--in three dimensions. *Hum Brain Mapp*, 9(4), 212-225.
- Ashburner, J., & Friston, K. J. (1999). Nonlinear spatial normalization using basis functions. *Hum Brain Mapp*, 7(4), 254-266.
- Ashburner, J., & Friston, K. J. (2000). Voxel-based morphometry--the methods. *Neuroimage*, 11(6 Pt 1), 805-821.
- Ashburner, J., & Friston, K. J. (2005). Unified segmentation. *Neuroimage*, 26(3), 839-851.
- Barbas, H. (1995). Anatomic basis of cognitive-emotional interactions in the primate prefrontal cortex. *Neurosci Biobehav Rev*, 19(3), 499-510.
- Barker, D. J. (2004). The developmental origins of well-being. *Philos Trans R Soc Lond B Biol Sci*, 359(1449), 1359-1366.
- Baumeister, R. F., Campbell, J.K., Krueger, J.I., Vohs, K.D. (2003). Does high self esteem cause better performance, interpersonal success, happiness or healthier life styles? *Psychol. Sci. Public Interest*, 4, 1-44.
- Baumeister, R. F., & Leary, M. R. (1995). The need to belong: desire for interpersonal attachments as a fundamental human motivation. *Psychol Bull*, 117(3), 497-529.
- Beers, S. R., & De Bellis, M. D. (2002). Outcomes of child abuse. *Neurosurg Clin N Am*, 13(2), 235-241.
- Biro, F. M., Striegel-Moore, R. H., Franko, D. L., Padgett, J., & Bean, J. A. (2006). Self-esteem in adolescent females. *J Adolesc Health*, 39(4), 501-507.
- Block, J., & Robbins, R.W. (1993). A longitudinal study of consistency and change in the self esteem from early adolescence to early adulthood. *Child Development*(64), 909-923.
- Bookstein, F. L. (2001). "Voxel-based morphometry" should not be used with imperfectly registered images. *Neuroimage*, 14(6), 1454-1462.
- Bost, K. K., Shin, N., McBride, B. A., Brown, G. L., Vaughn, B. E., Coppola, G., et al. (2006). Maternal secure base scripts, children's attachment security, and mother-child narrative styles. *Attach Hum Dev*, 8(3), 241-260.
- Bowlby, J. (1969). *Attachment*. London: The Hogarth Press.
- Bredy, T. W., Grant, R. J., Champagne, D. L., & Meaney, M. J. (2003). Maternal care influences neuronal survival in the hippocampus of the rat. *Eur J Neurosci*, 18(10), 2903-2909.
- Bremner, J. D. (2002). Neuroimaging of childhood trauma. *Semin Clin Neuropsychiatry*, 7(2), 104-112.
- Buss, C., Lord, C., Wadiwalla, M., Hellhammer, D. H., Lupien, S. J., Meaney, M. J., et al. (2007). Maternal care modulates the relationship between prenatal risk and hippocampal volume in women but not in men. *J Neurosci*, 27(10), 2592-2595.

- Canetti, L., Bachar, E., Galili-Weisstub, E., De-Nour, A. K., & Shalev, A. Y. (1997). Parental bonding and mental health in adolescence. *Adolescence*, 32(126), 381-394.
- Carmichael, S. T., & Price, J. L. (1995). Limbic connections of the orbital and medial prefrontal cortex in macaque monkeys. *J Comp Neurol*, 363(4), 615-641.
- Castelli, F., Happe, F., Frith, U., & Frith, C. (2000). Movement and mind: a functional imaging study of perception and interpretation of complex intentional movement patterns. *Neuroimage*, 12(3), 314-325.
- Cicchetti, D., & Rogosch, F. A. (2001). The impact of child maltreatment and psychopathology on neuroendocrine functioning. *Dev Psychopathol*, 13(4), 783-804.
- Cicchetti, D., & Rogosch, F. A. (2002). A developmental psychopathology perspective on adolescence. *J Consult Clin Psychol*, 70(1), 6-20.
- Cicchetti, D., & Toth, S. L. (1995). A developmental psychopathology perspective on child abuse and neglect. *J Am Acad Child Adolesc Psychiatry*, 34(5), 541-565.
- Cicchetti, D., & Toth, S. L. (2000). Developmental processes in maltreated children. *Nebr Symp Motiv*, 46, 85-160.
- Clark, D. B., De Bellis, M. D., Lynch, K. G., Cornelius, J. R., & Martin, C. S. (2003). Physical and sexual abuse, depression and alcohol use disorders in adolescents: onsets and outcomes. *Drug Alcohol Depend*, 69(1), 51-60.
- Cocosco, C. A., Zijdenbos, A. P., & Evans, A. C. (2003). A fully automatic and robust brain MRI tissue classification method. *Med Image Anal*, 7(4), 513-527.
- Cole, D. A., Maxwell, S. E., Martin, J. M., Peeke, L. G., Seroczynski, A. D., Tram, J. M., et al. (2001). The development of multiple domains of child and adolescent self-concept: a cohort sequential longitudinal design. *Child Dev*, 72(6), 1723-1746.
- Collins, D. L., Neelin, P., Peters, T. M., & Evans, A. C. (1994). Automatic 3D intersubject registration of MR volumetric data in standardized Talairach space. *J Comput Assist Tomogr*, 18(2), 192-205.
- Corbo, V., Clement, M. H., Armony, J. L., Pruessner, J. C., & Brunet, A. (2005). Size versus shape differences: contrasting voxel-based and volumetric analyses of the anterior cingulate cortex in individuals with acute posttraumatic stress disorder. *Biol Psychiatry*, 58(2), 119-124.
- Costa, P. T., Jr., Busch, C. M., Zonderman, A. B., & McCrae, R. R. (1986). Correlations of MMPI factor scales with measures of the five factor model of personality. *J Pers Assess*, 50(4), 640-650.
- Crocker, J., & Wolfe, C. T. (2001). Contingencies of self-worth. *Psychol Rev*, 108(3), 593-623.
- D'Argembeau, A., Collette, F., Van der Linden, M., Laureys, S., Del Fiore, G., Degueldre, C., et al. (2005). Self-referential reflective activity and its relationship with rest: a PET study. *Neuroimage*, 25(2), 616-624.
- De Bellis, M. D. (2005). The psychobiology of neglect. *Child Maltreat*, 10(2), 150-172.
- De Bellis, M. D., Narasimhan, A., Thatcher, D. L., Keshavan, M. S., Soloff, P., & Clark, D. B. (2005). Prefrontal cortex, thalamus, and cerebellar volumes in adolescents and young adults with adolescent-onset alcohol use disorders and comorbid mental disorders. *Alcohol Clin Exp Res*, 29(9), 1590-1600.
- DeYoung, C. G., Peterson, J. B., & Higgins, D. M. (2005). Sources of openness/intellect: cognitive and neuropsychological correlates of the fifth factor of personality. *J Pers*, 73(4), 825-858.

- Domingo, M., Keppley, S., & Chambliss, C. (1997). Relations of early maternal employment and attachment in introvertive and extravertive adults. *Psychol Rep*, 81(2), 403-410.
- Eslinger, P. J., Flaherty-Craig, C. V., & Benton, A. L. (2004). Developmental outcomes after early prefrontal cortex damage. *Brain Cogn*, 55(1), 84-103.
- Eslinger, P. J., Grattan, L. M., Damasio, H., & Damasio, A. R. (1992). Developmental consequences of childhood frontal lobe damage. *Arch Neurol*, 49(7), 764-769.
- Fossati, P., Hevenor, S. J., Graham, S. J., Grady, C., Keightley, M. L., Craik, F., et al. (2003). In search of the emotional self: an fMRI study using positive and negative emotional words. *Am J Psychiatry*, 160(11), 1938-1945.
- Fossati, P., Hevenor, S. J., Lepage, M., Graham, S. J., Grady, C., Keightley, M. L., et al. (2004). Distributed self in episodic memory: neural correlates of successful retrieval of self-encoded positive and negative personality traits. *Neuroimage*, 22(4), 1596-1604.
- Friston, K. J., & Ashburner, J. (2004). Generative and recognition models for neuroanatomy. *Neuroimage*, 23(1), 21-24.
- Frith, C., & Frith, U. (2005). Theory of mind. *Curr Biol*, 15(17), R644-646.
- Frith, C. D., & Frith, U. (1999). Interacting minds--a biological basis. *Science*, 286(5445), 1692-1695.
- Frith, U., & Frith, C. D. (2003). Development and neurophysiology of mentalizing. *Philos Trans R Soc Lond B Biol Sci*, 358(1431), 459-473.
- Gallagher, I. I. (2000). Philosophical conceptions of the self: implications for cognitive science. *Trends Cogn Sci*, 4(1), 14-21.
- Gilbertson, M. W., Shenton, M. E., Ciszewski, A., Kasai, K., Lasko, N. B., Orr, S. P., et al. (2002). Smaller hippocampal volume predicts pathologic vulnerability to psychological trauma. *Nat Neurosci*, 5(11), 1242-1247.
- Glaser, D. (2000). Child abuse and neglect and the brain--a review. *J Child Psychol Psychiatry*, 41(1), 97-116.
- Gould, E., & Tanapat, P. (1999). Stress and hippocampal neurogenesis. *Biol Psychiatry*, 46(11), 1472-1479.
- Gould, M. S., Marrocco, F. A., Kleinman, M., Thomas, J. G., Mostkoff, K., Cote, J., et al. (2005). Evaluating iatrogenic risk of youth suicide screening programs: a randomized controlled trial. *Jama*, 293(13), 1635-1643.
- Greenough, W. T., Black, J. E., & Wallace, C. S. (1987). Experience and brain development. *Child Dev*, 58(3), 539-559.
- Gur, R. C., Alsop, D., Glahn, D., Petty, R., Swanson, C. L., Maldjian, J. A., et al. (2000). An fMRI study of sex differences in regional activation to a verbal and a spatial task. *Brain Lang*, 74(2), 157-170.
- Gusnard, D. A. (2005). Being a self: considerations from functional imaging. *Conscious Cogn*, 14(4), 679-697.
- Gusnard, D. A., Akbudak, E., Shulman, G. L., & Raichle, M. E. (2001). Medial prefrontal cortex and self-referential mental activity: relation to a default mode of brain function. *Proc Natl Acad Sci U S A*, 98(7), 4259-4264.
- Gusnard, D. A., Ollinger, J. M., Shulman, G. L., Cloninger, C. R., Price, J. L., Van Essen, D. C., et al. (2003). Persistence and brain circuitry. *Proc Natl Acad Sci U S A*, 100(6), 3479-3484.
- Gusnard, D. A., & Raichle, M. E. (2001). Searching for a baseline: functional imaging and the resting human brain. *Nat Rev Neurosci*, 2(10), 685-694.

- Gusnard, D. A., Raichle, M. E., & Raichle, M. E. (2001). Searching for a baseline: functional imaging and the resting human brain. *Nat Rev Neurosci*, 2(10), 685-694.
- Hall, L. A., Peden, A. R., Rayens, M. K., & Beebe, L. H. (2004). Parental bonding: a key factor for mental health of college women. *Issues Ment Health Nurs*, 25(3), 277-291.
- Hartner, S. (1999). *The Construction of the Self: A Developmental Perspective*. New York: Guildford.
- Heider, D., Matschinger, H., Bernert, S., Alonso, J., & Angermeyer, M. C. (2006). Relationship between parental bonding and mood disorder in six European countries. *Psychiatry Res*, 143(1), 89-98.
- Heinonen, K., Raikkonen, K., Keltikangas-Jarvinen. (2003). Maternal Perceptions and Adolescent self esteem: a six year Longitudinal Study. *Adolescence*, 38(152), 669-687.
- Hildyard, K. L., & Wolfe, D. A. (2002). Child neglect: developmental issues and outcomes. *Child Abuse Negl*, 26(6-7), 679-695.
- Hoyle, R., Kermis, M.H, Leary, M.R, Baldwin, M.W. (1999). *Selfhood: Identity ,Esteem, Control*. Westwood.
- James, W. (1890). *The Principles of Psychology*: Dover Publications.
- Johnson, C. K., Gilbert, M. D., & Herdt, G. H. (1979). Implications for adult roles from differential styles of mother-infant bonding: an ethological study. *J Nerv Ment Dis*, 167(1), 29-37.
- Keiley, M. K., Howe, T. R., Dodge, K. A., Bates, J. E., & Petti, G. S. (2001). The timing of child physical maltreatment: a cross-domain growth analysis of impact on adolescent externalizing and internalizing problems. *Dev Psychopathol*, 13(4), 891-912.
- Keogh, E., & Herdenfeldt, M. (2002). Gender, coping and the perception of pain. *Pain*, 97(3), 195-201.
- Krampen, G. (2001). *Fragebogen zu Kompetenz -und Kontrollueberzeugun-gen (FKK)*: Gottingen:Hogrefe.
- Leary, M. R. M., G. (2003). *Individual differences in Self Esteem*. New York: Guildford Press.
- LeCuyer-Maus, E. A. (2000). Maternal sensitivity and responsiveness, limit-setting style, and relationship history in the transition to toddlerhood. *Issues Compr Pediatr Nurs*, 23(2), 117-139.
- Leifer, M., Kilbane, T., & Skolnick, L. (2002). Relationships between maternal adult attachment security, child perceptions of maternal support, and maternal perceptions of child responses to sexual abuse. *J Child Sex Abus*, 11(3), 107-124.
- Liu, D., Diorio, J., Day, J. C., Francis, D. D., & Meaney, M. J. (2000). Maternal care, hippocampal synaptogenesis and cognitive development in rats. *Nat Neurosci*, 3(8), 799-806.
- Liu, D., Diorio, J., Tannenbaum, B., Caldji, C., Francis, D., Freedman, A., et al. (1997). Maternal care, hippocampal glucocorticoid receptors, and hypothalamic-pituitary-adrenal responses to stress. *Science*, 277(5332), 1659-1662.
- Matsuoka, N., Uji, M., Hiramura, H., Chen, Z., Shikai, N., Kishida, Y., et al. (2006). Adolescents' attachment style and early experiences: a gender difference. *Arch Womens Ment Health*, 9(1), 23-29.
- Mazziotta, J. C., Toga, A. W., Evans, A., Fox, P., & Lancaster, J. (1995). A probabilistic atlas of the human brain: theory and rationale for its development. The International Consortium for Brain Mapping (ICBM). *Neuroimage*, 2(2), 89-101.

- McEwen, B. S. (2003). Early life influences on life-long patterns of behavior and health. *Ment Retard Dev Disabil Res Rev*, 9(3), 149-154.
- McGarvey, E. L., Kryzhanovskaya, L. A., Koopman, C., Waite, D., & Canterbury, R. J. (1999). Incarcerated adolescents' distress and suicidality in relation to parental bonding styles. *Crisis*, 20(4), 164-170.
- Mead, G. H. (1934). *Mind, Self and Society from the standpoint of a social Behaviorist*. Chicago: University of Chicago Press.
- Mignini, F., Vitaoli, L., Sabbatini, M., Tomassoni, D., & Amenta, F. (2004). The cerebral cortex of spontaneously hypertensive rats: a quantitative microanatomical study. *Clin Exp Hypertens*, 26(4), 287-303.
- Miguel-Hidalgo, J. J., Wei, J., Andrew, M., Overholser, J. C., Jurjus, G., Stockmeier, C. A., et al. (2002). Glia pathology in the prefrontal cortex in alcohol dependence with and without depressive symptoms. *Biol Psychiatry*, 52(12), 1121-1133.
- Miller, E. K., & Cohen, J. D. (2001). An integrative theory of prefrontal cortex function. *Annu Rev Neurosci*, 24, 167-202.
- Mitelman, S. A., Buchsbaum, M. S., Brickman, A. M., & Shihabuddin, L. (2005). Cortical intercorrelations of frontal area volumes in schizophrenia. *Neuroimage*, 27(4), 753-770.
- Nelson, E. E., Leibenluft, E., McClure, E. B., & Pine, D. S. (2005). The social re-orientation of adolescence: a neuroscience perspective on the process and its relation to psychopathology. *Psychol Med*, 35(2), 163-174.
- Northoff, G., & Bermpohl, F. (2004). Cortical midline structures and the self. *Trends Cogn Sci*, 8(3), 102-107.
- Ochsner, K. N., Beer, J. S., Robertson, E. R., Cooper, J. C., Gabrieli, J. D., Kihlstrom, J. F., et al. (2005). The neural correlates of direct and reflected self-knowledge. *Neuroimage*, 28(4), 797-814.
- Parker, G. (1979). Parental deprivation and depression in a non-clinical group. *Aust N Z J Psychiatry*, 13(1), 51-56.
- Parker, G. (1990). The Parental Bonding Instrument. A decade of research. *Soc Psychiatry Psychiatr Epidemiol*, 25(6), 281-282.
- Parker, G., Tupling, H., Brown, L.D. (1979). A Parental Bonding Instrument. *British Journal of Medical Psychology*(52), 1-10.
- Pearlin, L. I., & Schooler, C. (1978). The structure of coping. *J Health Soc Behav*, 19(1), 2-21.
- Pelcovitz, D., Kaplan, S. J., DeRosa, R. R., Mandel, F. S., & Salzinger, S. (2000). Psychiatric disorders in adolescents exposed to domestic violence and physical abuse. *Am J Orthopsychiatry*, 70(3), 360-369.
- Pelham, B. W., & Swann, W. B., Jr. (1989). From self-conceptions to self-worth: on the sources and structure of global self-esteem. *J Pers Soc Psychol*, 57(4), 672-680.
- Petrides, M. (2000a). Dissociable roles of mid-dorsolateral prefrontal and anterior inferotemporal cortex in visual working memory. *J Neurosci*, 20(19), 7496-7503.
- Petrides, M. (2000b). The role of the mid-dorsolateral prefrontal cortex in working memory. *Exp Brain Res*, 133(1), 44-54.
- Petrides, M. (2005). Lateral prefrontal cortex: architectonic and functional organization. *Philos Trans R Soc Lond B Biol Sci*, 360(1456), 781-795.
- Petrides, M., & Pandya, D. N. (1999). Dorsolateral prefrontal cortex: comparative cytoarchitectonic analysis in the human and the macaque brain and corticocortical connection patterns. *Eur J Neurosci*, 11(3), 1011-1036.

- Piccinelli, M., & Wilkinson, G. (2000). Gender differences in depression. Critical review. *Br J Psychiatry*, 177, 486-492.
- Pine, D. S. (2002). Brain development and the onset of mood disorders. *Semin Clin Neuropsychiatry*, 7(4), 223-233.
- Pomenrantz, E. M., & Ruble, D.N. (1998). The Role of Maternal Control in the Development of Sex Differences in Child Self Evaluative Factors. *Child Development*, 69(2), 458-478.
- Posner, M. I., & Rothbart, M. K. (1998). Attention, self-regulation and consciousness. *Philos Trans R Soc Lond B Biol Sci*, 353(1377), 1915-1927.
- Prasad, K. M., Sahni, S. D., Rohm, B. R., & Keshavan, M. S. (2005). Dorsolateral prefrontal cortex morphology and short-term outcome in first-episode schizophrenia. *Psychiatry Res*, 140(2), 147-155.
- Pruessner, J. C., Li, L. M., Serles, W., Pruessner, M., Collins, D. L., Kabani, N., et al. (2000). Volumetry of hippocampus and amygdala with high-resolution MRI and three-dimensional analysis software: minimizing the discrepancies between laboratories. *Cereb Cortex*, 10(4), 433-442.
- Pyszczynski, T., Greenberg, J., Solomon, S., Arndt, J., & Schimel, J. (2004). Why do people need self-esteem? A theoretical and empirical review. *Psychol Bull*, 130(3), 435-468.
- Raichle, M. E., & Gusnard, D. A. (2005). Intrinsic brain activity sets the stage for expression of motivated behavior. *J Comp Neurol*, 493(1), 167-176.
- Rosenberg. (1965). *Society and the Adolescent self-image*: Princeton, NJ: Princeton University Press.
- Russek, L. G., & Schwartz, G. E. (1996). Narrative descriptions of parental love and caring predict health status in midlife: a 35-year follow-up of the Harvard Mastery of Stress Study. *Altern Ther Health Med*, 2(6), 55-62.
- Russek, L. G., & Schwartz, G. E. (1997a). Feelings of parental caring predict health status in midlife: a 35-year follow-up of the Harvard Mastery of Stress Study. *J Behav Med*, 20(1), 1-13.
- Russek, L. G., & Schwartz, G. E. (1997b). Perceptions of parental caring predict health status in midlife: a 35-year follow-up of the Harvard Mastery of Stress Study. *Psychosom Med*, 59(2), 144-149.
- Salzinger, S., Feldman, R. S., Ng-Mak, D. S., Mojica, E., & Stockhammer, T. F. (2001). The effect of physical abuse on children's social and affective status: a model of cognitive and behavioral processes explaining the association. *Dev Psychopathol*, 13(4), 805-825.
- Sato, T., Uehara, T., Narita, T., Sakado, K., & Fujii, Y. (2000). Parental bonding and personality in relation to a lifetime history of depression. *Psychiatry Clin Neurosci*, 54(2), 121-130.
- Schmitt, D. P., & Allik, J. (2005). Simultaneous administration of the Rosenberg Self-Esteem Scale in 53 nations: exploring the universal and culture-specific features of global self-esteem. *J Pers Soc Psychol*, 89(4), 623-642.
- Schmitz, T. W., Kawahara-Baccus, T. N., & Johnson, S. C. (2004). Metacognitive evaluation, self-relevance, and the right prefrontal cortex. *Neuroimage*, 22(2), 941-947.
- Simpson, J. R., Jr., Drevets, W. C., Snyder, A. Z., Gusnard, D. A., & Raichle, M. E. (2001). Emotion-induced changes in human medial prefrontal cortex: II. During anticipatory anxiety. *Proc Natl Acad Sci U S A*, 98(2), 688-693.

- Simpson, J. R., Jr., Snyder, A. Z., Gusnard, D. A., & Raichle, M. E. (2001). Emotion-induced changes in human medial prefrontal cortex: I. During cognitive task performance. *Proc Natl Acad Sci U S A*, 98(2), 683-687.
- Simpson, J. R., Ongur, D., Akbudak, E., Conturo, T. E., Ollinger, J. M., Snyder, A. Z., et al. (2000). The emotional modulation of cognitive processing: an fMRI study. *J Cogn Neurosci*, 12 Suppl 2, 157-170.
- Sled, J. G., Zijdenbos, A. P., & Evans, A. C. (1998). A nonparametric method for automatic correction of intensity nonuniformity in MRI data. *IEEE Trans Med Imaging*, 17(1), 87-97.
- Smith, D. M., & Mizumori, S. J. (2006). Hippocampal place cells, context, and episodic memory. *Hippocampus*, 16(9), 716-729.
- Smith, J., & Prior, M. (1995). Temperament and stress resilience in school-age children: a within-families study. *J Am Acad Child Adolesc Psychiatry*, 34(2), 168-179.
- Somerville, L. H., Wig, G. S., Whalen, P. J., & Kelley, W. M. (2006). Dissociable medial temporal lobe contributions to social memory. *J Cogn Neurosci*, 18(8), 1253-1265.
- Sowell, E. R., Thompson, P. M., Mattson, S. N., Tessner, K. D., Jernigan, T. L., Riley, E. P., et al. (2001). Voxel-based morphometric analyses of the brain in children and adolescents prenatally exposed to alcohol. *Neuroreport*, 12(3), 515-523.
- Stockmeier, C. A., Mahajan, G.J. et al. (2004). Cellular changes in postmortem hippocampus in major depression. *Biological Psychiatry*, 56(9), 640-650.
- Stroink, M. L. (2004). The conflicting standards dilemma and gender: a mediating model of its affective implications and coping styles. *J Soc Psychol*, 144(3), 273-292.
- Stroud, L. R., Salovey, P., & Epel, E. S. (2002). Sex differences in stress responses: social rejection versus achievement stress. *Biol Psychiatry*, 52(4), 318-327.
- Talairach, J., Tournoux, P., (1988). *Co-planer Stereotactic Atlas of the Human Brain; 3-Dimensional Proportional System: an approach to cerebral imaging*. New York: Thieme.
- Teicher, M. H. (2002). Scars that won't heal: the neurobiology of child abuse. *Sci Am*, 286(3), 68-75.
- Teicher, M. H., Andersen, S. L., Polcari, A., Anderson, C. M., & Navalta, C. P. (2002). Developmental neurobiology of childhood stress and trauma. *Psychiatr Clin North Am*, 25(2), 397-426, vii-viii.
- Teicher, M. H., Andersen, S. L., Polcari, A., Anderson, C. M., Navalta, C. P., & Kim, D. M. (2003). The neurobiological consequences of early stress and childhood maltreatment. *Neurosci Biobehav Rev*, 27(1-2), 33-44.
- Teicher, M. H., Ito, Y., Glod, C. A., Andersen, S. L., Dumont, N., & Ackerman, E. (1997). Preliminary evidence for abnormal cortical development in physically and sexually abused children using EEG coherence and MRI. *Ann N Y Acad Sci*, 821, 160-175.
- Teicher, M. H., Tomoda, A., & Andersen, S. L. (2006). Neurobiological consequences of early stress and childhood maltreatment: are results from human and animal studies comparable? *Ann N Y Acad Sci*, 1071, 313-323.
- Tisserand, D. J., Pruessner, J. C., Sanz Arigita, E. J., van Boxtel, M. P., Evans, A. C., Jolles, J., et al. (2002). Regional frontal cortical volumes decrease differentially in aging: an MRI study to compare volumetric approaches and voxel-based morphometry. *Neuroimage*, 17(2), 657-669.
- Uehara, T., Sato, T., Sakado, K., & Someya, T. (1998). Parental Bonding Instrument and the Inventory to Diagnose Depression Lifetime version in a volunteer sample of Japanese workers. *Depress Anxiety*, 8(2), 65-70.

- Vermetten, E., & Bremner, J. D. (2002). Circuits and systems in stress. I. Preclinical studies. *Depress Anxiety*, 15(3), 126-147.
- Vitaliano, P. P., Russo, J., Carr, J.E., Maiuro, R.D., Becker, J. (1985). The Ways of Coping Checklist: Revision and Psychometric Properties. *Multivariate Behavioral Research*, 20(1), 3-26.
- Vohs, K. D., & Heatherton, T. F. (2003). The effects of self-esteem and ego threat on interpersonal appraisals of men and women: a naturalistic study. *Pers Soc Psychol Bull*, 29(11), 1407-1420.
- Weaver, I. C., Meaney, M. J., & Szyf, M. (2006). Maternal care effects on the hippocampal transcriptome and anxiety-mediated behaviors in the offspring that are reversible in adulthood. *Proc Natl Acad Sci U S A*, 103(9), 3480-3485.
- Weber, D. A., & Reynolds, C. R. (2004). Clinical perspectives on neurobiological effects of psychological trauma. *Neuropsychol Rev*, 14(2), 115-129.
- Worsley, K. J., Taylor, J. E., Tomaiuolo, F., & Lerch, J. (2004). Unified univariate and multivariate random field theory. *Neuroimage*, 23 Suppl 1, S189-195.
- Zijdenbos, A. P., & Dawant, B. M. (1994). Brain segmentation and white matter lesion detection in MR images. *Crit Rev Biomed Eng*, 22(5-6), 401-465.

7. FIGURES

Figure 1: The Medial Prefrontal Cortex (MPFC) template

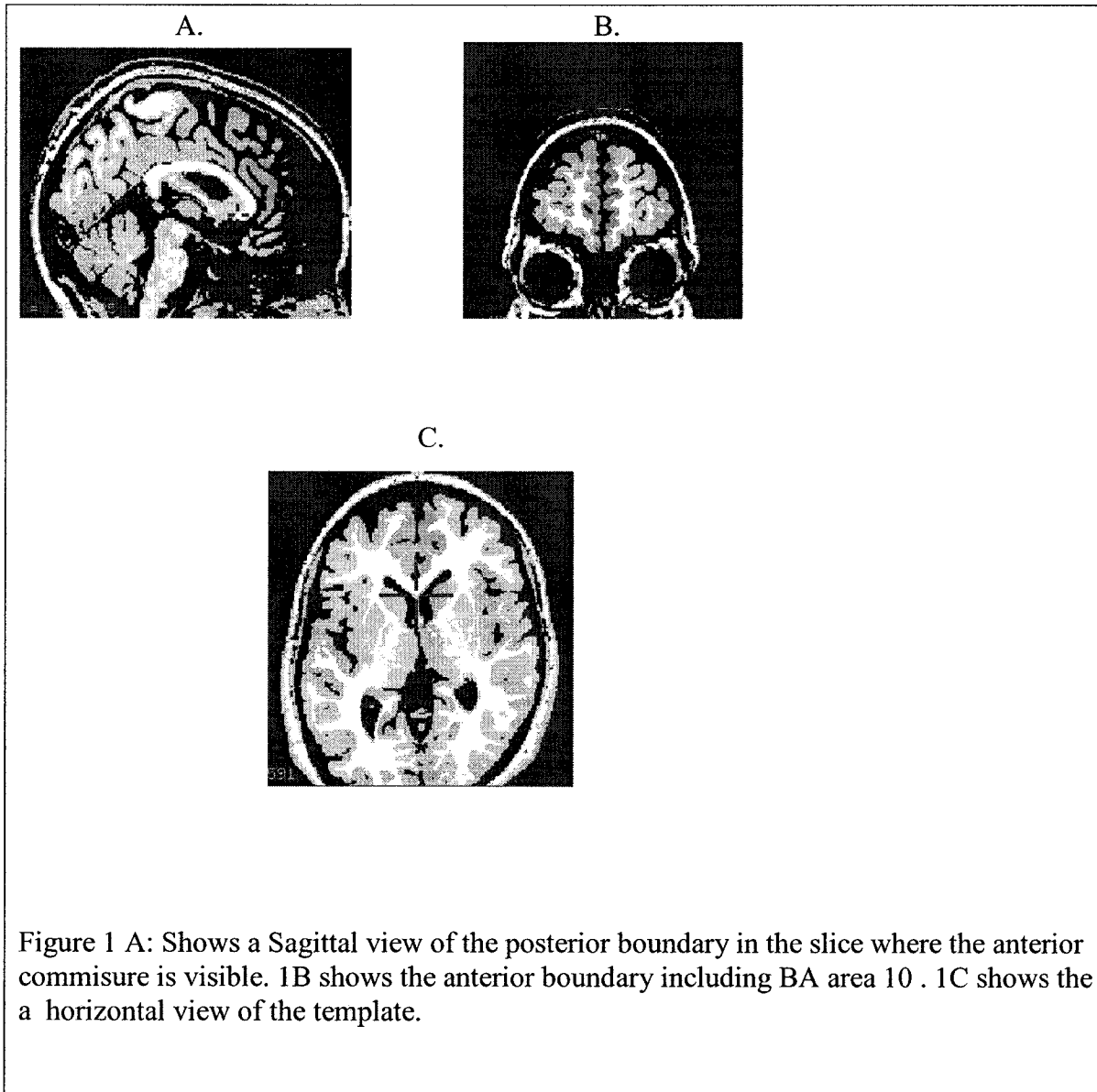


Figure 2: The Dorsolateral Prefrontal Cortex (DLPFC) Template.

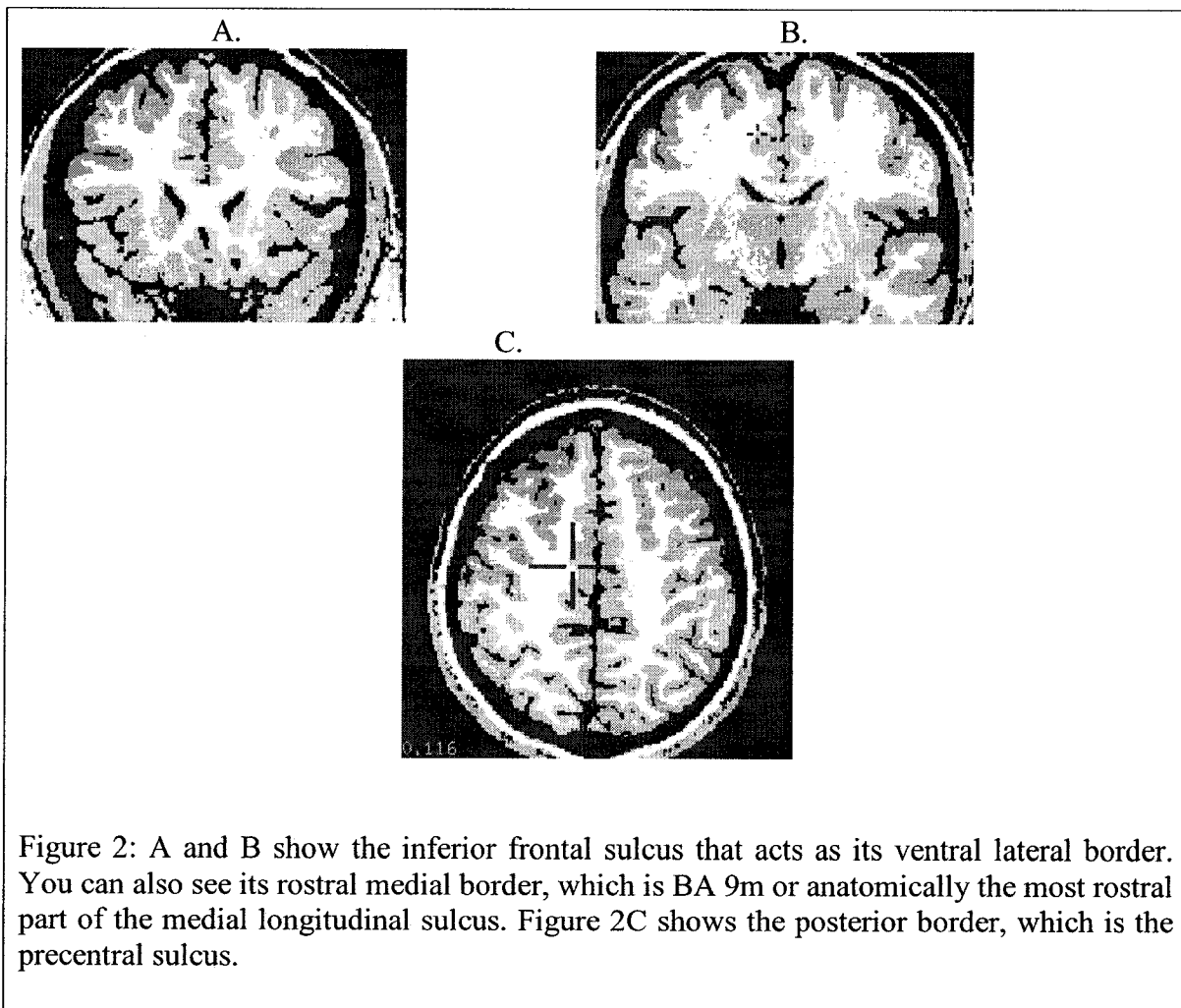


Figure 3: Volumetric Group Difference Between Low and High Mother care

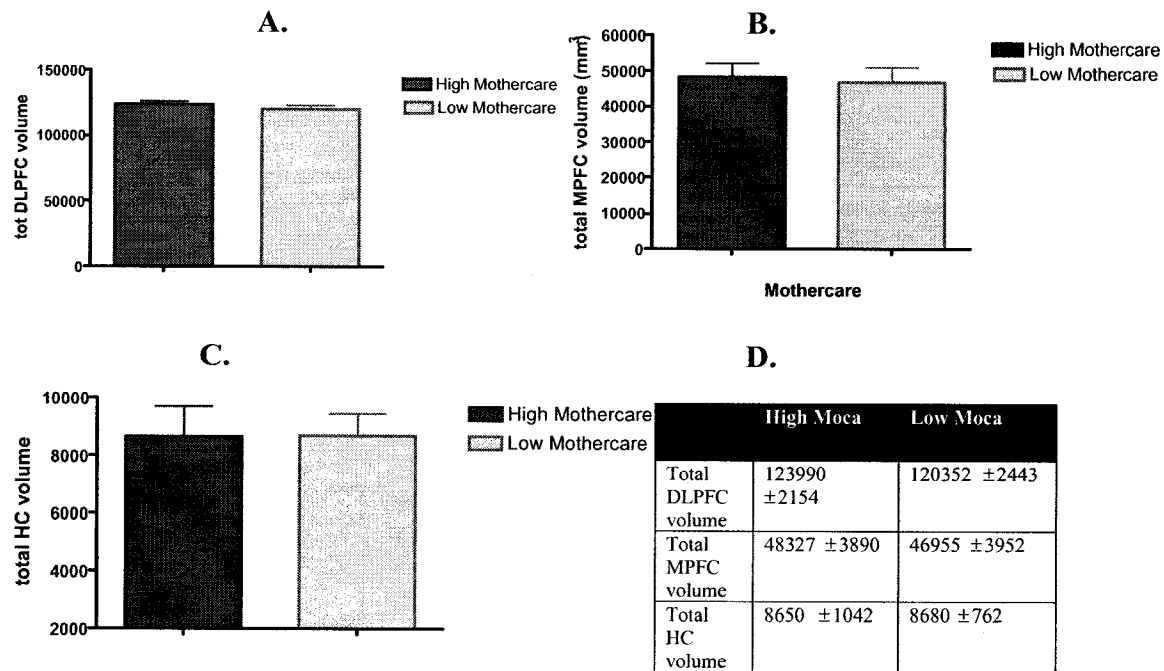


Figure 3: There were no volumetric differences observed between groups in (A): Total DLPFC, (B): total MPFC and (C): total HC volumes. D is the total mean volumes in both groups and the Standard deviation.

Figure 4: Associations between DLPFC and self esteem in high and low MOCA groups.

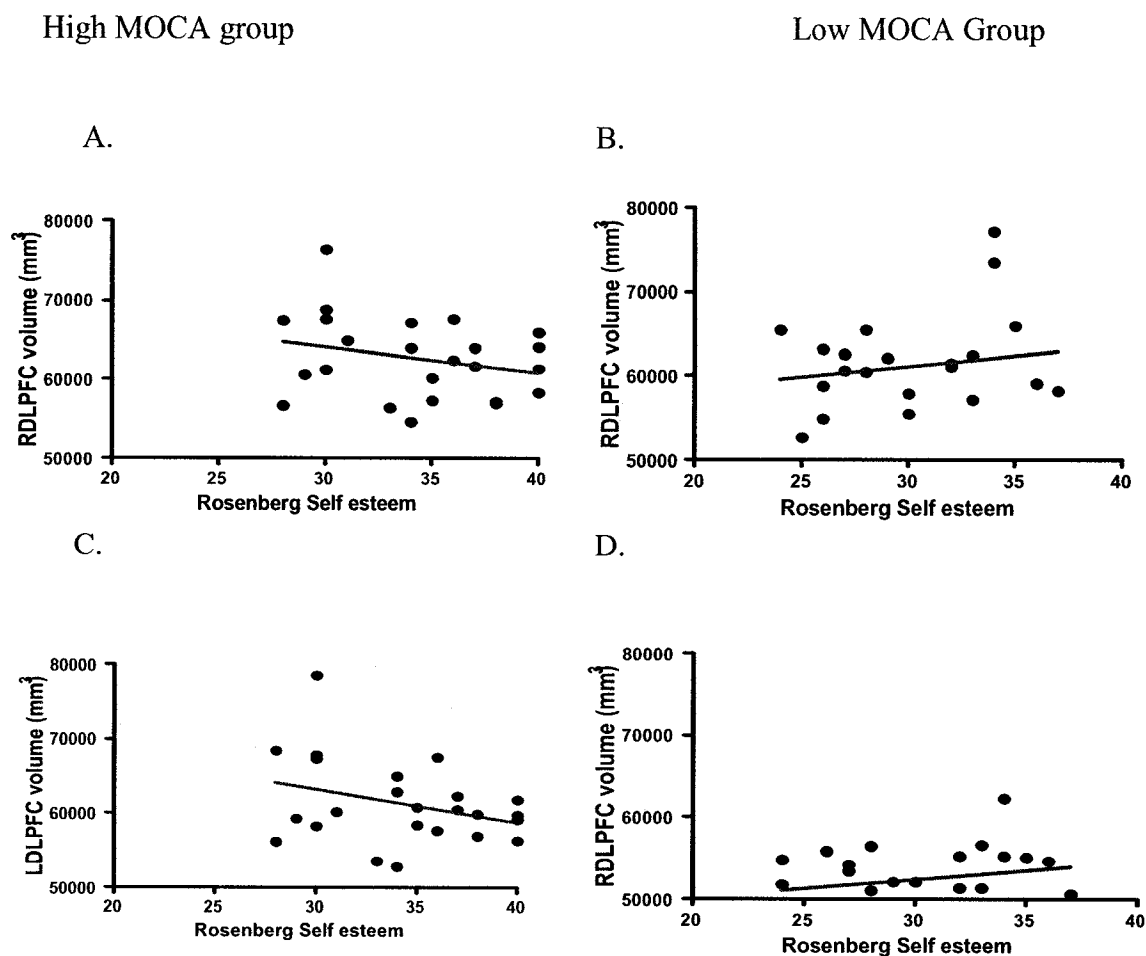


Figure 4: In the high MOCA group (A & C), there is an association between self esteem and DLPFC volume in both Right DLPFC (A) ($r = -.399$, $p < .05$) and left DLPFC (C) ($r = -.409$, $p < .05$). This association is not present in the Low MOCA group (B & D)

Figure 5: Association between HC volume and self esteem in High and Low MOCA groups

High MOCA group

Low MOCA group

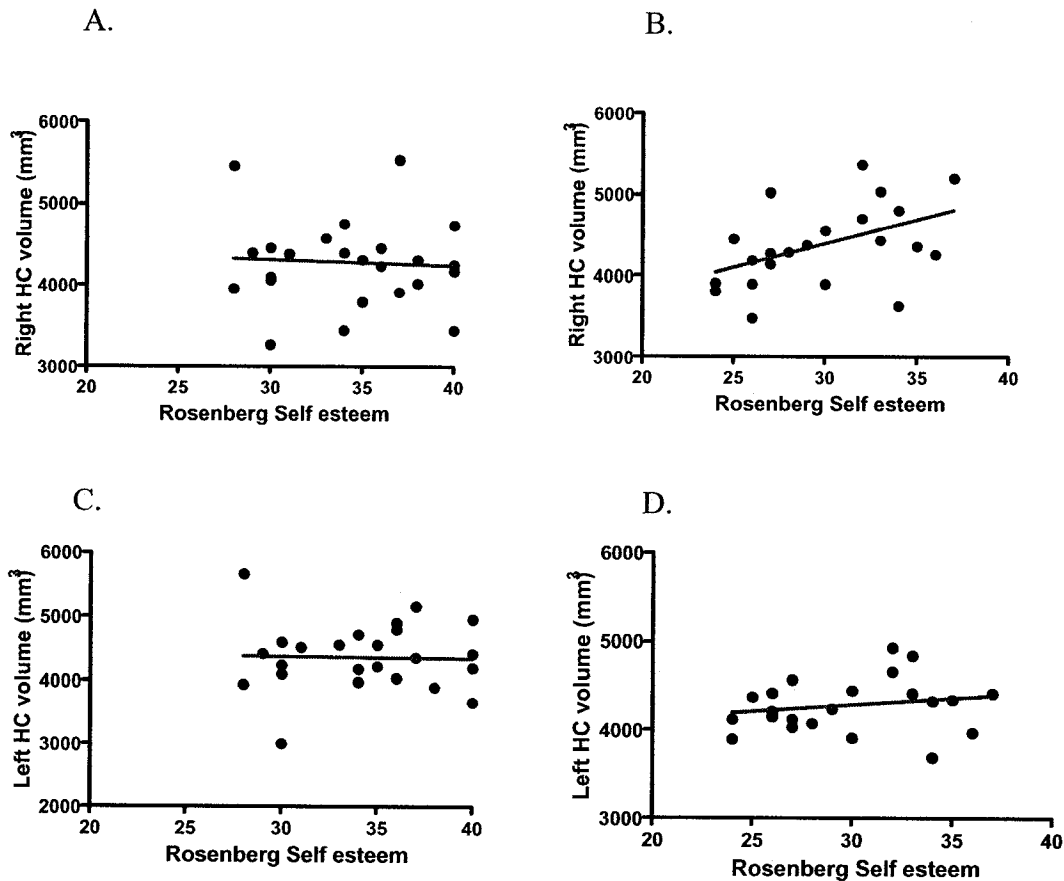


Figure 5: (A& C) are the associations between right and left HC and Rosenberg self esteem, in the high MOCA group. Both were non significant. In case of the Low MOCA group (B & D), there was a positive association observed in the Right HC = ($r=.486$, $p=.02$) (B). The association between the left HC and Rosenberg self esteem was insignificant (D).

Figure 6: Bar graph for DLPFC volumes as a function of self esteem and sex in High MOCA group

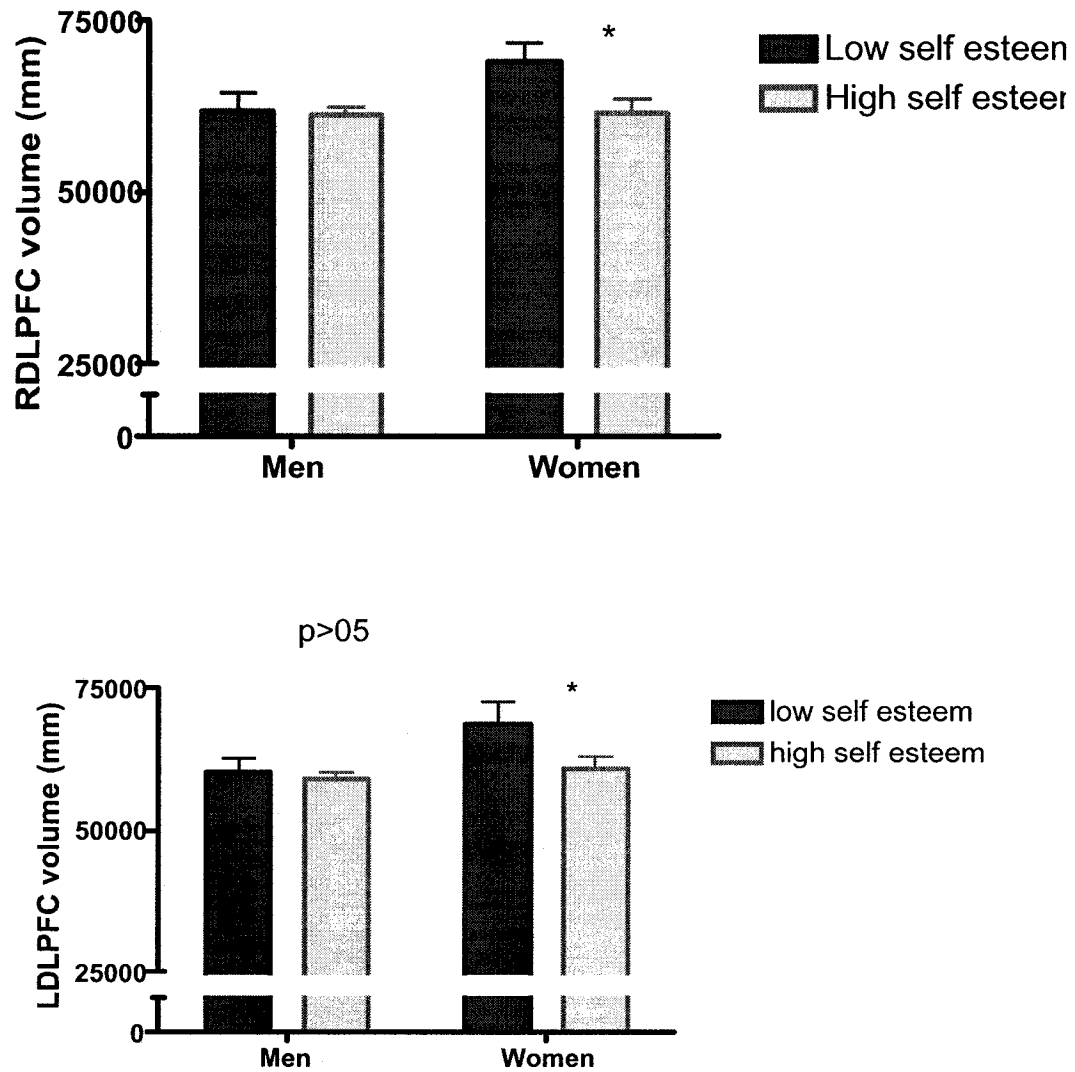


Figure 6: There was an 11% difference in the right DLPFC volume and 12% difference in the left DLPFC volume *only* in women as a function of self-esteem in the high MOCA group.

Figure 7: Bar Graph for HC volumes as a function of self esteem and sex in High MOCA group

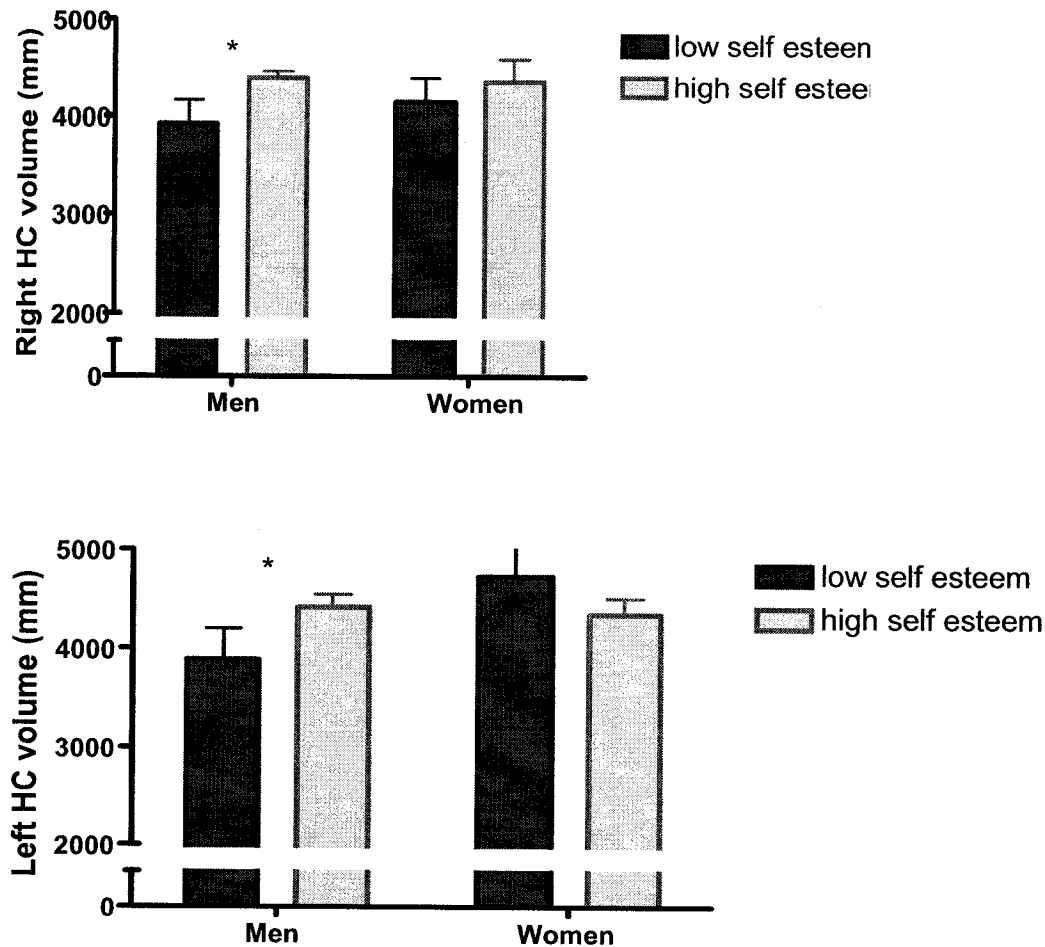


Figure 7: There was a 12% difference in the right DLPFC volume and 13% difference in the left HC volume *only* in men as a function of self-esteem in the high MOCA group.

Figure 8: Association between self esteem and DLPFC in men and women on low and high MOCA groups

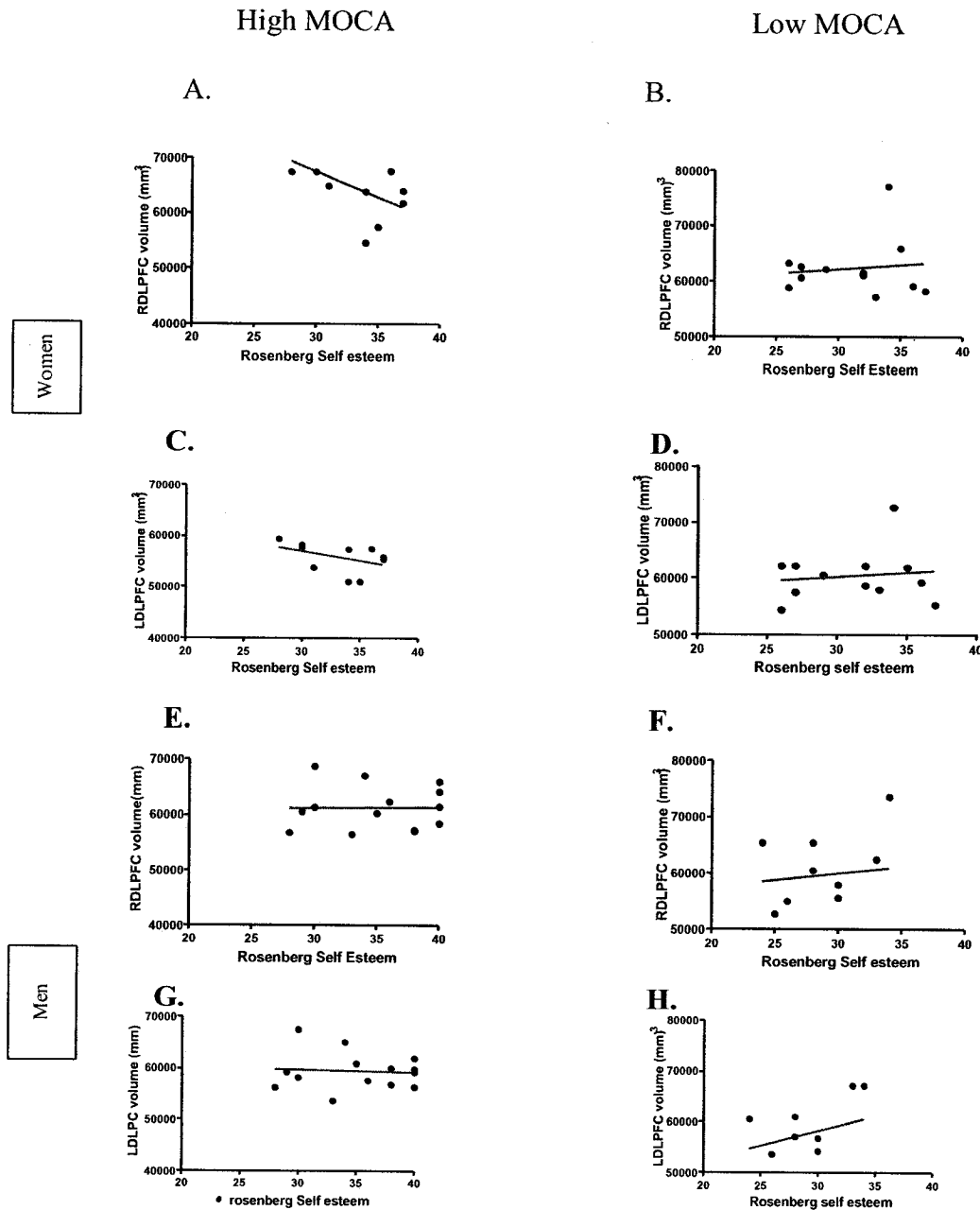


Figure 8: (A & C): In the high MOCA group there was a negative association between Rosenberg self esteem and right (A) ($r = -.644$, $p = .04$,) and partially in the left (C) DLPFC ($r = -.544$, $p = .07$). (B & D): There was no significant association between Rosenberg self-esteem and DLPFC volumes in Low MOCA women. (E-H): There were no significant associations to report in either high or low MOCA men.

Figure 9: Association between self esteem and HC in men and women on low and high MOCA groups

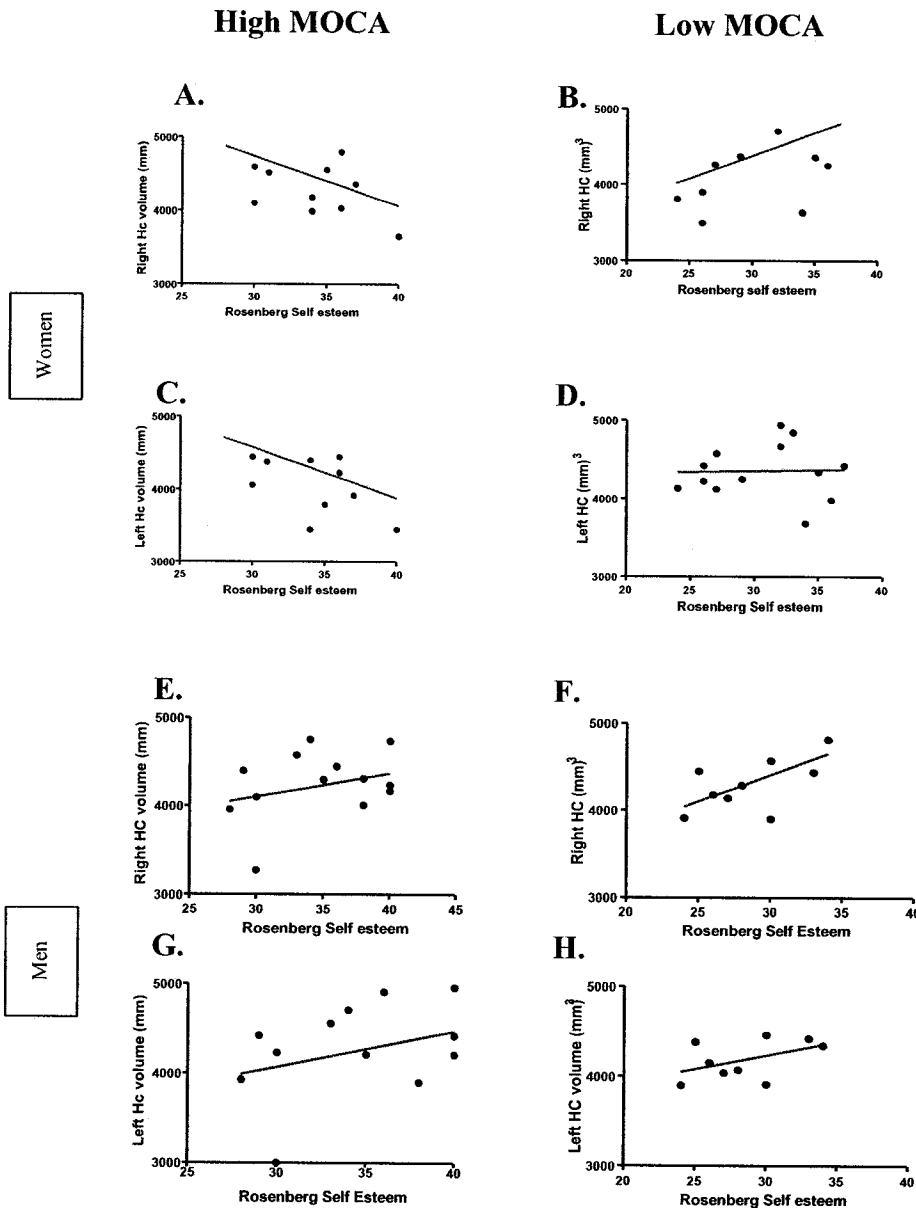
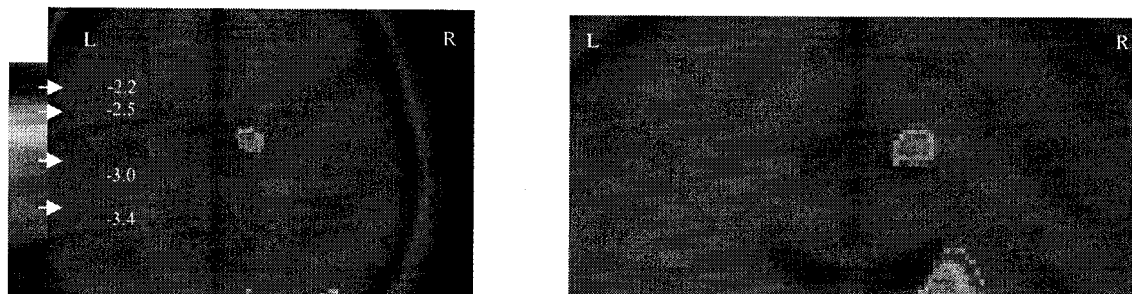


Figure 9: (A-D): represents women in high and low MOCA groups and there were no significant associations to report in either group. (E & G): show the correlation in High MOCA men between right(E) and left (G) hippocampus and Rosenberg self esteem. In the right there was a positive association reported ($r = .57, p = .04$) this association was non significant in the left ($p > .05$). (F & H) represent associations in low MOCA men. The only significant association was between right HC volume (F) and Rosenberg ($r = .71, p = .02$).

Figure 10: VBM results



Initial VBM results for women suggesting a possible link between self-esteem a prefrontal cortex at $t = -3.43$, $x=27, y=60, z=18$.

Figure 11: Probabilistic Maps

Figure11a. Probabilistic maps in the Low MOCA female groups.

Higher self-esteem women (left hand column) had relatively homogenous structure in comparison to low self esteem individuals. This is denoted by the color of the voxel , that is, the whiter the voxels the more homogenous the structure is, or in detail, there is a 100% probability that these voxels are present in all subjects. Thus the column of the left has a higher number of voxels which are white in comparison to the 2nd column which has a preponderance of green voxels which in turn suggests that the structure is homogenous in only 75% of all subjects.

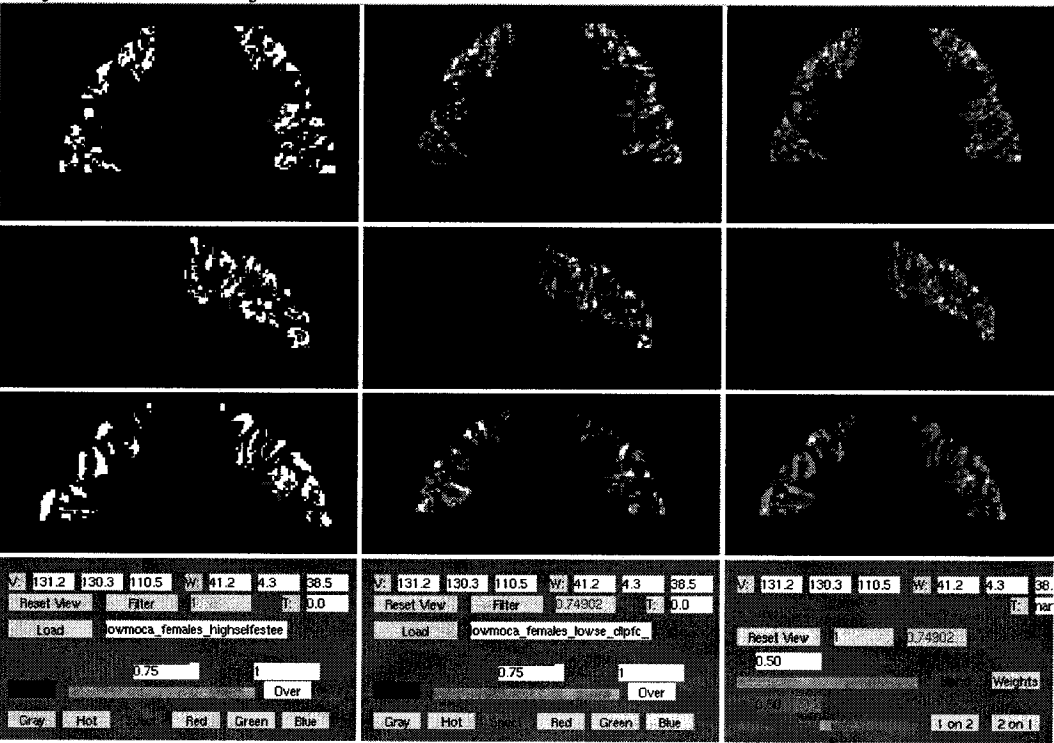
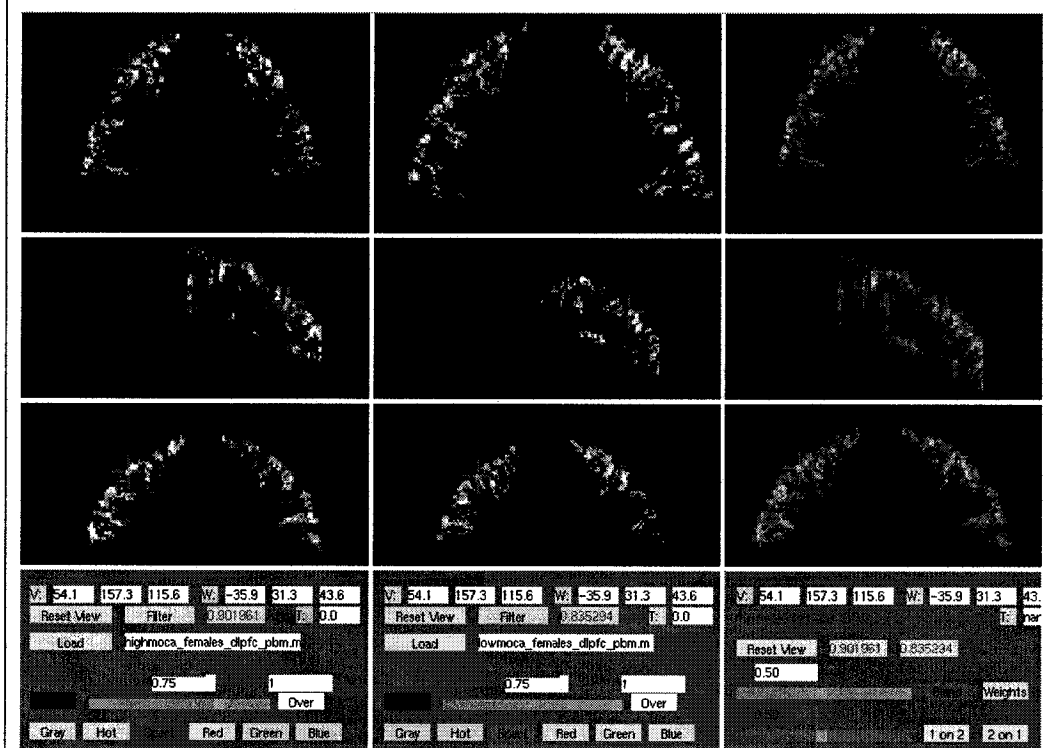


Figure 11b: Probabilistic maps for just Mother care in women.

At 75% probability for any voxel to be included in the structure, indicated that there was no difference in shape as function of low or high mother are alone.



At 75% confirmed that the optimal combination of high mother care high self esteem has more homogenous shape.



8. Tables

Table 1: Demographics for the whole group split by Mother care

	High MOCA	Low MOCA	Significance
Mother care	32 (3)	19 (6)	$T_{(49)} = 11.11$ $p < .001$
Age	22 (2)	24 (4)	$P > .05$
RSE	33 (6)	29 (3)	$T_{(49)} = 2.57$ $p < .05$
Internality	32 (4)	31 (4)	$P > .05$
	N= 27	N= 24	

Table 1: Indicates the mean scores with the standard deviation denoted by the parenthesis. The last column indicates that the two main variable of interest, that is mother care and Rosenberg self esteem were significantly different between our two groups.

Table 2: PBI scores on all four quadrants.

	High Mother Care group	Low Mother Care group	Statistical significance
Mother care	32 (3.2)	19(5.5)	$t_{(42)}=9.9$; $p=.000$
Mother over protection	10(5.8)	19 (6.6)	$t_{(42)}= -4.9$; $p= .000$
Father care	27(7.1)	20(7.1)	$t_{(41)}= 2.8$ $p = .000$
Father Overprotection	7.2 (4.1)	12(4.2)	$t_{(42)} = 3.5$ $p= .000$

Table 2: Indicates that along with mothercare, the two groups were also significantly different in the mean scores in the subscales of Mother over protection, father care and father overprotection, thus ensuring that we did indeed have two discreet groups. The Parenthesis denotes the standard deviation.

Table 3: Group demographics for men and women in high and low MOCA group

	High MOCA Males	Low MOCA Males	Stat Sig.	High MOCA Females	Low MOCA Females	Stat Sig.	Stat Sig. high moca males- females	Stat Sig. Low Moca males- females
Right HC(mm)	4255±389 N=13	4353±328 N=11	>.05	4331±647 N=11	4421±617 N=12	>.05	>.05	.05*
Left HC(mm)	4260±518 N=13	4203±213 N=10	>.05	4370±636 N=12	4356±350 N=13	>.05	>.05	>.05
LMPFC(mm)	24436±2090 N=14	23676±2337 N=10	>.05	24488±2070 N=11	23958±1741 N=13	>.05	>.05	>.05
RMPFC(mm)	23389±1992 N=14	22826±2121 N=10	>.05	24272±1833 N=11	23346±2213 N=13	>.05	>.05	>.05
LDLPFC(mm)	59468±3600 N=14	57709±6247 N=10	>.05	63271±7094 N=11	60400±4764 N=12	>.05	<.05*	>.05
RDLPFC(mm)	61275±3988 N=14	59800±7196 N=10	>.05	63939±6068 N=11	62324±5253 N=12	>.05	>.05	<.05*
PBI MOCA	31±3 N=14	19±6 N=11	<.001*	33±3 N=13	19±5 N=13	<.001*	<.05	>.05
RSE	35±5 N=14	29±4 N=11	<.001*	32±7 N=13	31±4 N=13	>.05	>.05	>.05
Age	22±2 N=14	25±4 N=11	<.05*	22±2 N=13	24±4 N=12	>.05	>.05	>.05

Table 3: is the demographics for the group split by mother care and sex. The statistical significance as ascertained by *t*-values is a comparison between high and low MOCA men and women and within group men and women on the neural substrates, age as well as the parameters of self-esteem and Mother care. As indicated, in the low MOCA group there is a difference between men and women in left HC volume. Similarly there is a difference in the mean volume of the Right DLPFC between low MOCA men and women as well. On the other hand, there seems to be gender difference between High MOCA men and women in the Left DLPFC. As expected, there is considerable difference between low and high mother care on both men and women, but no difference within the group is noticed.

9. Appendices

1. compliance certificate

2. Questionnaires used.

2a: Parental Bonding index (Parker et al 1979)

2b: Rosenberg self esteem (Rosenberg 1965)

2c: Questionnaire for competency and control (Krampen 2001)

PBI

This questionnaire lists various attitudes and behaviours of parents. As you remember your **mother** in the **first 16 years of your life**, please mark the most appropriate response to each statement.

		Very true	Rather true	Rather untrue	Very untrue
1	Spoke to me with a warm and friendly voice.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
2	Did not help me as much as I needed.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
3	Let me do those things that I liked to do.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
4	Seemed emotionally cold to me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
5	Appeared to understand my problems and worries.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
6	Was affectionate to me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
7	Wanted me to make my own decisions.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
8	Did not want me to grow up.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
9	Tried to control everything I did.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
10	Invaded my privacy.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
11	Enjoyed talking things over with me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
12	Frequently smiled at me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
13	Tended to baby me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
14	Did not seem to understand what I needed or wanted.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
15	Let me decide things for myself.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
16	Made me feel I wasn't wanted.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
17	Could make me feel better when I was upset.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
18	Did not talk to me very much.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
19	Tried to make me dependent on her.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
20	Felt I could not look after my-self unless she was around.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
21	Gave me as much freedom as I wanted.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
22	Was overprotective of me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
23	Did not praise me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
24	Let me dress in any way I wanted.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

As you remember your **father** in the **first 16 years of your life**, please mark the most appropriate response to each statement.

		Very true	Rather true	Rather untrue	Very untrue
25	Spoke to me with a warm and friendly voice.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
26	Did not help me as much as I needed.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
27	Let me do those things that I liked to do.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
28	Seemed emotionally cold to me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
29	Appeared to understand my problems and worries.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
30	Was affectionate to me	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
31	Wanted me to make my own decisions.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
32	Did not want me to grow up.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
33	Tried to control everything I did.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
34	Invaded my privacy.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
35	Enjoyed talking things over with me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
36	Frequently smiled at me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
37	Tended to baby me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
38	Did not seem to understand what I needed or wanted.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
39	Let me decide things for my-self.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
40	Made me feel I wasn't want-ed.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
41	Could make me feel better when I was upset.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
42	Did not talk to me very much.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
43	Tried to make me dependent on him.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
44	Felt I could not look after my-self unless he was around.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
45	Gave me as much freedom as I wanted.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
46	Was overprotective on me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
47	Did not praise me.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
48	Let me dress in any way I wanted.	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

ROSENBERG SCALE OF SE

Below is a list of statements dealing with your general feelings about yourself. If you **STRONGLY AGREE**, circle 1. If you **AGREE** with the statement, circle 2. If you **DISAGREE**, circle 3. If you **STRONGLY DISAGREE**, circle 4.

		1 Strongly Agree	2 Agree	3 Disagree	4 Strongly Disagree
1	I feel that I am a person of worth, at least on an equal plane with others.	1	2	3	4
2	I feel that I have a number of qualities.	1	2	3	4
3	All in all, I am inclined to feel that I am a failure.	1	2	3	4
4	I am able to do things as well as other people.	1	2	3	4
5	I feel I do not have much to be proud of.	1	2	3	4
6	I take a positive attitude towards myself.	1	2	3	4
7	On the whole, I am satisfied with myself.	1	2	3	4
8	I wish I could have more respect for myself.	1	2	3	4
9	I certainly feel useless at times.	1	2	3	4
10	At times I think I am no good at all.	1	2	3	4

Subject: _____

Date: _____

Questionnaire of Competence and Control

The following thirty-two questions deal with the way you perceive situations and events. Please answer each question by deciding whether the statement is

- 1 - totally wrong
- 2 - wrong
- 3 - more wrong than right
- 4 - more right than wrong
- 5 - right
- 6 - totally right

and circle the appropriate number next to the question. Please answer all questions, even if some of them will appear very similar to you.

-
- | | | | | | | |
|--|---|---|---|---|---|---|
| 1. It mainly depends on me whether other people do what I want. | 1 | 2 | 3 | 4 | 5 | 6 |
| 2. A good proportion of my life is determined by coincidental events. | 1 | 2 | 3 | 4 | 5 | 6 |
| 3. I feel that a lot of things in my life depend on other people. | 1 | 2 | 3 | 4 | 5 | 6 |
| 4. Sometimes I feel passive and not creative. | 1 | 2 | 3 | 4 | 5 | 6 |
| 5. Whether I have an accident only depends on me and my behavior. | 1 | 2 | 3 | 4 | 5 | 6 |
| 6. When I make plans I am sure that they will come true. | 1 | 2 | 3 | 4 | 5 | 6 |
| 7. I often don't have the possibilities to protect myself from misfortune. | 1 | 2 | 3 | 4 | 5 | 6 |
| 8. I often do not know what to do in ambiguous situations. | 1 | 2 | 3 | 4 | 5 | 6 |

- | | | | | | | |
|--|---|---|---|---|---|---|
| 9. When wishes come true,
good fortune plays a role. | 1 | 2 | 3 | 4 | 5 | 6 |
| 10. Other people often prevent the
fulfillment of my plans. | 1 | 2 | 3 | 4 | 5 | 6 |
| 11. I have ways to protect myself from getting ill. | 1 | 2 | 3 | 4 | 5 | 6 |
| 12. I often do not know how to make
my dreams come true. | 1 | 2 | 3 | 4 | 5 | 6 |
| 13. A lot of things in my life
happen by chance. | 1 | 2 | 3 | 4 | 5 | 6 |
| 14. Other people influence my
everyday life in many ways. | 1 | 2 | 3 | 4 | 5 | 6 |
| 15. Whether I have an accident
depends on how fortunate I am. | 1 | 2 | 3 | 4 | 5 | 6 |
| 16. I know a lot of ways to protect
myself from illnesses. | 1 | 2 | 3 | 4 | 5 | 6 |
| 17. I only have little possibility to
defend my interests against others. | 1 | 2 | 3 | 4 | 5 | 6 |
| 18. It is not a good idea to plan things
in advance because of destiny. | 1 | 2 | 3 | 4 | 5 | 6 |
| 19. Being nice and friendly
helps in being successful. | 1 | 2 | 3 | 4 | 5 | 6 |
| 20. I always know what to
do when I am in danger. | 1 | 2 | 3 | 4 | 5 | 6 |
| 21. It is pure coincidence if
other people do what I want. | 1 | 2 | 3 | 4 | 5 | 6 |
| 22. Other people impose on how I feel. | 1 | 2 | 3 | 4 | 5 | 6 |
| 23. I can determine a lot of things
that happen in my life. | 1 | 2 | 3 | 4 | 5 | 6 |
| 24. I sometimes feel very helpless
and don't know what to do. | 1 | 2 | 3 | 4 | 5 | 6 |

- | | | | | | | |
|--|---|---|---|---|---|---|
| 25. I usually follow my interests
and reach my goals. | 1 | 2 | 3 | 4 | 5 | 6 |
| 26. Other people determine
whether I have an accident or not. | 1 | 2 | 3 | 4 | 5 | 6 |
| 27. Reaching my goals is a consequence
of my personal effort and ambition. | 1 | 2 | 3 | 4 | 5 | 6 |
| 28. Even in difficult situations I know what to do. | 1 | 2 | 3 | 4 | 5 | 6 |
| 29. In order to have a greater chance
of reaching my goals, I adjust them
to the wishes of other people. | 1 | 2 | 3 | 4 | 5 | 6 |
| 30. My life is determined entirely
by me and my wishes. | 1 | 2 | 3 | 4 | 5 | 6 |
| 31. Destiny determines whether I get sick. | 1 | 2 | 3 | 4 | 5 | 6 |
| 32. I always find ways for solving my problems. | 1 | 2 | 3 | 4 | 5 | 6 |
-

Mehereen Wadiwalla (2007) The Influence of Mother Care on the relationship between Self-esteem and Neural Substrates in young men and women: A Neuroimaging Study.