Preventing Breast Cancer in Our Daughters

A Community-Based Participatory Research Study of the Effect of Changing Personal Care Products on Healthy Human Breast Cells

Funded by the California Breast Cancer Research Program Pink Ribbon Day Peninsula Jewish Community Center October 27, 2019

Polly Marshall Breast Cancer Over Time polly@breastcancerovertime.org

Dedication



Luisa Preciutti Tumini



Luisa's daughter

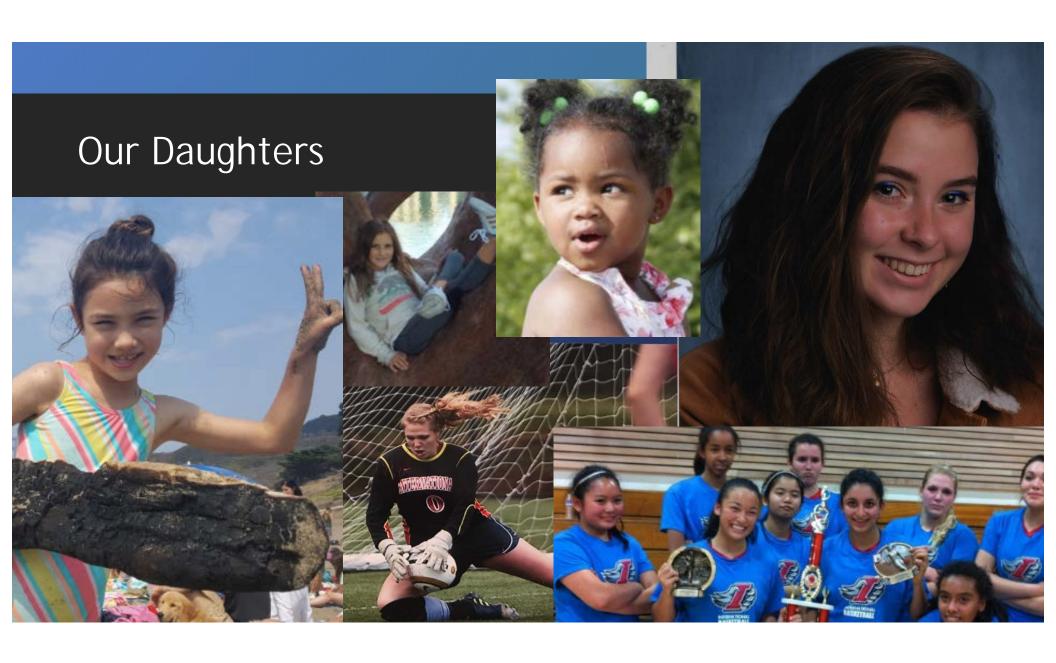
Breast Cancer Over Time



BCOT Steering Committee 2015

Created and controlled by breast cancer survivors to support and assist scientific research on the PREVENTION of breast cancer





Our Research Team



Dr. William Goodson, M.D. Breast Surgeon California Pacific Medical Center



Polly Marshall, J.D. Executive Director Breast Cancer Over Time

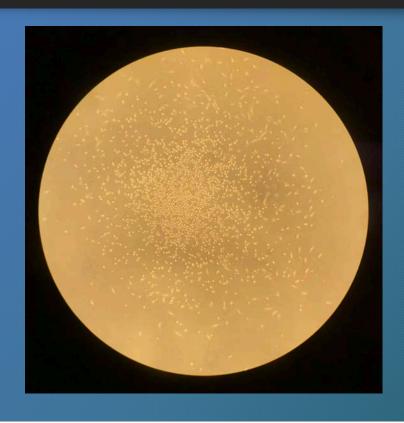


Dr. Shanaz Dairkee, Ph.D Senior Scientist California Pacific Medical Center Research Institute



Samantha Torres, MPH Study Coordinator Assistant Director Breast Cancer Over Time

Normal Human Breast Cells



Propagated as a breast cell culture in Dr. Shanaz Dairkee's lab at CPMCRI

Healthy Breast Cell Donors



Our Research Question

Does reduced exposure to common chemicals, known as xenoestrogens (XEs), in personal care products curb estrogenic hyper-signaling and its adverse effects on normal cell function within the healthy breast tissue of women volunteers participating in the XE-Low (XEL) intervention?

Preceding studies

HERMOSA Study - 2016

- Community-based participatory research study
- Measured phthalate, paraben, and phenol levels in urine of adolescent girls
- Significant drop after 3 day intervention
- Also funded by CBCRP



Reducing Phthalate, Paraben, and Phenol Exposure from Personal Care Products in Adolescent Girls: Findings from the HERMOSA Intervention Study

Kim G. Harley, Katherine Kogut, Daniel S. Madrigal, Maritza Cardenas, Irene A. Vera, Gonzalo Meza-Alfaro, Jianwen She, Qi Gavin, Rana Zahedi, Asa Bradman, Brenda Eskenazi, and Kimberly L. Parra

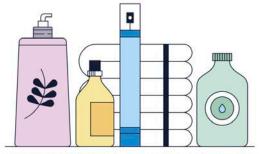
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Personal Care Products and Breast Cancer Risk



San Francisco-based breast cancer prevention study seeking volunteers who meet all the following criteria:

Are women 18-50 years old

Are premenopausal Have not had any kind of cancer (except basal cell skin cancer) Use selfcare products (such as shampoo, moisturizer, sunscreen, etc.)

Volunteers will participate in a healthy intervention in which they use parabenand phthalate-free self-care products provided by the study, and donate samples of blood, urine, and breast cells (via fine needle aspiration) in a medical office in San Francisco. Volunteers receive over \$200 worth of locally-made healthy cosmetics and \$40 in Peet's cards.

Sponsored by CPMC Research Institute and Breast Cancer Over Time



Interested?

Take our Intake Questionnaire at: bit.ly/breastcancerovertime



or contact Samantha Torres: 951.486.8285

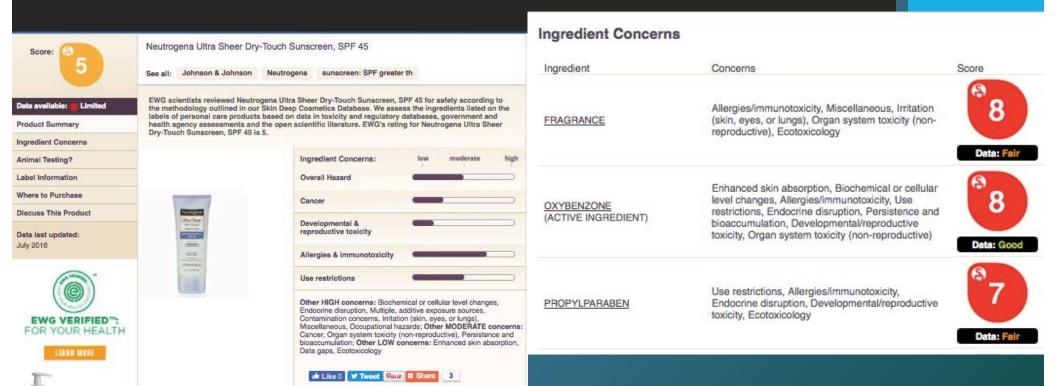
storres@breastcancerovertime.org

What xenoestrogens are in the products we use?





Neutrogena Ultra Sheer Dry-Touch Sunscreen, SPF 45



http://www.ewg.org/skindeep/product/593518/Neutrogena_Ultra_Sheer_Dry-Touch_Sunscreen%2C_SPF_45/#.WgCghBOPKT-

RETINYL PALMITATE (VITAMIN A PALMITATE)

Use restrictions, Developmental/reproductive toxicity, Biochemical or cellular level changes, Cancer, Organ system toxicity (non-reproductive)

9

Data: Fair

OXYBENZONE

Enhanced skin absorption, Biochemical or cellular level changes, Allergies/immunotoxicity, Use restrictions, Endocrine disruption, Persistence and bioaccumulation, Developmental/reproductive toxicity, Organ system toxicity (non-reproductive)



Data: Good

OCTINOXATE

Enhanced skin absorption, Biochemical or cellular level changes, Endocrine disruption,

Allergies/immunotoxicity, Persistence and bioaccumulation, Developmental/reproductive toxicity, Organ system toxicity (non-reproductive)



Data: Fair

HOMOSALATE

Enhanced skin absorption, Use restrictions, Organ system toxicity (non-reproductive), Endocrine disruption, Ecotoxicology, Contamination concerns (SALICYLIC ACID, TRIMETHYLCYCLOHEXANOL)



Patas Pata

Data Sources

Boehnlein J, Sakr A, Lichtin JL, Bronaugh RL. 1994. Characterization of esterase and alcohol dehydrogenase activity in skin. Metabolism of retinyl palmitate to retinol (vitamin A) during percutaneous absorption. Pharm Res 11(8): 1155-9.

CIR (Cosmetic Ingredient Review). 2006. CIR Compendium, containing abstracts, discussions, and conclusions of CIR cosmetic ingredient safety assessments. Washington DC.

Cherng SH, Xia Q, Blankenship LR, Freeman JP, Wamer WG, Howard PC, et al. 2005. Photodecomposition of retinyl palmitate in ethanol by UVA light-formation of photodecomposition products, reactive oxygen species, and lipid peroxides. Chem Res Toxicol 18(2): 129-38

Duell EA, Kang S, Voorhees JJ. 1997. Unoccluded retinol penetrates human skin in vivo more effectively than unoccluded retinyl palmitate or retinoic acid. J Invest Dermatol 109(3): 301-5.

EC (Environment Canada). 2008. Domestic Substances List Categorization. Canadian Environmental Protection Act (CEPA) Environmental Registry.

FDA (U.S. Food and Drug Administration) 2006. Food Additive Status List. Downloaded from http://www.cfsan.fda.gov/%7Edms/opa-appa.html, Oct 16, 2006.

Common Endocrine Disruptors in Cosmetics

- Propylparaben
- Butylparaben
- Isobutylparaben
- Methylparaben
- Ethylparaben
- Benzyl Salicylate
- Triclosan

- Oxybenzone
- Oxtinoxate
- Homosalate
- Cyclopentasiloxone
- BHT
- Lillial butylphenyl methylpropional

Other hazardous chemicals in cosmetics

- DMDM Hydantoin (formadehyde releaser)
- 2-Bromo-2-Nitropropane-1,3 Diol (formadehyde releaser)
- Retinyl palmitate (biochemical and cellular level changes)
- Methylisothiazolinone (human immune toxicant banned in Europe, Germany; restricted in Japan and Canada)
- Cocamide DEA (possible carcinogen, nitrosamine contamination)
- Octisalate (enhanced skin absorption)

Fragrance



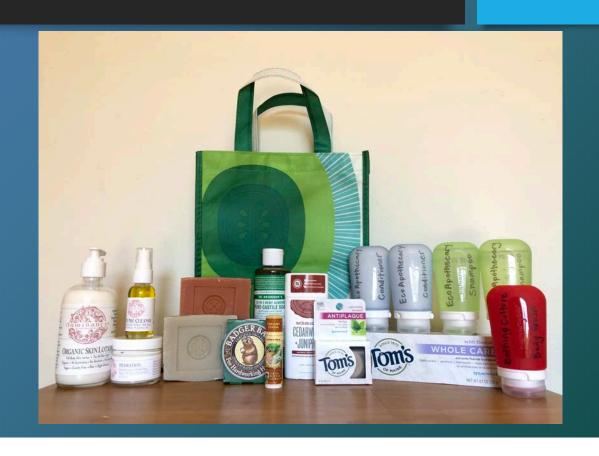
- "Trade secret" Ingredients not disclosed
- Very likely to include phthalates, per EWG

Our Study Protocol

- Obtain blood, urine, and breast cell samples before and after 28 day XE-low healthy intervention.
- 2. Test blood for levels of natural hormones
- 3. Test urine for paraben and phthalate levels
- Propagate live breast cells and perform tests to measure ERα activation, cell proliferation, and apoptosis signaling (higher levels of which are associated with increased breast cancer risk)

Our Healthy Intervention

- Participants used only XElow personal care products provided by Breast Cancer Over Time for 28 days between cell donations
- Participants kept logs of all products used for 28 days between cell donations



Analysis and Comparison of Samples

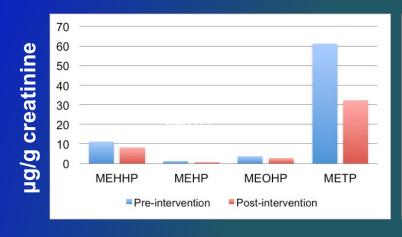
- Blood: analyzed for levels of natural hormones (estradiol, progesterone, sex hormone-binding globulin)
- Urine: analyzed for paraben and phthalate levels
- Breast cells: cultured and tests performed for functional differences in live cells

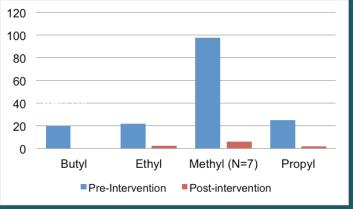
Results

- Participants: no attrition and positive reviews
- Blood: hormone levels not significantly different
- Urine: significant drops in paraben and phthalate levels after healthy intervention
- Cells: differences on functional tests on live cells measuring estrogen receptor isoform levels, cell proliferation, and cell death

Change in XE metabolite levels of urine samples 1 and 2 from XEL volunteers

POST-MENOPAUSAL (N=8)





Phthalates

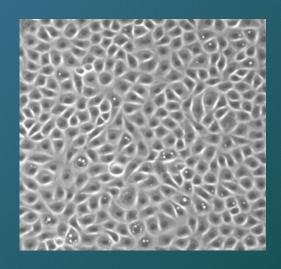
Parabens

Exit Surveys

- Very positive experience
- Liked the "healthy" personal care products
- Liked having breast cancer survivor buddies
- Reported starting to read ingredient labels in cosmetics
- Gratified and proud to have donated breast cells for a scientific study
- Felt they were making a difference
- FNAs "not a big deal" they would do it again

Study Conclusions

- Women will donate normal breast cells for research
- Volunteers will comply with study protocol
- Positive changes to human breast cells <u>are</u> observable after both 14 and 28 day interventions
- Study of responses of HEALTHY LIVING HUMAN BREAST CELLS to environmental exposures is feasible and can provide important information on human breast carcinogenesis



Additional Community Conclusions

- Women want to help in scientific research
- Their participation is meaningful and educational
- Survivors are fantastic recruiters and supporters
- More stringent government regulation of environmental chemicals, including cosmetic products, is essential to public health and must be demanded by all of us!



New Full Study: In Vivo Impact of Estrogen Exposure on the Human Breast

- Funded by California Breast Cancer Research Program (cigarette tax monies)
- 60 participants: 40 "Intervention" and 20 "Controls"
- Nanostring gene expression test added for all participants, both visits
- To volunteer, email Samantha Torres at storres@breastcancerovertime.org or scan our QR code:



Current Legislation to Support

- California: The Cosmetic Fragrance and Flavor Ingredient Right to Know Act of 2019 (SB 574)
 - Requires cosmetics companies to report toxic fragrance and flavor ingredients in their products to the California Safe Cosmetics Database within the California Department of Public Health
- Federal: <u>Safe Cosmetics and Personal Care Act of 2019</u> (H.R. 4296)
 - Requires full fragrance and flavor ingredient disclosure on a company's website and disclosure of a product's toxic fragrance and flavor ingredients on the product label
 - Bans from cosmetics 20 chemicals known to be toxic

Acknowledgements

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- CPMC Research Institute
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- San Francisco Public Health Foundation

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