List of references supporting the assessment of *Melaleuca alternifolia* (Maiden and Betch) Cheel, *M. linariifolia* Smith, *M. dissitiflora* F. Mueller and/or other species of *Melaleuca*, aetheroleum

**Final**

The European Medicines Agency acknowledges that copies of the underlying works used to produce this monograph were provided for research only with exclusion of any commercial purpose.

Altman P. Assessment of the skin sensitivity and irritancy potential of tea tree oil; A report prepared for the Australian Rural Industries Research and Development Corporation (RIRDC) 15 October 1991. [Final results of this study were published by Aspres and Freeman 2003], 1991


Aspres N, Freeman S. Predictive testing for irritancy and allergenicity of tea tree oil in normal human subjects. *Exogenous Dermatology* 2003, 2:258-261


AUSTTEAM (Australian Tea Tree Export & marketing) Ltd. Tea Tree Oil conference, 1995


Belaiche P. Treatment of Vaginal Infections of Candida albicans with the Essential Oil of *Melaleuca alternifolia*. *Phytotherapie* 1985a, 15:13-14

Belaiche P. Treatment of Skin infections, with the Essential Oil of *Melaleuca alternifolia*. *Phytotherapie* 1985b, 15:15-17


Bhushan M, Beck MH. Allergic contact dermatitis from tea tree oil in a wart paint. *Contact dermatitis* 1997, 36(2):117-118


Cal K. Commentary to the article "Human skin penetration of the major components of Australian tea tree oil applied in its pure form and as a 20% solution in vitro". *Eur J Pharm Biopharm* 2008, 68(3):838-839


List of references supporting the assessment of *Melaleuca alternifolia* (Maiden and Betch) Cheel, *M. linariifolia* Smith, *M. dissitiflora* F. Mueller and/or other species of *Melaleuca*, aetheroleum

EMA/HMPC/320929/2012  Page 2/11


Carson CF. Research in Progress - TTO- Demonstrating proof of concept/efficacy for innovative applications of tea tree oil: A report prepared for the Australian Rural Industries Research Development Corporation (RIRDC) Project: Microbial Adaptation and Tolerance to Tea Tree Oil PRJ-002403, 2009. Report No.: PRJ-002403


Carson CF, Hammer KA, Riley TV. *Melaleuca alternifolia* (Tea Tree) oil: a review of antimicrobial and other medicinal properties. Clin Microbiol Rev 2006, 19(1):50-62. Available at: [http://cmr.asm.org/content/19/1/50.full#ref-121](http://cmr.asm.org/content/19/1/50.full#ref-121)


Carson CF, Riley TV. Safety, efficacy and provenance of tea tree (*Melaleuca alternifolia*) oil. *Contact Dermatitis* 2001, 45(2):65-67

Carson CF, Riley TV and Cookson BD. Efficacy and safety of tea tree oil as a topical antimicrobial agent *Journal of Hospital Infection* 1998, 40:1 75-1 78


List of references supporting the assessment of *Melaleuca alternifolia* (Maiden and Betch) Cheel, *M. linariifolia* Smith, *M. dissitiflora* F. Mueller and/or other species of *Melaleuca*, aetheroleum

CMEC 13. Extracted ratified minutes from the 13. Meeting of the Complementary Medicines Evaluation Committee (CMEC, Australia) on 11 June 1999


Cox SD, Mann CM, Markham JIL, Bell HC, Gustafson JE, Warmington JR et al. The mode of antimicrobial action of the essential oil of *Melaleuca alternifolia* (tea tree oil). *Journal of Applied Microbiology* 2000, 88, 170–175

Coutts I, Shaw S, Orton D. Patch testing with pure tea tree oil 12 months experience. *Journal of Dermatology* 2002, 147(Suppl 62):70


Drury S. L’huile de Melaleuca, Un merveilleux remède naturel. Éditions Médecis_Entrelacs Orasy, France, 1995


EFSA Journal - Scientific Opinion on the safety and efficacy of aliphatic and alicyclic ethers (chemical group 16) when used as flavourings for all animal species 2012, 10(11):2967


Entwistle P. Australian Tea Tree Industry Code of Practice: Australian Tea Tree Industry Association (ATTIA), 2005 (unpublished)


European Pharmacopoeia 7th ed. Tea tree oil – Melaleuca aetheroleum 01/2008:1837


Feinblatt HM. Cajeput-type oil for the treatment of furunculosis. *Journal of the National Medical Association* 1960, 52:32-34


Faoagali J, George N and Leditschke JF. Does tea tree oil have a place in the topical treatment of burns? *Burns* 1997, 23(4):349-351


Greig JE, Carson CF, Stuckey MS, Riley TV. Safety of Tea Tree Oil - second stage A report prepared for the Australian Rural Industries Research Development Corporation (RIRDC) RIRDC Publication No 02/036, RIRDC Project No UWA-51A, 2002


Hammer KA, Carson CF, Riley TV. In vitro activities of ketoconazole, econazole, miconazole, and Melaleuca alternifolia (tea tree) oil against Malassezia species. Antimicrob Agents Chemother 2000, 44(2):467-469

Hammer KA, Carson CF, Riley TV. Antifungal activity of the components of Melaleuca alternifolia (tea tree) oil. J Appl Microbiol 2003b, 95(4):853-860


Hammer KA, Riley TV. Serial passage of Staphylococcus aureus and S. epidermidis in tea tree oil does not induce resistance to tea tree oil. Australian Society for Microbiology (ASM) Annual Scientific Meeting & Exhibition. Perth 2009

Harkenthal M, Hausen BM, Reichling J. 1,2,4-Trihydroxy menthane, a contact allergen from oxidized tea tree oil. Pharamazie 2000, 55(2):153-154

Hausen BM, Reichling J, Harkenthal M. Degradation products of monoterpenes are the sensitizing agents in tea tree oil. American Journal of Contact Dermatitis 1999, 10(2):68-77


Hostynck JJ, Maibach HI. Thresholds of elicitation depend on induction conditions. Could low level exposure induce sub-clinical allergic states that are only elicited under the severe conditions of clinical diagnosis? Food and Chemical Toxicology 2004, 42(11):1859-1865


ICP Firefly Pty Ltd. In vivo micronucleus test of Australian Tea Tree Oil (Melaleuca alternifolia) Batch ATTIA/0501: ICP Firefly Pty Ltd, PO Box 6198, Alexandria, NSW, Australia, 2005


Lawless J. Das Tea Tree Öl. The new guide to one of nature’s most remarkable gifts. Wilhelm Goldmann Verlag, Muenchen 1996


MB Research Laboratories. Local lymph node assay in mice (LLNA) - Tea Tree Oil (Melaleuca alternifolia). MB Research Laboratories. 1765 Wentz Road, Spinnerstown PA USA, 2007 (unpublished)

McMahon MA, Blair IS, Moore JE, McDowell DA. Habituation to sub-lethal concentrations of tea tree oil (Melaleuca alternifolia) is associated with reduced susceptibility to antibiotics in human pathogens. J Antimicrob Chemother 2007, 59(1):125-127


Messager S, Hammer KA, Carson CF, Riley TV. Effectiveness of hand-cleansing formulations containing tea tree oil assessed ex vivo on human skin and in vivo with volunteers using European standard EN 1499. The Journal of hospital infection 2005, 59(3):220-228


Mondello F, De Bernardis F, Girolamo A, Cassone A, Salvatore G. In vivo activity of terpinen-4-ol, the main bioactive component of Melaleuca alternifolia Cheel (tea tree) oil against azole-susceptible and -resistant human pathogenic Candida species. BMC Infect Dis 2006 3, 6:158


Nielsen JB. Literature review on tea tree oil: Toxicity profiles for tea tree oil, constituents of tea tree oil and known oxidation products: A report prepared for the Australian Rural Industries Research Development Corporation (RIRDC) 2005, Report No.: PRJ-002403

Nielsen JB. What you see may not always be what you get – Bioavailability and extrapolation from in vitro tests. Toxicology in Vitro 2008


RCC Ltd. Study A69041 Tea tree oil: Local lymph node assay (LLNA) in mice (identification of contact allergens): RCC Ltd, Zelgliweg 1, CH-4452 Itengen, Switzerland, 2006

RCC Ltd. Study A78682 Tea tree oil: Local lymph node assay (LLNA) in mice (identification of contact allergens): RCC Ltd, Zelgliweg 1, CH-4452 Itengen, Switzerland, 2006

RCC Ltd. Study A78816 Tea tree oil: Local lymph node assay (LLNA) in mice (identification of contact allergens): RCC Ltd, Zelgliweg 1, CH-4452 Itengen, Switzerland, 2006


Rutherford T, Nixon R, Tam M, Tate B. Allergy to tea tree oil: retrospective review of 41 cases with positive patch tests over 4.5 years. The Australasian journal of dermatology 2007, 48(2):83-87

Satchell AC, Saurajen A, Bell C, Barnetson RS. Treatment of interdigital tinea pedis with 25% and 50% tea tree oil solution: a randomized, placebo-controlled, blinded study. *The Australasian journal of dermatology* 2002a, 43(3):175-178


Scientific Committee on Consumer Products (SCCP). Opinion on Tea Tree Oil - SCCP/08438/04: European Commission, 7 December 2004

Scientific Committee on Consumer Products (SCCP). Opinion on Tea Tree Oil - SCCP/1155/08: European Commission, 16 December 2008


Selvaag E, Eriksen B, Thune P. Contact allergy due to tea tree oil and cross-sensitization to colophony. *Contact dermatitis*. 1994, 31(2):124-125


Southwell IA, Russell MF, Davies NW. Detecting traces of methyl eugenol in essential oils: tea tree oil, a case study. *Flavour Fragr J* 2011, 26: 336-340

Stevenson OE, Finch TM. Allergic contact dermatitis from rectified camphor oil in Earex ear drops. *Contact dermatitis* 2003, 49(1):51


Thomsen NA, Carson CF, Hammer KA, Riley TV. Antimicrobial susceptibility of Staphylococci after exposure to Melaleuca alternifolia (tea tree) oil. Australian Society for Microbiology (ASM) Annual Scientific Meeting & Exhibition. Perth 2009


Treudler R, Richter G, Geier J, Schnuch A, Orfanos CE, Tebbe B. Increase in sensitization to oil of turpentine: recent data from a multicenter study on 45,005 patients from the German-Austrian Information Network of Departments of Dermatology (IVDK). *Contact dermatitis* 2000, 42(2):68-73


Walker M. Clinical investigation of Australian Melaleuca alternifolia oil for a variety of common foot problems. *Current Podiatry* 1972:7–15
