



Aromatherapy and massage for antenatal anxiety: Its effect on the fetus

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KEYWORDS

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Summary Antenatal anxiety has been linked to maternal hypothalamic–pituitary–adrenal axis changes which can affect fetal development and may have lasting effects on the child's psychological development. Treatments for anxiety have hitherto focused on psychotherapy techniques or antidepressant drugs but these do not always effect long term improvement. Aromatherapy and massage have successfully been used to produce significantly greater improvement in reduction of anxiety. Midwives may highlight anxiety in some of the mothers in their care and can incorporate the holistic approach of aromatherapy and massage into their practice. However, further research is required to establish the efficacy and cost-effectiveness of aromatherapy and massage in the antenatal period.

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Introduction

A degree of anxiety during pregnancy is normal for most women, particularly in the first trimester,¹ as they adjust to its physical and psychosocial impact. As pregnancy progresses altered body image can increase anxiety,² and external factors, such as stress at work, financial, social or relationship difficulties may also contribute.^{3–6}

Anxiety is a complex phenomenon which may increase between 18 and 32 weeks gestation, a period which coincides with various antenatal screening tests,⁷ and can easily be exacerbated by an over-zealously supportive professional struc-

ture and excessive medicalisation.^{8–10} Antenatal testing for fetal abnormalities can increase anxiety in pregnancy and if women are confronted with a positive malformation scan the levels of anxiety are unchanged.¹¹ Studies have confirmed that women's fears of pregnancy, childbirth, previous loss of a baby, lack of support, financial and unrealistic work expectations manifest as symptoms of anxiety.^{5,12,13} It is important that midwives allow women time to express their anxieties, as pressures of a busy clinic can create constraints and essential information missed.^{14–16}

Identifying, a reliable screening anxiety scale could help the midwife to clarify the nature and relationship of the anxiety, but many screening tools are not suitable for pregnancy.¹⁷ Scales to measure antenatal anxiety have been validated,

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but in order to obtain an adequate evaluation of anxiety; it may be necessary to measure the influences of psychological and body changes at different stages of pregnancy.¹⁸⁻²² It is important that these anxiety scales are interpreted correctly, as professionals may see that the benefits of antidepressants outweigh the risks in pregnancy and could encourage their use in late pregnancy.²³ Although the risks of teratogenesis is low, there is no conclusive evidence on the safety of antidepressants in late pregnancy.^{24,25}

Parent education classes provide opportunities for women to express their fears and worries and these opportunities can go some way towards alleviating the severity of anxiety,²⁶⁻²⁸ although classes may have little effect on those more severely affected.⁴ Treatments for anxiety principally focus on psychotherapy techniques, but these do not always result in long term improvement and have resourcing implications, as they are facilitated by trained professionals.²⁹

Bio-psycho-social effects of anxiety on the fetus

Anxiety is a complex phenomenon but the current clinical definition is as a psychosocial condition in which worry, fear, and apprehension are combined with physical symptoms (Box 1,³⁰). A mother's response to anxiety is determined by the hypothalamus, which acts on the autonomic nervous system and the endocrine system resulting in physiological changes.³¹ Prolonged anxiety (more than three weeks and up to six months' duration) without

Box 1 Symptoms of anxiety.

- Palpitations, accelerated heart rate
- Trembling or shaking
- Difficulty in breathing
- Chest pain or discomfort
- Feeling dizzy, faint, light headed
- Fear of losing control, passing out
- Sweating
- Dry mouth
- Feeling of choking
- Nausea or abdominal discomfort
- Feeling that objects are unreal
- Fear of dying
- Numbness or tingling sensations
- Hot flushes or cold chills³⁰

intervention may result in a maladaptive response to pregnancy.³²

Recent studies have attempted to disentangle the different environmental and genetic factors, and to link specific exposures to specific effects.³³⁻³⁵ Antenatal anxiety has been linked with changes to the maternal hypothalamus-pituitary-adrenal axis (HPA). During pregnancy the corticotrophin releasing hormone (CRH) increases substantially before delivery, with placental CRH relating to the duration of pregnancy.^{36,37} Antenatal anxiety increases CRH and has been associated with premature labour.^{38,39} Mancuso et al.⁴⁰ demonstrated that women, who delivered prematurely had significantly higher levels of CRH compared with women who delivered at term and may predispose the mother to spontaneous abortion.⁴¹ Gitau et al.⁴² found a potential link in pregnant women between clinically-indicated fetal blood sampling and acute fetal distress, suggesting that the increase in CRH derives from the placenta rather than the fetus in response to stress.

It is increasingly apparent that antenatal anxiety can have lasting effects on the psychological development of the child.⁴³ The changes in the functioning of the HPA axis are linked to antenatal anxiety and the subsequent disturbances in offspring account partly for cognitive and neurological development.⁴⁴ In a cohort study of 10,000 pregnant women, O'Connor et al.⁴⁵ identified a link between maternal anxiety in the third trimester and behavioural/emotional problems in the resulting children at four years. The researchers found that self-reported antenatal anxiety at 32 weeks predicted severe behavioural/emotional problems in both sexes, but anxiety in late gestation appears to be significantly associated with hyperactivity/inattention in boys.⁴⁵ Mixed handedness, dyslexia and autism may also result.⁴⁶

It is interesting to note that anxiety occurring at different gestations produces variable fetal effects.⁴⁷ For example, severe life events in the first trimester has been shown to result in a 50% increase in neural tube abnormalities⁴⁸ and other anomalies, such as cleft palate.⁴⁹ In contrast, anxiety in late pregnancy, when neural connections are being made in the brain, produces behavioural/emotional outcomes.⁴⁶

Massage and aromatherapy for anxiety in pregnancy

It is known that massage improves self-image during pregnancy, and may aid acceptance of

physical changes in the mother.⁵⁰ The well-researched relaxation effects also extend to the fetus.^{51,52} Massage stimulates production of endorphins and decreases blood pressure, through its effect on the parasympathetic nervous system,^{53,54} which may reduce the severity and prognosis of pregnancy-induced hypertension^{55,56} and potential maternal and fetal morbidity and mortality. With these claims of transiently reducing blood pressure midwives should be mindful of supine hypotension and the woman's position when performing massage.⁵⁷ Massage has been found to reduce anxiety in labour⁵⁸ and may be more effective than breath coaching,⁵⁹ although it is difficult to extrapolate from these studies whether the reduction in anxiety arose from the massage or the use of essential oils.^{60,61}

Massage facilitates absorption of essential oils via the skin,⁶² and aromatherapy is one of the primary complementary therapies used for the treatment of anxiety.⁶³ Although essential oils are known to cross the placental barrier and can be transferred to the infant in breast milk,^{64,65} aromatherapy is largely non-interventionist when compared to the side-effects and long-term complications of anxiolytic drugs. As the pregnant woman's body mass increases, it would be prudent of the therapist to reduce the area massaged (e.g. back or feet and hands), as the penetration of the essential oils will increase, due to their affinity with the cellular membranes.⁶⁶

Essential oils are partly absorbed through the process of olfaction by stimulating areas associated with smell in the limbic system of the brain and evidence that odours effect emotions and cognition have received empirical support,⁶⁷ although odour memories can sometimes heighten women's anxieties and fears.^{68,69} It may be significant that olfaction is the first of the senses to develop in utero: many connections from the olfactory nerve to the higher brain centres are made before the baby is delivered.⁷⁰⁻⁷² However, the hyperosmic state of many women during pregnancy, caused by vascular congestion as a result of increased circulating oestrogen, suggests judicious use of essential oils.⁷³ Studies have demonstrated the choice of area to treat a woman could increase her anxieties, especially in the hospital environment.^{74,75}

Certain essential oils, such as lavender and rosemary, have been investigated repeatedly for their effects on anxiety in general,^{51,74,76,77} and specific chemical constituents, for example, linalool and linalyl acetate, have been identified as having anxiolytic actions.⁷⁸⁻⁸⁰

There have been very few studies using essential oils specifically in pregnancy, therefore it is necessary to apply the findings from animal studies and from human studies in which aromatherapy has been used both for healthy volunteers and for ill patients. However, animal studies should be viewed with caution, as short-term, low-dose applications, given for therapeutic reasons to pregnant women, are noticeably different from tests performed on rats or mice given excessively large doses.^{81,82} In clinical studies, cancer patients found that their anxiety levels were reduced when given massage with citrus sinensis oil⁷⁴ and it is interesting to note a gender-related issue, with citrus oils appearing to have a greater effect on women than on men.⁸³⁻⁸⁵

There are few empirical studies that demonstrate the efficacy and safety of essential oils with humans, but there are studies relating to anxiety in labour, palliative care, dementia, but not pregnancy.^{74,75,86-95,108} The complexity of anxiety as a variable is not suitable to be analysed statistically, as fall in blood pressure does not necessarily imply fall in anxiety levels.⁹⁶ Researchers have suggested that patients who received aromatherapy reported significantly greater improvement in their level of anxiety for short periods, and a better quality of life.⁷⁷ The largest maternity-specific study was conducted by Burns et al.⁹⁷ who offered aromatherapy to 8058 women in labour using ten essential oils, with no detrimental effects on mother and baby. The effect of aromatherapy reduced fear and anxiety by 62%, as well as relieving physical symptoms such as pain and nausea; lavender and frankincense oils being the most frequently used.

The use of aromatherapy can be perceived as a non-interventional therapy in contrast to anxiolytic drugs and there is insufficient evidence to support their safety in pregnancy, due to the possibility of teratogenic and mutagenic effects on the fetus.^{64,65} On investigation benzodiazepines and cannabis have been found in breast milk, and been associated with sedation and hypothermia in babies.^{32,98-100} Likewise, Menella et al.¹⁰¹ established when pregnant women ingested garlic it altered the odour of the amniotic fluid, which suggests that essential oils cross the placenta. A working knowledge of essential oil toxicology is required and crossing the placenta does not necessarily mean that there is a risk of toxicity to the fetus, as each component of the essential oil depends on the toxicity and the plasma concentrations of the compound.^{102,103} Mothers see aromatherapy, which does not have the stigma associated with psychiatric drugs, as a treatment

with nice smelly oils and having another option of aromatherapy, instead of prescription drugs could empower women.¹⁰⁴

Anxiety can magnify minor discomforts of early pregnancy, but many of the essential oils are contraindicated during early fetal formation due to their abortifacient and emmenagogic effects and the challenge for midwives is to apply the same principles of prescribing essential oils, as they do to conventional pharmaceuticals.¹⁰² For example, high doses of citral administered orally to rats resulted in intrauterine growth retardation and bone abnormalities.¹⁰⁵ In view of the higher than normal circulating levels of melanocytic hormone, women's skin sensitivities are increased in pregnancy and the phototoxic effects of citrus oils may be potentiated, therefore patch testing would be advised.^{106,107}

Conclusion

Although pregnancy is a normal physiological life event, high levels of anxiety increase physiopathological complications and adversely affect maternal and fetal outcome.

Midwives could be pivotal in identifying women who are anxious and reduce anxiety by introducing aromatherapy to a maternity unit. The evidence indicates that the pharmacological activity of essential oils facilitates optimal health in anxious pregnant women thus benefiting the fetus. Researchers have intimated that essential oil components are able to cross the placenta and reach the fetus, but this does not mean there is a risk of toxicity, as studies have demonstrated the safe use of essential oils has no detrimental effects on mother and baby.

The greatest challenge that lies before us involves adapting the scientific method, so that it can accurately test and explore the complexity of human variables, and midwives need to draw on both the data from science and the wisdom from their human experience. The current concept of benefit is often limited to clinical benefits, but considerations to social and lifestyle benefits needs to occur and these medical decisions should include costs effectiveness to decrease adverse neonatal outcomes. Audit and evaluation of the aromatherapy is important to identify costs for future research and to evaluate the effectiveness and safety. Patients perceive it as reducing anxiety levels, but what is not known is for how long the reduction in the level of anxiety is maintained. We need to integrate aromatherapy more fully into

midwifery practice, as it would be a great disservice for the future of mother and baby by limiting and narrowing our view.

References

1. Homer C, Farrall T, Davis G, Brown M. Women's worry in the antenatal period. *Br J Midwifery* 2002;10(6).
2. Hedegaard M. Life style, work and stress and pregnancy outcome. *Curr Opin Obstet Gynecol* 1999;11(6):553-6.
3. Dragonas T, Christodoulou G. Prenatal care. *Clin Psychol Rev* 1998;18(2):127-42.
4. Hayes B, Muller R, Bradley B. Perinatal depression: a randomized controlled trial of an antenatal education intervention for primiparas. *Birth* 2001;28(1):28-9.
5. Davidson J. Aromatherapy & work-related stress. *Int J Aromatherapy* 2002;12(3):145-51.
6. Lappin J. Depressed mood during pregnancy and after childbirth: time points for assessing prenatal mood must be optimised. *BMJ* 2001;323(7325):1367.
7. O'Cathain A, Walters S, Nicholl P, Thomas J, Kirkham M. Use of evidence based leaflets to promote informed choice in maternity care: controlled trial in everyday practice. *BMJ* 2002;324(7338):643.
8. Johanson R, Newborn M, Macfarlane A. Has the medicalisation of childbirth gone too far? *BMJ* 2002;324:802-4.
9. Maloni J, Kane J, Suen L-J, Wang K. Dysphoria among high-risk pregnant hospitalised women on bed rest: a longitudinal study. *Nurs Res* 2002;51:92-9.
10. Marcin M, Nemeroff C. The neurobiology of social anxiety disorder: the relevance of fear and anxiety. *Acta Psych Scand* 2003;108(Suppl 417):51-64.
11. Kowalcek I, Huber G, Lammers C, Brunk J, Bieniakiewicz I, Gembruch U. Anxiety scores before and after prenatal testing for congenital anomalies. *Arch Gynecol Obstet* 2003;267(3):126-9.
12. Cote-Arsenault D. The influence of perinatal loss on anxiety in multigravidas. *J Obstet Gynaecol Neonatal Nurs* 2003;32(5):623-9.
13. Melender H-L. Experiences of fears associated with pregnancy and childbirth: a study of 329 pregnant women. *Birth* 2002;29(2):101-11.
14. Stapleton H, Kirkham M, Thomas G, Curtis P. Language use in antenatal consultations. *Br J Midwifery* 2002;10(5):273-6.
15. Stapleton H, Kirkham M, Curtis P, Thomas G. Silence and time in antenatal care. *Br J Midwifery* 2002;10(6):393-6.
16. Achat H, Kawachi I, Levine S, Berkey C, Coakley E, Colditz G. Social networks, stress and health related quality of life. *Qual Life Res* 1998;7:735-50.
17. Jomeen J, Martin C. Is the hospital anxiety and depression scale (HAPS) a reliable screening tool in early pregnancy? *Psychol Health* 2004;19(6):787-800.
18. Bower P, Richards D, Lovell K. The clinical and cost effectiveness of self-help treatments for anxiety and depressive disorders in primary care: a systematic review. *Br J General Practice* 2001;51(471):838-45.
19. Bennett V, Brown L. *Myles textbook for midwives*, 13th ed. London: Churchill Livingstone; 1999.
20. Doyle-Waters M, Kishor N, Doyle-Baker P. Development and validation of an anxiety scale for pregnancy. *Med Sci Sports Exercise* 2001;33(5 Suppl 1):S168.
21. Ohman S, Grunewald C, Waldenstrom U. Womens' worries during pregnancy testing the Cambridge Worry Scale on 200

- Swedish women. *Scand J Caring Sciences* 2003;17(2): 148–52.
22. Spielberger C. *State-trait anxiety inventory: a comprehensive bibliography*, 2nd ed. Palo Alto London: Consulting Psychologists Press; 1989.
 23. Matthey S. Detection and treatment of postnatal depression (perinatal depression or anxiety). *Curr Opin Psychiatry* 2004;17(1):21–9.
 24. Casper R, Fleischer B, Lee-Ancas J, Gilles A, Gaylor E, DeBattista A, et al. Follow-up of children of depressed mothers exposed or not exposed to anti-depressant drugs during pregnancy. *J Pediatr* 2003;142(4):402–8.
 25. Oates M, Lee A. Depressed mood during pregnancy and after childbirth: data do not support idea that depression is more common antenatally than postnatally. *BMJ* 2001; 323(325):1365–8.
 26. Bascom A. Complementary and alternative therapies in occupational health. Part 11—Specific therapies. *AAOHN J* 2002;50(10):468–74.
 27. Durham R, Murphy T, Allan T, Richard K, Treiving L, Fenton G. Cognitive therapy, analytic psychotherapy and anxiety management training for anxiety disorder. *Br J Psychiatry* 1994;165:315–23.
 28. Cannon R. High risk pregnancy in the workplace—influencing positive outcomes. *AAOHN J* 2000;48(9):435–46.
 29. Ng C-C-M, Lai F-M, Yeo G-S-H. Assessment of maternal anxiety levels before and after amniocentesis. *Singapore Med J* 2004;45(8):370–4.
 30. House A, Stark D. ABC of psychological medicine: anxiety in medical patients. *BMJ* 2002;325:207–9.
 31. Gregson O, Looker T. The biological basis of stress management. *Br J Guidance Counselling* 1994;22(1):13–22.
 32. Gale C. Anxiety disorder. *BMJ* 2000;321(7270):1204–7.
 33. Morgan C, Wang S, Rasmusson A, Hazlett G, Anderson G, Charney D. Relationship among plasma cortisol catecholamines, neuropeptide Y and human performance during exposure to uncontrollable stress. *Psychosom Med* 2001; 63(3):412–22.
 34. Field T, Diego M, Hernandez-Reif M, Schanberg K, Kuhn. Right frontal EEG and pregnancy/neonatal outcomes. *Psychiatry* 2002;65(1):35–47.
 35. Matthew S, Coplan J, Gorman J. Neurobiology mechanisms of social anxiety disorder. *Am J Psychiatry* 2001;158(10): 1558–67.
 36. Wadhwa P, Porto M, Garite T, Chicz-DeMet A, Sandman C. Maternal corticotropin-releasing hormone levels in the early third trimester predict length of gestation in human pregnancy. *Am J Obstet Gynecol* 1998;179(4): 1079–85.
 37. Ruiz R, Fullerton J, Dudley D. The inter-relationship of maternal stress, endocrine factors and inflammation on gestational length. *Obstet Gynaecol* 2003;58(6):415–28.
 38. Teixeira J, Glover V, Fisk N. Acute cerebral redistribution in response to invasive procedures in the human fetus. *Am J Obstet Gynecol* 1999;181(4):1018–25.
 39. Dayan J, Creveuil C, Herlicoviez M, Herbel C, Baranger E, Savoye C, et al. Role of anxiety and depression and the onset of spontaneous preterm labor. *Am J Epidemiol* 2002; 155(4):293–301.
 40. Mancuso R, Schetter C, Rini C, Roesch-Scott C, Hobel-Calvin J. Maternal prenatal anxiety and corticotropin-releasing hormone associated with timing of delivery. *Psychosomat Med* 2004;66(5):762–9.
 41. Hedegaard M, Henriksen T, Sabroe S, Jorgen N. Psychological distress in pregnancy and preterm delivery. *BMJ* 1993;307(6898):234–9.
 42. Gitau R, Fisk N, Glover V. Human fetal and maternal corticotropin releasing hormone response to acute stress. *BMJ* 2004;89(1):F29–32.
 43. Field T, Hernandez-Reif, Hart S, Theakston H, Schanberg S, Kuhn C. Pregnant women benefit from massage therapy. *J Psychosom Obstet Gynecol* 1999;20(1):31–8.
 44. Glover V. Maternal stress or anxiety in pregnancy and emotional development of the child. *Br J Psych* 1997; 171(8):105–6.
 45. O'Connor T, Heron J, Golding J, Beveridge M, Glover V. Maternal antenatal anxiety and children's behavioural/emotional problems at 4 years: report from the avon longitudinal study of parents and children. *Br J Psych* 2002;180:502–8.
 46. Glover V, O'Connor T. Effects of antenatal stress and anxiety: implications for development and psychiatry. *Br J Psych* 2002;180:389–91.
 47. Huizink A, De Medina R, Pascale M, Mulder E, Gerard H, Biutelaar J. Psychological measures of prenatal stress as predictors of infant temperament. *J Am Acad Child Adolesc Psych* 2002;41(9):1078–85.
 48. Suarez L, Cardarelli K, Hendricks K. Maternal stress, social support and risk of neural tube defects among Mexican Americans. *Epidemiology* 2003;14(5):612–6.
 49. Hobel C, Dunker-Schetter C, Roesch S, Castro L, Arora C. Maternal plasma corticotropin-releasing hormone associated with stress at 20 weeks' gestation in pregnancy ending in preterm delivery. *Am J Obstet Gynecol* 1999;180: 257–63.
 50. Kimber L. Massage for childbirth and pregnancy. *Practising Midwife* 2002;5(3):21–3.
 51. Diego M, Dieter J, Field T, Lecanuet J, Hernandez-Rief M, Beutler J, et al. Fetal activity following stimulation of the mothers' abdomen, feet and hands. *Develop Psychobiol* 2002;41(4):396–406.
 52. Holey E, Cook E. *Therapeutic massage*. London: WB Saunders Company Ltd.; 1998.
 53. Cassar M. Massage in pregnancy. *Practising Midwife* 2001; 4(3):10–4.
 54. Chaitow L, Walker DJ. *Clinical application of neuromuscular techniques: the upper body*, vol. 1. USA: Churchill Livingstone; 2002.
 55. Hernandez-Reif M, Field T, Krasnegor J, Theafston H, Hossain Z, Burman I. High blood pressure and associated symptoms were reduced by massage therapy. *J Bodywork Movement Therap* 2000;4(1):31–8.
 56. Marcoux S, Berube S, Brisson C, Mondor M. Job strain and pregnancy-induced hypertension. *Epidemiology* 1999;10: 376–82.
 57. Sheu S, Irvin B, Huey-Shyan L, Chun-Lin M. Effects of progressive muscle relaxation on blood pressure and psychological status for clients with essential hypertension in Taiwan. *Holistic Nurs Pract* 2003;17(1): 41–7.
 58. Chang MY, Wang SY, Chen CH. Effects of massage on pain and anxiety during labour: a randomized controlled trial in Taiwan. *J Adv Nurs* 2002;38(1):68–73.
 59. Field T, Hernandez-Rief M, Taylor S, Quintano O, Burman I. Labour pain is reduced by massage therapy. *J Psychosom Obstet Gynecol* 1997;18(1):28–91.
 60. Hollis M. *Massage for therapists*, 2nd ed. London: Blackwell Science; 2000.
 61. Lee B. Caring for the emotions in pregnancy, birth and beyond. *Midwives* 2003;6(10):438–41.
 62. Buckley J. Massage and aromatherapy massage: nursing art and science. *Int J Palliative Nurs* 2002;8(6):276–80.

63. Long L, Huntley A, Ernst E. Which complementary and alternative therapies benefit which conditions? A survey of the opinions of 223 professional organizations. *Complement Therap Med* 2001;9:178-85.
64. Duddridge E. Using complementary therapies during the childbearing process. *Br J Midwifery* 2002;10(11):691-704.
65. Araujo I, Souza C, De-Carvalho R, Kuriyama S, Rodrigues R, Volimer R, et al. Study of the embryofetotoxicity of alpha-terpinene in the rat. *Food Chem Toxicol* 1996;34(5):477-82.
66. Jager W, Buchbauer G, Jirovertz L. Evidence of the sedative effects of neroli oil, citronellal and phenylethyl acetate on mice. *J Essent Oil Res* 1992;4:387-90.
67. Alexander M. Aromatherapy & immunity: how the use of essential oils aid immunity potentiality part iv modulating immunity with aromatherapy: conditioning, suppression and stimulation of the immune system. *Int J Aromatherapy* 2002;12(1):49-57.
68. Jellinek J. Odours and mental states. *Int J Aromatherapy* 1998/1999;9(3):115-6.
69. Field T, Ironson G, Scafidi F, Nawrocki T, Goncalves A, Burman I, et al. Massage therapy reduces anxiety and enhances EEG pattern of alertness and math computations. *Int J Neurosci* 1996;86:197-205.
70. Balazs T. The fragrant brain. Process and meaning in olfactory communication. *Int J Aromatherapy* 1998/1999;9(2):57-60.
71. Schaal B, Marlier L, Soussignan R. Human fetuses learn odours from their pregnant mother's diet. *Chem Senses* 2000;25(6):729-37.
72. Kawakami K, Takai-Kawakami K, Okazaki Y, Kurihara H, Shimizu Y, Yanaihara T. The effect of odors on human newborn infants under stress. *Infant Behavior Develop* 1998;2(4):531-5.
73. Stables D. *Physiology in childbearing with anatomy and biosciences*. London: Bailliere and Tindall; 2000.
74. De Valois B, Clarke E. A retrospective assessment of 3 years of patient audit for an aromatherapy massage service for cancer patients. *Int J Aromatherapy* 2001;11(3):134-43.
75. Dunn C, Sleep J, Collett D. Sensing an improvement: an experimental study to evaluate the use of aromatherapy, massage and periods of rest in an intensive care unit. *J Adv Nurs* 1995;21(1):34-40.
76. Moss M, Cook J, Wesnes K, Duckett P. Aromas of rosemary and lavender essential oils differentially affect cognition and mood in healthy adults. *Int J Neurosci* 2003;113(1):15-38.
77. Giordano J, Boatwright D, Stapleton S, Huff L. Blending the boundaries: steps toward an integration of complementary and alternative medicine into mainstream practice. *J Alternat Complement Med* 2002;8(6):897-906.
78. Bradshaw R, Marchant J, Meredith M, Broom D. Effects of lavender straw on stress and travel sickness in pigs. *J Alternat Complement Med* 1998;4(3):271-5.
79. Buchbauer G, Jivovetz L, Jager W, Dietrich H, Plank C. Fragrance compounds and essential oils with sedative effects upon inhalation. *J Pharmacol Sci* 1993;82(6):660-4.
80. Elisabetsky E, Marschner J, Souza D. Effects of linalool on glutamatergic system in the rat cerebral cortex. *Neurochem Res* 1995;20:461-5.
81. Fujiwara R, Tehruhisu K, Yokoyama M. Psychoneuroimmunological aromatherapy. *Int J Aromatherapy* 2002;12(2):77-82.
82. Guba R. Toxicity myths—essential oils and their carcinogenic potential. *Int J Aromatherapy* 2001;11(2):76-83.
83. Ceccarelli L, Masi F, Fiorenzani P, Aloisi A. Sex differences in the citrus lemon essential oil-induced increase of hippocampal acetylcholine release in rats exposed to a persistent painful stimulation. *Neurosci Lett* 2002;330:25-8.
84. Lehner A. Ambient odour of orange in a dental office reduces anxiety and improves mood in female patients. *Physiol Behav* 2000;71:83-6.
85. Marchand S, Arsenault P. Odors modulate pain perception. A gender-specific effect. *Physiol Behav* 2002;76:251-6.
86. Wilkerson S, Aldridge J, Salmon I, Cain E, Wilson B. An evaluation of aromatherapy massage in palliative care. *Palliative Med* 1999;13(5):409-17.
87. Hadfield N. The role of aromatherapy massage in reducing anxiety in patients with malignant brain tumours. *Int J Palliative Nurs* 2001;7(6):279-85.
88. Wiebe E. A randomized trial of aromatherapy to reduce anxiety before abortion. *Effectiv Clin Pract* 2000;3(4):166-9.
89. Ballard C, O'Brien J, Reichelt K, Perry E. Aromatherapy as a safe and effective treatment for the management of agitation in severe dementia: the results of a double-blind, placebo-controlled trial with Melissa. *J Clin Psych* 2002;63(7):553-8.
90. Holmes C, Hopkins V, Hensford C, Maclaughlin V, Wilkinson D, Rosenvinge H. Lavender oil as a treatment for agitated behaviour in severe dementia: a placebo controlled study. *Int J Geriatric Psych* 2002;17(4):305-8.
91. Kilstoff K, Chenoweth L. New approaches to health and well-being for dementia day-care clients, family carers and day care staff. *Int J Nurs Pract* 1998;4(2):70-83.
92. Vance D. Considering olfactory stimulation for adults with age-related dementia. *Perceptual Motor Skills* 1999;88:398-400.
93. Akhondzadeh S, Noroozian M, Mohammadi M, Ohadinia S. Melissa officinalis extract in the treatment of patients with mild to moderate alzheimer's disease: A double blind, randomised, placebo controlled trial. *J Neurol, Neurosurg Psych* 2003;74(7):863-9.
94. Beshara M, Giddings D. Use of plant essential oils in treating agitation in a dementia unit: 10 case studies. *Int J Aromatherapy* 2002;12(4):207-12.
95. Burns A, Byrne J, Ballard C, Holmes C. Sensory stimulation in dementia. *BMJ (International Edition)* 2002;325(7376):1312-4.
96. Mann S. The mind/body link in essential hypertension: time for a new paradigm. *Alternat Therapies* 2000;6(2):39-44.
97. Burns E, Blamey C, Ersser S, Lloyd A, Barnetson L. *The use of aromatherapy in intrapartum midwifery practice. The Women's Centre. Oxford Centre for Health Care Research and Development*. England: Oxford Brookes University; 1999.
98. Dolovitch L, Addis A, Regis Vaillancourt J. Benzodiazepine use in pregnancy and major malformations of oral cleft: meta-analysis of cohort and case-control studies. *BMJ* 1998;317:839-43.
99. Bernstein J. *Handbook of drug therapy in psychiatry*, 3rd ed. USA: Mosby Year Book; 1995.
100. Grotenhermen F. Clinical pharmacokinetics of cannabis. *J Cannabis Therap* 2003;3(1):3-51.
101. Menella J, Johnson A, Beauchamp G. Garlic ingestion by pregnant women alters the odor of amniotic fluid. *Chem Senses* 1995;20(2):207-9.
102. Bowles J. *The basic chemistry of aromatherapeutic essential oils*, 2nd ed. Australia: Good Scents Aromapleasures; 2002.

103. Tisserand R. Essential oil: safety 11. *Int J Aromatherapy* 1996;7(4):26-9.
104. Buckle R. Aromatherapy in the USA. *Int J Aromatherapy* 2003;13(1):42-6.
105. Nogueira A, Carvalho R, Souza C, Chahoud I, Paumgarten F. Study of the embryofeto-toxicity of citral in the rat. *Toxicology* 1995;96/2:105-13.
106. Anderson K, Johansen J, Bruze M, Frosch P, Goossens L. The time-dose-response relationship for elicitation of contact dermatitis in isoeugenol allergic individuals. *Int J Aromatherapy* 2001;11(4):206-12.
107. De Groot A, Frosch P. Adverse reactions to fragrance—a clinical review. *Contact Dermatitis* 1997;36:57-86.
108. Louis M, Kowalski S. Use of aromatherapy with hospice patients to decrease pain, anxiety and depression and to promote an increased sense of well being. *American Journal of Hospice Palliative Care* 2002;19(6):381-6.

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