BMJ 2012;345:e5909 doi: 10.1136/bmj.e5909 (Published 2 October 2012)

RESEARCH

Indoor tanning and non-melanoma skin cancer: systematic review and meta-analysis

Mackenzie R Wehner *medical student*¹ *MPhil scholar in epidemiology*², Melissa L Shive *medical student*³, Mary-Margaret Chren *professor*⁴, Jiali Han *associate professor*⁵⁶, Abrar A Qureshi *associate professor*⁵, Eleni Linos *assistant professor*⁴

¹Stanford University School of Medicine, Stanford, CA, USA; ²Department of Public Health and Primary Care, University of Cambridge, Cambridge, UK; ³University of California San Francisco (UCSF) School of Medicine, San Francisco, CA, USA; ⁴Department of Dermatology, University of California San Francisco (UCSF), 2340 Sutter Street, San Francisco, CA, 94143-0808, USA; ⁵Department of Dermatology, Channing Laboratory, Department of Medicine, Brigham and Women's Hospital, Harvard Medical School, Boston, MA, USA; ⁶Department of Epidemiology, Harvard School of Public Health, Boston, MA, USA

Abstract

Objective To synthesise the literature on indoor tanning and non-melanoma skin cancer.

Design Systematic review and meta-analysis.

Data sources PubMed (1966 to present), Embase (1974 to present), and Web of Science (1898 to present).

Study selection All articles that reported an original effect statistic for indoor tanning and non-melanoma skin cancer were included. Articles that presented no data, such as review articles and editorials, were excluded, as were articles in languages other than English.

Data extraction Two investigators independently extracted data. Random effects meta-analysis was used to summarise the relative risk of ever use versus never use of indoor tanning. Dose-response effects and exposure to indoor tanning during early life were also examined. The population attributable risk fraction for the United States population was calculated.

Results 12 studies with 9328 cases of non-melanoma skin cancer were included. Among people who reported ever using indoor tanning compared with those who never used indoor tanning, the summary relative risk for squamous cell carcinoma was 1.67 (95% confidence interval 1.29 to 2.17) and that for basal cell carcinoma was 1.29 (1.08 to 1.53). No significant heterogeneity existed between studies. The population attributable risk fraction for the United States was estimated to be 8.2% for squamous cell carcinoma and 3.7% for basal cell carcinoma. This corresponds to more than 170 000 cases of non-melanoma skin cancer each year attributable to indoor tanning. On the basis of data from three studies, use of indoor tanning before age 25 was more strongly associated with both squamous cell carcinoma

(relative risk 2.02, 0.70 to 5.86) and basal cell carcinoma (1.40, 1.29 to 1.52).

Conclusions Indoor tanning is associated with a significantly increased risk of both basal and squamous cell skin cancer. The risk is higher with use in early life (<25 years). This modifiable risk factor may account for hundreds of thousands of cases of non-melanoma skin cancer each year in the United States alone and many more worldwide. These findings contribute to the growing body of evidence on the harms of indoor tanning and support public health campaigns and regulation to reduce exposure to this carcinogen.

Introduction

The incidence of basal cell carcinoma and squamous cell carcinoma of the skin, collectively termed non-melanoma skin cancer (NMSC), has increased dramatically over previous decades, in what some have termed an epidemic.^{1 2} NMSC is by far the most common human malignancy, and nearly 30% of white people living in areas of exposure to high ultraviolet radiation will develop an NMSC in their lifetime.³ Because of its high prevalence, NMSC is a considerable financial burden to healthcare systems.⁴⁻⁶

Although NMSC is the most common cancer, it is often excluded from national cancer registries and cancer databases because it typically does not affect survival. NMSC is more common in older men, but more tumours have recently been documented in women and in younger people of both sexes.^{7 8} Besides older age and male sex, major risk factors for development of NMSC include light skin, family history, residence at latitudes near the equator, and exposure to ultraviolet radiation.^{9 10}

Correspondence to: E Linos linose@derm.ucsf.edu

Extra material supplied by the author (see http://www.bmj.com/content/345/bmj.e5909?tab=related#webextra)

Because it is potentially modifiable, indoor tanning is a particularly important type of exposure to ultraviolet radiation. Indoor tanning is a class I carcinogen, considered "carcinogenic to humans" by the International Agency for Research on Cancer.¹¹ Indoor tanning is significantly associated with increased risk of malignant melanoma and was shown to be a potential risk factor for NMSC in a previous, smaller meta-analysis.¹² ¹³ Several studies have examined the link between NMSC and indoor tanning.¹⁴⁻²⁹ However, the study sizes and the percentage of participants reporting exposure have been small, yielding varied results. Our aim in this study was to synthesise the available data on indoor tanning and NMSC.

Methods

We carried out this review in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines.³⁰

Literature search

We defined NMSC as either basal cell carcinoma or squamous cell carcinoma and indoor tanning as the use of an ultraviolet emission device to produce a cosmetic tan. The terminology used in the literature is diverse. In this analysis, we considered indoor tanning, sunbed, sunlamp, tanning bed, tanning booth, solarium, artificial tanning, artificial ultraviolet tanning, non-solar ultraviolet tanning, and variations of these to be synonymous with indoor tanning.

We identified studies through searches of electronic databases and by scanning reference lists of articles. We searched PubMed (1966 to present), Embase (1974 to present), and Web of Science (1898 to present). Two authors (MRW and MLS) did the search, and the last search was run on 16 March 2012. Additionally, we reviewed identified articles and reviews on the topics of NMSC and ultraviolet exposure closely to locate additional articles. Specifics of the search strategy used in each database are detailed in the supplementary materials.

All published articles in English that reported an effect statistic for indoor tanning and NMSC or that reported measuring or adjusting for indoor tanning in a study including participants with NMSC were eligible for inclusion. Two authors (MRW and MLS) assessed the eligibility of studies by using the title and abstract for initial screening followed by review of the full text. We excluded articles that presented no data, such as review articles and editorials, and articles in languages other than English. Inclusion criteria for quantitative meta-analysis were studies reporting an effect estimate, such as an odds ratio or hazard ratio, or reporting measurement of or adjustment for indoor tanning that could be used to calculate an effect estimate.

We used a data extraction sheet, which was developed on the basis of the Cochrane Consumers and Communication Review Group's data extraction template. We extracted the following data items from each study: characteristics of study participants (including age, sex, type of NMSC) and inclusion/exclusion criteria, characteristics of study design (including design type, presence of matching in case-control studies, matching characteristics, and number of controls per case), outcomes (including effect estimates of different doses of exposure to indoor tanning), and statistical methods (including univariate or multivariate analyses, logistic regressions, and variables included).

Statistical methods

For the primary meta-analysis, we used the odds ratio or hazard ratio for ever exposure to indoor tanning, which was the exposure measure used by most of the studies. Ever exposure (for example, participants were asked: "Have you ever used an indoor ultraviolet tanning device to produce a cosmetic tan?") was available for 10 studies. For two studies, the only available measure was "regular" exposure (Bakos et al¹⁷) or greater than five exposures per year (Walther et al²⁸). We thus did the primary analysis on the 10 studies reporting ever exposure and a subsequent sensitivity analysis that included all 12 studies. In one instance, we calculated an unadjusted odds ratio and 95% confidence interval from the raw data of a matched study.¹⁴ Data comparing ever use versus never use of indoor tanning came directly from the authors of one study that did not show this comparison in the manuscript.²⁹ We also did a sensitivity analysis excluding four retrospective studies that did not fully adjust for confounders (see supplementary table).14 19 21 2

We did additional analyses on studies that reported effect statistics for high dose exposures and on studies that reported effect statistics for young age at exposure. High dose exposure was assessed by four studies, which used exposure measures of "regular use of tanning beds in lifetime,"¹⁷ 6-26 years of regular exposure,²⁰ exposure more than five times per year,²⁸ and exposure four times per year.²⁹ Three studies reported young age at first exposure, using 16 years of age or younger,²⁰ less than 20 years of age,²⁵ and "high school/college," which is typically 14-25 years of age.²⁹ Although these studies used different doses of exposure for high dose exposure and different cut-off ages for young age, these represent the only and thus best available data.

We used Stata 11 statistical software to do random effects model meta-analyses, yielding summary relative risks and 95% confidence intervals. All statistical tests were two sided. We analysed data for basal cell carcinoma and squamous cell carcinoma separately. To investigate variability (heterogeneity) in study outcomes, we used a χ^2 test for heterogeneity and an I² statistic.

We used the STROBE statement guidelines to assess the quality of individual studies.³¹ Of 16 studies identified, two did not state when data was collected,^{18 26} three case-control studies did not match cases and controls,^{19 21 28} and three case-control studies used frequency rather than individual matching.^{20 25 27} Additionally, nine studies did not clearly report participation rates,^{15 17 18 19 21 23 26 28 29} and three did not state ages or sexes of participants.^{22 23 26}

To assess potential small study effects and publication bias across studies, we created funnel plots by plotting the effect found by each study against the inverse of its standard error. We reviewed the funnel plot visually and used Begg's rank correlation test and Egger's weighted linear regression test for formal testing. This aimed to investigate the possibilities that small studies showing no effects may not be published and that small studies are more likely to be done with less methodological rigor, leading to inaccurate effect estimates.

We calculated population proportional attributable risk as (prevalence of exposure×(RR-1))/(1+prevalence of exposure×(RR-1)), where RR is relative risk. We calculated this for the United States, for which we had representative data on both the incidence of NMSC (estimated at 3 507 693 new cases in 2006^2) and one year prevalence of indoor tanning (estimated at 13.4% for 2005^{32}).

Results

Figure 1 || shows the study selection process. Database searches yielded a total of 558 unique publications whose title and abstract were screened. We discarded 536 because of no relevance (n=416), no unique data presented (for example, editorial article or review, n=110), or language other than English (n=10). We examined the remaining 22 articles in full text. Sixteen studies met the inclusion criteria and are included in table 1 \Downarrow . We included 12 studies in the quantitative meta-analysis. Excluded from meta-analysis were one study that reported only mean numbers of exposures (numbers of indoor tanning visits),¹⁸ one study that grouped patients with basal cell carcinoma, squamous cell carcinoma, and melanoma,²⁴ and two studies that grouped basal cell carcinoma and squamous cell carcinoma together.^{23 26} These last two studies also used patients admitted to hospital with solid organ cancer as controls, which are unlikely to be representative of the population from which the cases arose.^{23 26} All four studies showed non-significant associations, although one showed a statistically significant positive association when sunbed exposure was grouped with medical phototherapy.24

The 12 studies included in the meta-analysis were published between 1985 and 2012, used data collected between 1977 and 2010 in six different countries, and included 80 661 total participants and 9328 cases of NMSC. All 12 studies reported or provided raw data for effect estimates as odds ratios or hazard ratios (table $2\downarrow$). When available, we preferentially present effect estimates reported using multivariate models.

Effect estimates for ever exposure to indoor tanning compared with never exposure were available for 10 out of 12 studies. A meta-analysis of these studies yielded summary relative risks of 1.29 (95% confidence interval 1.08 to 1.53) for basal cell carcinoma and 1.67 (1.29 to 2.17) for squamous cell carcinoma (fig 2)). A χ^2 test for heterogeneity was non-significant for both basal cell carcinoma (P=0.14) and squamous cell carcinoma (P=0.09); I² statistics were 36.8% (95% confidence interval 0% to 72%) for basal cell carcinoma and 47.1% (0% to 79%) for squamous cell carcinoma. To include all available studies, we did a sensitivity analysis using the effect statistics from all 12 studies, including the two studies that reported only higher dose exposure.^{17 28} Both studies considered only basal cell carcinoma; with these two studies included, the summary relative risk for basal cell carcinoma was 1.25 (1.01 to 1.55). Funnel plots assessing publication bias were symmetrical for both basal cell carcinoma and squamous cell carcinoma, and all Begg's and Egger's tests were not statistically significant, suggesting that publication bias was unlikely. Our results did not change appreciably in a sensitivity analysis excluding four retrospective studies that did not fully adjust for confounders.14 19 21 27

To assess the presence of a dose-response effect, we did a sub-analysis on studies that included effect estimates for frequent or multiple (high dose) exposures to indoor tanning (table $3\downarrow$, top). High dose exposure was associated with a relative risk of 1.50 (0.81 to 2.77) for basal cell carcinoma. To assess the potential effect of exposure to indoor tanning at a young age, we did a sub-analysis on studies that included effect estimates for early life exposure (table $3\downarrow$, bottom). Indoor tanning exposure before age 25 was associated with a relative risk of 1.40 (1.29 to 1.52) for basal cell carcinoma and 2.02 (0.70 to 5.86) for squamous cell carcinoma.

Population attributable risk

Applying our summary risk estimates to the prevalence of exposure to indoor tanning in the United States, we calculated the population attributable risk fraction at 3.7% for basal cell carcinoma and at 8.2% for squamous cell carcinoma. This corresponds to 98 408 cases of basal cell carcinoma and 72 244 cases of squamous cell carcinoma, making 170 652 cases of non-melanoma skin cancer each year attributable to indoor tanning (see supplementary figure).

Discussion

In this systematic review and meta-analysis of more than 9300 cases of non-melanoma skin cancer from 12 studies, we found a positive, statistically significant association between exposure to indoor tanning and NMSC. Ever exposure to indoor tanning was associated with a 67% higher risk for squamous cell carcinoma and a 29% higher risk for basal cell carcinoma. Exposure to indoor tanning at a young age was significantly associated with an increased risk for basal cell carcinoma and showed a non-significant increased risk for squamous cell carcinoma. High dose exposure to indoor tanning showed a non-significant increased risk for basal cell carcinoma. This suggests a critical period for exposure during early life and a potential dose-response effect.

Possible explanations for findings

A causal link between indoor tanning and NMSC is one possible explanation for our findings. This link is biologically plausible, because both ultraviolet A and ultraviolet B radiations are established carcinogens in animal models and human studies.^{11 33} Several population based studies have documented rapid rises in skin cancer among young women, coinciding with the adoption of indoor tanning and supporting this causal relation.^{8 34 35} The dose-response effect noted in several studies further supports this argument.^{20 29} The temporal relation in which indoor tanning at a young age is a stronger risk factor not only supports a causal interpretation but also implies a critical period of higher susceptibility during early life.

Alternative explanations for these findings include the possibility that these observational and mostly retrospective studies were confounded by skin type or outdoor exposure to ultraviolet radiation. For example, fair skinned people who are more susceptible to NMSC might use indoor tanning more often, and indoor tanners may also get more outdoor ultraviolet exposure.35 Because most of the studies included in the primary meta-analysis (8/10) controlled for skin type or sun sensitivity and many (5/10) controlled for outdoor ultraviolet exposure, we do not feel that this confounding alone could account for the significant associations observed. Also, differential recall of use of indoor tanning in people already diagnosed as having skin cancer is a concern in case-control studies. However, given that prospective cohort studies found similar effect sizes, and are heavily weighted in this meta-analysis, we do not think that recall bias significantly affects our conclusion. Although they are always a concern in literature based meta-analyses, publication bias and small study effects are unlikely to fully explain these findings on the basis of our analyses.

Comparison with other studies

These findings are consistent with a previous meta-analysis of indoor tanning and NMSC,¹³ which, using only five studies, found a significant increase in risk for squamous cell carcinoma (summary relative risk 2.25, 95% confidence interval 1.08 to 4.70) and a non-significant effect for basal cell carcinoma (summary relative risk 1.03, 0.56 to 1.90). Our analysis, using substantially more studies, found a similarly significantly increased risk for squamous cell carcinoma and, notably, found

a significantly increased risk for basal cell carcinoma. Our findings add to the growing body of evidence on the harms of indoor tanning. Indoor tanning is already considered a class I carcinogen on the basis of its effect on malignant melanoma.^{11 12} Although NMSC is generally not a lethal cancer, it affects a vast number of people worldwide and accounts for a considerable disease burden.^{1 2 6 36} Although the population attributable risk fraction of indoor tanning on NMSC is modest, when applied to the 3.5 million new cases of NMSC diagnosed each year, indoor tanning may account for hundreds of thousands of new cases each year in the United States alone. Yet this is a global problem: recent studies from Australia, France, Denmark, Germany and Sweden suggest that 10.6-35% of people have used a tanning bed at least once in their lives, and 1.3-29.9% have done so in the previous year.^{37 38 39 40 41 42 43}

Limitations of study

This study is limited by the fact that it included only observational and mostly case-control studies. However, a randomised trial of indoor tanning is not realistic, and most of the included studies controlled for multiple potential confounders, making this the best level of evidence possible. Another potential limitation of this meta-analysis is the broad time period spanned by the data. Included studies collected data from the 1970s to the 2010s, which is important because indoor tanning devices have changed over time from high ultraviolet B output to predominately ultraviolet A output.^{44 45 46} However, multiple studies have indicated that both ultraviolet B and ultraviolet A seem to be capable of causing significant mutagenic damage to skin.^{11 33 47 48}

Conclusions

Indoor tanning, which is already an established risk factor for malignant melanoma, is probably a risk factor for both squamous cell carcinoma and basal cell carcinoma, which are the most common human cancers. We hope that these findings can support public health campaigns and motivate increased regulation to reduce exposure to this carcinogen, especially during early life.

We thank Stephen Bent of the University of California San Francisco and Emanuele Di Angelantonio of the University of Cambridge for their assistance.

Contributors: MRW, M-MC, and EL were involved in the conception and design of the study. MRW, MLS, and EL did the data collection and analysis. MRW, MLS, M-MC, and EL interpreted the data. MRW and EL drafted the manuscript, and all authors critically revised it for important intellectual content. All authors approved the final version to be published. EL is the guarantor.

Funding: This project was supported by award number KL2RR024130 from the National Center for Research Resources of the National Institutes of Health, and by award number K24 AR052667 from the National Institute of Arthritis and Musculoskeletal and Skin Diseases of the National Institutes of Health. The study sponsors were not involved in the study design and the collection, analysis, and interpretation of data, nor the writing of the article or the decision to submit it for publication. The authors were independent from the study sponsors.

Competing interests: All authors have completed the Unified Competing Interest form at www.icmje.org/coi_disclosure.pdf (available on request from the corresponding author) and declare: no support from any organisation for the submitted work; MMC does consultancy for Genentech; no other relationships or activities that could appear to have influenced the submitted work.

Ethical approval: Not needed.

Data sharing: No additional data available.

- 1 Lomas A, Leonardi-Bee J, Bath-Hextall F. A systematic review of worldwide incidence of nonmelanoma skin cancer. Br J Dermatol 2012;166:1069-80.
- 2 Rogers HW, Weinstock MA, Harris AR, Hinckley MR, Feldman SR, Fleischer AB, et al. Incidence estimate of nonmelanoma skin cancer in the United States, 2006. Arch Dermatol 2010;146:283-7.
- 3 Miller DL, Weinstock MA. Nonmelanoma skin cancer in the United States: incidence. J Am Acad Dermatol 1994;30:774-8.
- 4 Chen JG, Fleischer AB Jr, Smith ED, Kancler C, Goldman ND, Williford PM, et al. Cost of nonmelanoma skin cancer treatment in the United States. *Dermatol Surg* 2001;27:1035-8.
- 5 Housman TS, Feldman SR, Williford PM, Fleischer AB Jr, Goldman ND, Acostamadiedo JM, et al. Skin cancer is among the most costly of all cancers to treat for the Medicare population. J Am Acad Dermatol 2003;48:425-9.
- 6 Mudigonda T, Pearce DJ, Yentzer BA, Williford P, Feldman SR. The economic impact of non-melanoma skin cancer: a review. J Natl Comp Canc Netw 2010;8:888-96.
- 7 Roewert-Huber J, Lange-Asschenfeldt B, Stockfleth E, Kerl H. Epidemiology and aetiology of basal cell carcinoma. Br J Dermatol 2007;157(suppl 2):47-51.
- 8 Christenson LJ. Incidence of basal cell and squamous cell carcinomas in a population younger than 40 years. JAMA 2005;294:681-90.
- 9 Armstrong BK, Kricker A. The epidemiology of UV induced skin cancer. J Photochem Photobiol B 2001;63:8-18.
- 10 Wei-Passanese EX, Han J, Lin W, Li T, Laden F, Qureshi AA. Geographical variation in residence and risk of multiple nonmelanoma skin cancers in US women and men. *Photochem Photobiol* 2012:88:483-9.
- 11 El Ghissassi F, Baan R, Straif K, Grosse Y, Secretan B, Bouvard V, et al. A review of human carcinogens—part D: radiation. *Lancet Oncol* 2009;10:751-2.
- 12 Gallagher RP, Spinelli JJ, Lee TK. Tanning beds, sunlamps, and risk of cutaneous malignant melanoma. *Cancer Epidemiol Biomarkers Prev* 2005;14:562-66.
- 13 International Agency for Research on Cancer Working Group on Artificial Ultraviolet (UV) Light And Skin Cancer. The association of use of sunbeds with cutaneous malignant regioneers and check this access a surface to access a cutation of the Cancer 2002/120/11/12 20
- melanoma and other skin cancers: a systematic review. Int J Cancer 2007;120:1116-22.
 Asgari MM, White E, Warton EM, Hararah MK, Friedman GD, Chren M-M. Association of tea consumption and cutaneous squamous cell carcinoma. Nutr Cancer 2011;63:314-8.
- 15 Aubry F, MacGibbon B. Risk factors of squamous cell carcinoma of the skin: a case-control study in the Montreal region. *Cancer* 1985;55:907-11.
- 16 Bajdik CD, Gallagher RP, Astrakianakis G, Hill GB, Fincham S, McLean DI. Non-solar ultraviolet radiation and the risk of basal and squamous cell skin cancer. *Br J Cancer* 1996;73:1612-4.
- 17 Bakos RM, Kriz M, Mühlstädt M, Kunte C, Ruzicka T, Berking C. Risk factors for early-onset basal cell carcinoma in a German institution. *Eur J Dermatol* 2011;21:705-9.
- 18 Boyd AS, Shyr Y, King LE Jr. Basal cell carcinoma in young women: an evaluation of the association of tanning bed use and smoking. J Am Acad Dermatol 2002;46:706-9.
- 19 Corona R, Dogliotti E, D'Errico M, Sera F, Iavarone I, Baliva G, et al. Risk factors for basal cell carcinoma in a Mediterranean population: role of recreational sun exposure early in life. Arch Dermatol 2001;137:1162-8.
- 20 Ferrucci LM, Cartmel B, Molinaro AM, Leffell DJ, Bale AE, Mayne ST. Indoor tanning and risk of early-onset basal cell carcinoma. J Am Acad Dermatol 2011; published online 8 Dec.
- 21 Gon A, Minelli L. Risk factors for basal cell carcinoma in a southern Brazilian population: a case-control study. *Int J Dermatol* 2011;50:1286-90.
- 22 Han J, Colditz GA, Hunter DJ. Risk factors for skin cancers: a nested case-control study within the Nurses' Health Study. Int J Epidemiol 2006;35:1514-21.
- 23 Herity B, O'Loughlin G, Moriarty MJ, Conroy R. Risk factors for non-melanoma skin cancer. Ir Med J 1989;82:151-2.
- 24 Hogan DJ, To T, Wilson ER, Miller AB, Robson D, Holfeld K, et al. A study of acne treatments as risk factors for skin cancer of the head and neck. *Br J Dermatol* 1991:125:343-8.
- 25 Karagas MR, Stannard VA, Mott LA, Slattery MJ, Spencer SK, Weinstock MA. Use of tanning devices and risk of basal cell and squamous cell skin cancers. J Natl Canc Inst 2002;94:224-6.
- 26 O'Loughlin C, Moriarty MJ, Herity B, Daly L. A re-appraisal of risk factors for skin carcinoma in Ireland: a case control study. *Ir J Med Sci* 1985;154:61-5.
- 27 Rosso S, Joris F, Zanetti R. Risk of basal and squamous cell carcinomas of the skin in Sion, Switzerland: a case-control study. *Tumori* 1999;85:435-42.
- 28 Walther U, Kron M, Sander S, Sebastian G, Sander R, Peter RU, et al. Risk and protective factors for sporadic basal cell carcinoma: results of a two-centre case-control study in southern Germany. Clinical actinic elastosis may be a protective factor. Br J Dermatol 2004;151:170-8.
- 29 Zhang M, Qureshi AA, Geller AC, Frazier L, Hunter DJ, Han J. Use of tanning beds and incidence of skin cancer. J Clin Oncol 2012;30:1588-93.
- 30 Moher D, Liberati A, Tetzlaff J, Altman DG. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *BMJ* 2009;339:b2535.
- 31 Von Elm E, Altman DG, Egger M, Pocock SJ, Gøtzsche PC, Vandenbroucke JP. Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) statement: guidelines for reporting observational studies. *BMJ* 2007;335:806-8.
- 32 Heckman CJ, Coups EJ, Manne SL. Prevalence and correlates of indoor tanning among US adults. J Am Acad Dermatol 2008;58:769-80.
- Council on Scientific Affairs. Harmful effects of ultraviolet radiation. JAMA 1989;262:380-4.
 Hausauer AK, Swetter SM, Cockburn MG, Clarke CA, Increases in melanoma among
- 34 Hausauer AN, Swetter SM, Cockourn MG, Clarke CA. Increases in melanoma among adolescent girls and young women in California: trends by socioeconomic status and UV radiation exposure. Arch Dermatol 2011;147:783-9.
- 35 Purdue MP, Freeman LEB, Anderson WF, Tucker MA. Recent trends in incidence of cutaneous melanoma among US Caucasian young adults. *J Invest Dermatol* 2008;128:2905-8.
- 36 Guy GP, Ekwueme DU. Years of potential life lost and indirect costs of melanoma and non-melanoma skin cancer: a systematic review of the literature. *Pharmacoeconomics* 2011;29:863-74.
- 37 Francis K, Dobbinson S, Wakefield M, Girgis A. Solarium use in Australia, recent trends and context. Aust N Z J Public Health 2010;34:427-30.
- 38 Lawler SP, Kvaskoff M, DiSipio T, Whiteman D, Eakin E, Aitken J, et al. Solaria use in Queensland, Australia. Aust N Z J Public Health 2006;30:479-82.

What is already known on this topic

Non-melanoma skin cancer (NMSC) is the most common human cancer

Indoor tanning is a known risk factor for malignant melanoma, but the data on NMSC are less clear because small individual studies have resulted in varied effect estimates

What this study adds

Indoor tanning is associated with NMSC, especially when exposure occurs early in life

This modifiable risk factor accounts for hundreds of thousands of new cancer cases in the United States each year

More countries, including the United States, should follow in the footsteps of Europe, Australia, and Canada and restrict tanning bed use for minors

- Ezzedine K, Malvy D, Mauger E, Nageotte O, Galan P, Hercberg S, et al. Artificial and 39 natural ultraviolet radiation exposure: beliefs and behaviour of 7200 French adults. J Eur Acad Dermatol Venereol 2008;22:186-94.
- 40 Køster B, Thorgaard C, Philip A, Clemmensen I. Sunbed use and campaign initiatives in the Danish population, 2007-2009: a cross-sectional study. J Eur Acad Dermatol Venereol 2011; published online 9 Jan.
- Börner FU, Schütz H, Wiedemann P. A population-based survey on tanning bed use in 41 Germany. BMC Dermatol 2009;9:6.
- Boldeman C. Bränström R. Dal H. Kristiansson S. Rodvall Y. Jansson B. et al. Tanning 42 habits and sunburn in a Swedish population age 13-50 years. Eur J Cancer 2001;37:2441-8.
- Bränström R, Ullén H, Brandberg Y. Attitudes, subjective norms and perception of 43 behavioural control as predictors of sun-related behaviour in Swedish adults. Prev Med 2004;39:992-9.
- Diffey BL, Farr PM. Tanning with UVB or UVA: an appraisal of risks. J Photochem Photobiol 44 B 1991;8:219.
- Gerber B. Mathys P. Moser M. Bressoud D. Braun-Fahrländer C. Ultraviolet emission 45 spectra of sunbeds. Photochem Photobiol 2002;76:664-8.

- Nilsen LTN, Hannevik M, Aalerud TN, Johnsen B, Friberg EG, Veierød MB. Trends in UV 46 irradiance of tanning devices in Norway: 1983-2005. Photochem Photobiol 2008;84:1100-8. Wang SQ, Setlow R, Berwick M, Polsky D, Marghoob AA, Kopf AW, et al. Ultraviolet A
- 47 and melanoma: a review. J Am Acad Dermatol 2001;44:837-46.
- 48 Hocker T, Tsao H. Ultraviolet radiation and melanoma: a systematic review and analysis of reported sequence variants. Hum Mutat 2007;28:578-88.

Accepted: 28 August 2012

Cite this as: BMJ 2012;345:e5909

This is an open-access article distributed under the terms of the Creative Commons Attribution Non-commercial License, which permits use, distribution, and reproduction in any medium, provided the original work is properly cited, the use is non commercial and is otherwise in compliance with the license. See: http://creativecommons.org/licenses/bync/2.0/ and http://creativecommons.org/licenses/by-nc/2.0/legalcode.

Tables

Table 1| Summary of studies

Reference	Country, year of publication	Data collection	Diagnosis*	Total No	No of cases	Age (years)	% male	Exposure†
Studies include	d in meta-analysis		-					
Asgari et al14	United States, 2011	2004-05	SCC	830	415	43-85	61.9	Ever/never
Aubry and McGibbon ¹⁵	Canada, 1985	1977-78	SCC	266	92	Unknown	65.8	Ever/never
Bajdik et al ¹⁶	Canada, 1996	1983-84	BCC; SCC	812	226 BCC; 180 SCC	25-79	100.0	Ever/never
Bakos et al ¹⁷	Germany, 2011	2004-08	BCC	50	25	19-40	40.0	"Regular" exposure
Corona et al19	Italy, 2001	1995-97	BCC	324	166	≥18	47.8	Ever/never
Ferrucci et al20	United States, 2011	2006-10	BCC	766	376	<40	30.8	Ever/never
Gon and Minelli ²¹	Brazil, 2011	2006-07	BCC	407	127	18-80	41.7	Ever/never
Han et al ²²	United States, 2006	1989-98	BCC; SCC	1362	283 BCC; 275 SCC	43-68	0.0	Ever/never
Karagas et al ²⁵	United States, 2002	1993-95	BCC; SCC	1436	603 BCC 293 SCC	25-74	59.5	Ever/never
Rosso et al ²⁷	Switzerland, 1999	1994-96	BCC; SCC	290	120 BCC; 25 SCC	20-75	52.4	Ever/never
Walther et al ²⁸	Germany, 2004	1997-99	BCC	624	213	19-92	48.1	>5 exposures per year‡
Zhang et al ²⁹	United States, 2012	1989-2009	BCC; SCC	73 494	5506 BCC; 403 SCC	25-62	0.0	Ever/never§
Studies not incl	uded in meta-analysis							
Boyd et al ¹⁸	United States, 2002	Unknown	BCC	60	30	20-40	0.0	No of exposures
Herity et al ²³	Ireland, 1989	1984-85	NMSC	792	202 BCC; 194 SCC	Unknown	Unknown	Ever/never
Hogan et al ²⁴	Canada, 1991	1989	Skin cancer¶	4820	791 BCC; 41 SCC; 15 MM**	10-56	37.3	Ever/never
O'Loughlin et al26	Ireland, 1985	Unknown	NMSC	242	58 BCC; 63 SCC	Unknown	71.9	"Often" exposed

*BCC=basal cell carcinoma; SCC=squamous cell carcinoma of skin; NMSC=non-melanoma skin cancer (BCC and SCC grouped together in publication); skin cancer (malignant melanoma, BCC, SCC all grouped together in publication).

†Ever/never: measurement of ever exposure to indoor tanning compared with never exposure; "regular" exposure: defined in publication as "regular" exposure to indoor tanning, with no further specifics; "often" exposed: defined in publication as "often" exposed to indoor tanning, with no further specifics.

 $\ddagger>5$ exposures to indoor tanning per year compared with ${\leq}5$ exposures per year.

§Ever/never comparison obtained directly from authors.

¶Skin cancer of head and neck.

**MM=malignant melanoma.

Table 2| Outcome effect sizes for studies included in primary meta-analysis

	Odds rat			
Reference	Basal cell carcinoma	Squamous cell carcinoma	Adjustments	
Studies comparing ever expos	sure with indoor tanning to never ex	kposure		
Asgari et al14	-	1.41 (0.90 to 2.22)	None (crude)	
Aubry and McGibbon ¹⁵	—	13.42 (1.38 to 130.48)	Age, sex, eye and hair colour, skin type*, ethnicity, sun exposure†	
Bajdik et al ¹⁶	1.2 (0.7 to 2.2)	1.4 (0.7 to2.7)	Age, (sex‡), hair colour, skin type*, ethnicity, sun exposure†	
Corona et al ¹⁹	0.6 (0.3 to 1.2)	_	Age, sex, skin type*, sun exposure†, family history of skin cancer	
Ferrucci et al ²⁰	1.69 (1.15 to 2.48)	_	Age, sex, body site, skin type*, sun sensitivity§, family history of skin cancer, melanocortin 1 receptor gene non-synonymous variants	
Gon and Minelli ²¹	0.31 (0.07 to 1.35)	_	Age, sex, eye colour, hair colour, skin type*, family history of skin cancer, presence of actinic keratosis	
Han et al ²²	1.32 (0.87 to 2.03)	1.44 (0.93 to 2.24)	Age, (sex‡), sun exposure†, sun sensitivity§, history of severe sunburns¶, geography**, family history of skin cancer	
Karagas et al ²⁵	1.5 (1.1 to 2.1)	2.5 (1.7 to 3.8)	Age, sex, sun sensitivity§††	
Rosso et al ²⁷	1.24 (0.53 to 2.88)	_	Age, sex	
Zhang et al ²⁹	1.29 (1.22 to 1.35)‡‡	1.50 (1.20 to 1.78)‡‡	Age, (sex‡), hair colour, sun exposure†, sun sensitivity§, history of severe sunburns¶, geography**, number of moles on legs, family history of melanoma	
Studies measuring higher dos	se exposure: "regular" exposure ¹⁷ o	r >5 exposures per year ²⁸		
Bakos et al ¹⁷	25.0 (2.26 to 277.36)		Superienuse parents' superienuse smoking	

Bakos et al ¹⁷	25.0 (2.26 to 277.36)	_	Sunscreen use, parents' sunscreen use, smoking
Walther et al ²⁸	0.7 (0.3 to 1.5)	_	None (crude)

*Skin type: complexion (Aubry and McGibbon), skin colour (Bajdik et al, Ferrucci et al), pigmentary traits (Corona et al).

†Sun exposure: non-occupational sun exposure (Aubry and McGibbon), lifetime occupational sun exposure (Bajdik et al), number of weeks spent at beach before age 20 and outdoor work (Corona et al), cumulative sun exposure while wearing bathing suit (Han et al), outdoor sun exposure (Zhang et al).

 $\ddagger Study not adjusted for sex because only included one sex.$

§Skin sensitivity to sun: skin response to first exposure of season to 1 hour of summer sun and to prolonged exposure to sun (Ferrucci et al), constitutional susceptibility score (Han et al), sun sensitivity (Karagas et al), childhood tendency to sunburn (Zhang et al).

(History of severe sunburns: lifetime sunburns which blistered (Han et al), number of severe sunburns between 15 and 20 (Zhang et al).

**Geography: geographic region at baseline (Han et al), ultraviolet index in area of residence at birth and 15 and 30 years of age (Zhang et al).

++Adjusted for age and sex in basal cell carcinoma analysis because adjustment for sun sensitivity did not change results.

 $\ddagger Hazard$ ratio (95% CI), obtained directly from authors.

Table 3| Outcome effect sizes for studies included in subset analyses of high dose exposure and young age at exposure

		Odds ra	atio (95% CI)			
Reference	Exposure/age	Basal cell carcinoma	Squamous cell carcinoma			
Studies assessing high dose exposure to indoor tanning						
Bakos et al17	"Regular use of tanning beds in lifetime"	25.0 (2.26 to 277.36)	—			
Ferrucci et al20	6-26 years of "regular" exposure	2.16 (1.34 to 3.48)	—			
Walther et al28	>5 exposures per year	0.7 (0.3 to 1.5)	—			
Zhang et al29	4 exposures per year	1.15 (1.11 to 1.19)*	1.15 (1.01 to 1.31)*			
Summary relative risk		1.50 (0.81 to 2.77)				
Studies assessing young age at exposure to indoor tanning						
Ferrucci et al20	≤16 years of age	1.83 (1.12 to 2.97)	—			
Karagas et al25	<20 years of age	1.8 (1.0 to 3.0)	3.66 (1.9 to 6.9)			
Zhang et al29	"High school/college"	1.38 (1.27 to 1.5)†	1.21 (0.85 to 1.71)†			
Summary relative risk		1.40 (1.29 to 1.52)	2.02 (0.70 to 5.86)			

*Hazard ratio (95% CI).

†Hazard ratio (95% CI) obtained directly from authors.

Figures

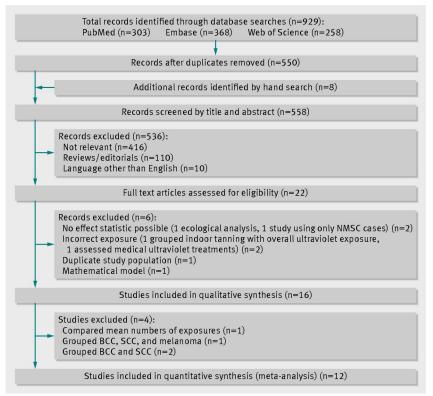


Fig 1 PRISMA flow diagram of literature search and study selection for meta-analysis of indoor tanning and non-melanoma skin cancer. BCC=basal cell carcinoma; NMSC=non-melanoma skin cancer; SCC=squamous cell carcinoma

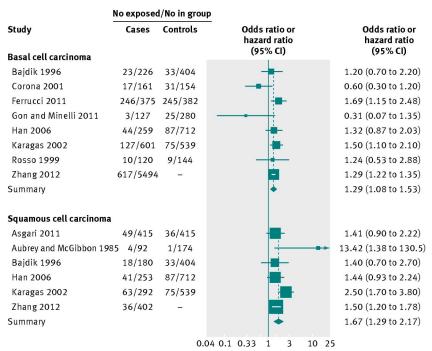


Fig 2 Relative risk of basal cell carcinoma and squamous cell carcinoma in participants ever exposed to indoor tanning compared with participants never exposed to indoor tanning