

Brussels, January 30, 2021











Oncogenetics / Supportive care / Risk prevention

Pr François Duhoux Medical Oncology and Center for Human Genetics Cliniques universitaires Saint-Luc



Financial Disclosure

My institution has received payments from the following pharmaceutical companies on my behalf:

- Amgen
- AstraZeneca
- Eli Lilly
- mundipharma
- Novartis
- Pfizer
- Roche
- Teva





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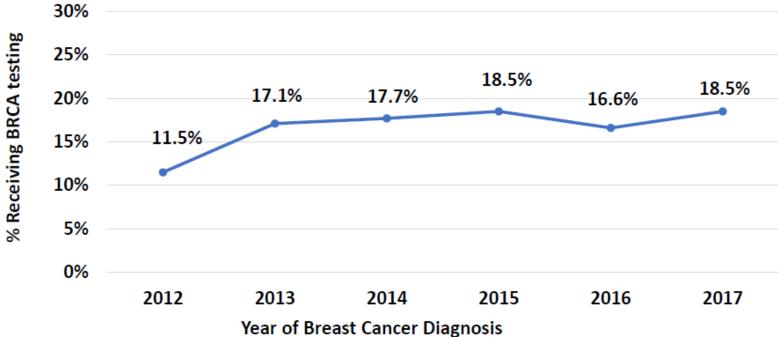


PS7-78: Trends in BRCA Testing Among Patients Diagnosed with Breast Cancer - A Retrospective Analysis of a United States Commercial Claims Database from the PRIOR-1 Study

Shelby L Corman¹, Hrishikesh Kale¹, Pooja Shah¹, Gboyega Adeboyeje² 1. Pharmerit – an OPEN Health Company; 2. Merck & Co. Inc.



Trend in prevalence of receiving a BRCA test





Real-world clinical outcomes of patients with BRCA-mutated (BRCAm) HER2-negative metastatic breast cancer: a CancerLinQ[®] study

Robert S Miller,¹ Stella Mokiou,² Aliki Taylor,² Miao Jiang,³ Ping Sun,² Susan McCutcheon²

CancerLinQ, American Society of Clinical Oncology, Alexandria, VA, USA; *AstraZeneca, Cambridge, UK; *AstraZeneca, Gaithersburg, MD, USA

PS7-66

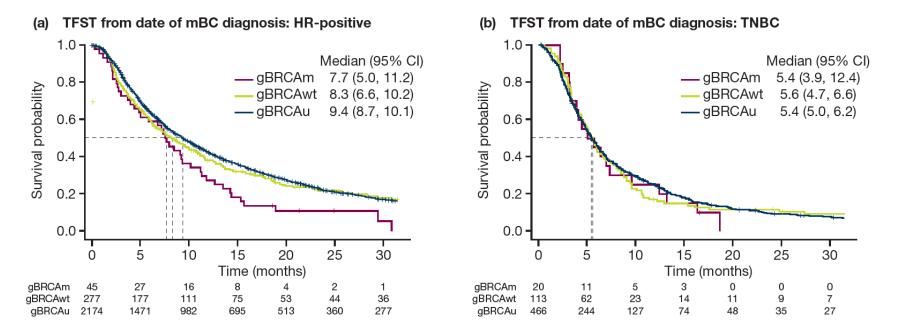
- Retrospective cohort study of patients diagnosed with HER2-negative mBC between January 1, 2010 and December 31, 2018, who were treated in routine clinical practice and included in the CancerLinQ Discovery database.
- Patients were grouped according to:
 - recorded gBRCA status: gBRCAm, gBRCA wild type (gBRCAwt) or unknown gBRCA (gBRCAu); and

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ABC

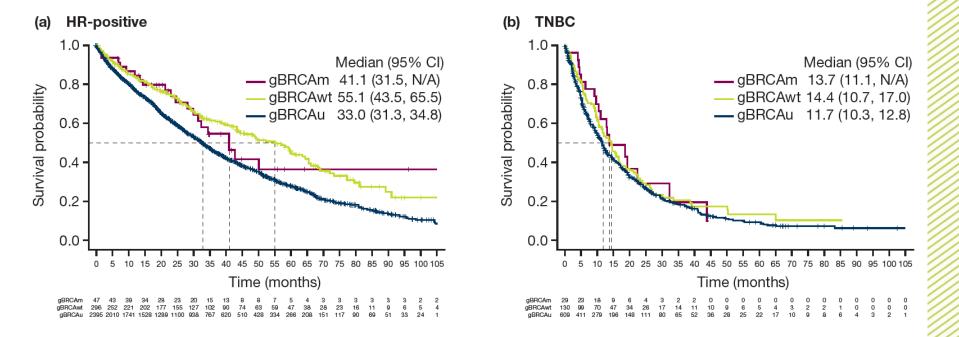
- hormone receptor (HR) status: HR-positive or triple-negative breast cancer (TNBC).
- Primary objective: time to first subsequent treatment (TFST), calculated from: 1) date of mBC diagnosis; and 2) date of initiation of first-line treatment for mBC.
- Secondary objective: real-world overall survival (OS), calculated from date of mBC diagnosis.
- TFST and OS (in months) were estimated using Kaplan–Meier methods; median (95% confidence intervals) are reported.





Median TFST was similar irrespective of gBRCA status when patients were stratified by HR status.





Median OS was similar irrespective of gBRCA status, when stratified by HR status



San Antonio Breast Cancer Symposium – December 8-12, 2020



Exploring the causal role of the human gut microbiome in breast cancer risk using Mendelian randomization (MR)

Tim Robinson, Grace Edmunds, Bryony Hayes and Kaitlin Wade

NIHR Academic Clinical Lecturer in Medical Oncology CRUK-ICEP University of Bristol, UK @drtimrobinson





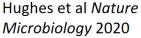
Breast Cancer Risk GWAS

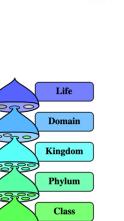
- Breast Cancer Association Consortium (BCAC) and Consortium of Investigators of Modifiers of BRCA1/2 (CIMBA)
- Overall breast cancer: 133 384 cases and 113 789 controls
- Subtype: 106 278 cases, 91 477 controls
- TNBC meta-analysis: 18 016 cases and 100 971 controls

Zhang et al Nature Genetics 2020

Microbiome GWAS

- Harmonized analytical pipeline with taxonspecific models
- Flemish Gut Flora Project (FGFP, n=2223)
- Food-Chain Plus (FoCus, n=950)
- PopGen (n=717)





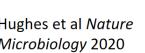
Order

Family

Genus

Species







Results – Microbial traits affect overall breast cancer risk

 G. Ruminococcus: OR 1.03 (1.00-1.06), p=0.043

C. Gammaproteobacteria F. Sutterellaceae G. Bifidobacterium G. Butyricicoccus G. Dialister	
G. Parabacteroides	
G. Ruminococcus	
C. Unalgosified F. Emisinal atrich geogra	
G. Unclassified F. Erysipelotrichaceae G. Unclassified F. Porphyromonadaceae G. Unclassified O. Bacteroidales G. Unclassified P. Firmicutes G. unclassified P. Firmicutes G Veillonella	

Odds Ratio Overall Breast Cancer Risk



Results – Microbial traits affect TNBC breast cancer risk

- G. Parabacteroides: OR 0.84 (0.72-0.98), p=0.031
- G. Unclassified O. Bacteroidales: OR 1.12(1.03-1.22), p= 0.012

C. Gammaproteobacteria F. Sutterellaceae G. Bifidobacterium						
G. Butyricicoccus						
G. Dialister			-	-		
G. Parabacteroides	—	-				
G. Ruminococcus				-		
G. Unclassified F. Erysipelotrichaceae						
G. Unclassified F. Porphyromonadaceae	?					
G. Unclassified O. Bacteroidales			-	-	_	
G. Unclassified P. Firmicutes				-		
G. unclassified P. Firmicutes				-		
G Veillonella				—		
	0.7	0.8	0.9 1	1.1 1	.2 1	1 .3

Odds Ratio TNBC Cancer Risk





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Targeting Depressive Symptoms in Younger Breast Cancer Survivors: A RCT of Mindfulness Meditation and Survivor Education

PA Ganz, JE Bower, AH Partridge, AC Wolff,

ED Thorner, H Joffe, MR Irwin, L Petersen, CM Crespi

UCLA Jonsson Comprehensive Cancer Center, Los Angeles; Dana Farber Cancer Institute, Boston; Johns Hopkins Medicine & Sidney Kimmel Comprehensive Cancer Center, Baltimore

Pathways to Wellness Study, funded by NCI R01 CA200977 & BCRF



Pathways to Wellness (PTW) Randomized Trial ClinicalTrials.gov NCT03025139

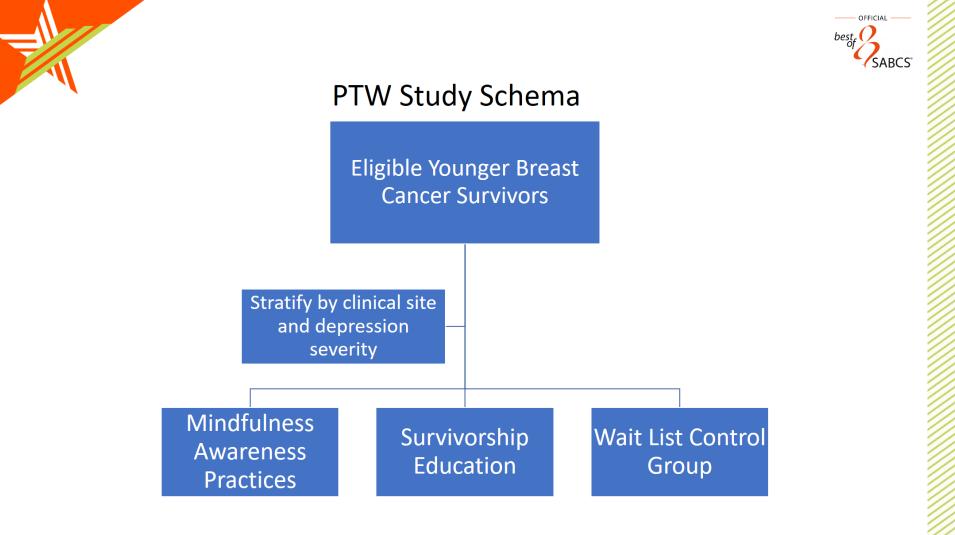
- Phase III, randomized, multi-institution trial testing efficacy of two interventions targeting depressive symptoms in younger breast cancer survivors
- Interventions were compared to a concurrent wait list control (WLC) group
- Interventions were 6-week group programs
 - Mindfulness Awareness Practices (MAPS)
 - Survivorship Education (SE)
- Site instructors delivered 2-hour structured content at each session with monitoring of fidelity

→ Inclusion of 247 patients

Intervention content - 6 group sessions

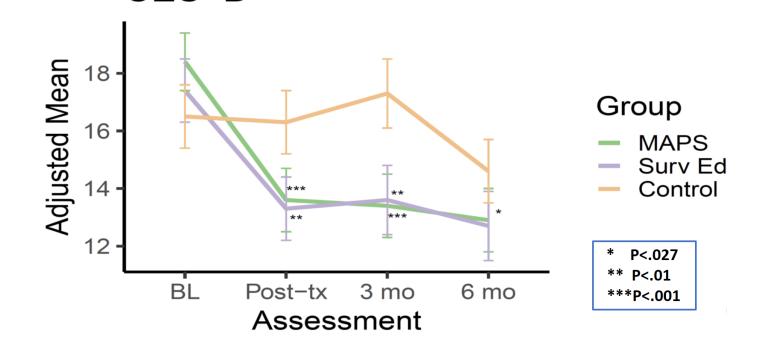


Mindfulness Awareness Practices (MAPS)	Survivorship Education (SE)
What is mindfulness? Listening, Embodiment and Obstacles Working with pain Working with Difficult Emotions, Cultivating Positive Emotions Working with Thoughts, Mindful Interactions Wrap-up	Breast cancer 101: important issues for younger survivors QOL in breast cancer survivors Energy balance, Nutrition, Physical Activity Cancer in the Family: Cancer Genetics and Testing Relationships and Work-Life Balance Body image, menopause, sexual health; moving forward





PTW Depression Outcomes CES-D

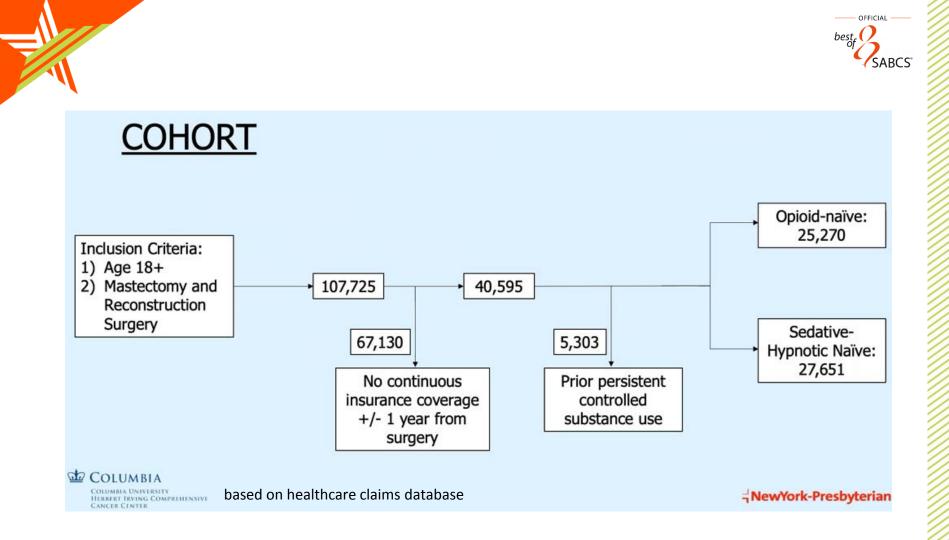




Persistent Controlled Substance Use Following Mastectomy with Reconstruction Surgery

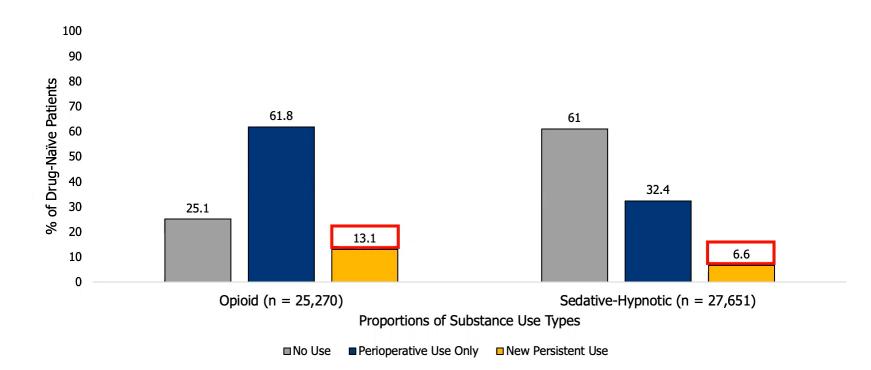
Jacob Cogan MD, Rohit Raghunathan MS, Melissa Beauchemin PhD RN, Melissa Accordino MD, Alexander Melamed MD, Jason Wright MD, and Dawn Hershman MD

HERBERT IRVING COMPREHENSIVE CANCER CENTER Columbia University Medical Center



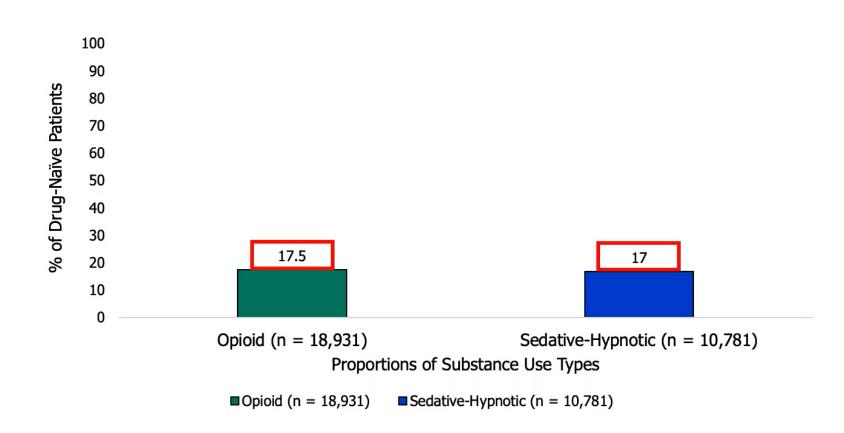


Rates of New Persistent Controlled Substance Use





Persistent Controlled Substance Use Among Those Who Initiate





Chances of Pregnancy after Breast Cancer, Reproductive and Disease Outcomes: a Systematic Review and Meta-analysis

Eva Blondeaux^{1,2}, Marta Perachino^{1,2}, Marco Bruzzone¹, Richard A. Anderson³, Evandro de Azambuja⁴, Philip D. Poorvu⁵, Hee Jeong Kim⁶, Cynthia Villarreal-Garza⁷, Barbara Pistilli⁸, Ines Vaz-Luis⁸, Cristina Saura⁹, Kathryn J. Ruddy¹⁰, Maria Alice Franzoi⁴, Chiara Sertoli², Marcello Ceppi¹, Hatem A. Azim Jr.⁷, Frederic Amant¹¹, Isabelle Demeestere¹², Lucia Del Mastro^{1,2}, Ann H. Partridge⁵, Olivia Pagani¹³, Fedro A. Peccatori¹⁴, Matteo Lambertini^{1,2}

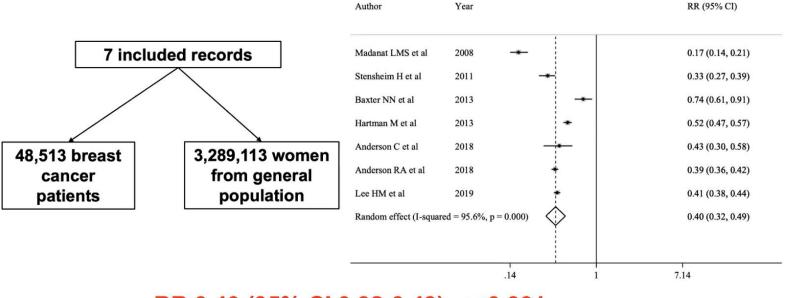
¹IRCCS Ospedale Policlinico San Martino, Genova, Italy. ²University of Genova, Genova, Italy. ³MRC Centre for Reproductive Health, The Queen's Medical Research Institute, The University of Edinburgh, Edinburgh, UK. ⁴Institut Jules Bordet, Université Libre de Bruxelles (U.L.B.), Brussels, Belgium. ⁵Dana-Farber Cancer Institute, Boston MA, USA. ⁶Asan Medical Center, Seoul, Korea. ⁷Hospital Zambrano Hellion, Tecnologico de Monterrey, San Pedro Garza Garcia, Nuevo Leon, Mexico. ⁸Institut Gustave Roussy, Villejuif, France. ⁹Vall d'Hebron University Hospital (HUVH), Vall d'Hebron Institute of Oncology (VHIO), Barcelona, Spain. ¹⁰Mayo Clinic, Rochester, Minnesota, USA. ¹¹Netherlands Cancer Institute and Amsterdam University Medical Centers, Amsterdam, The Netherlands. ¹²Erasme Hospital, Université Libre de Bruxelles (U.L.B.), Brussels, Belgium. ¹³Medical School, Geneva University Hospital, Geneva, Switzerland, European School of Oncology, Milan Italy. ¹⁴European Institute of Oncology IRCCS, European School of Oncology, Milan, Italy

GS3-09





Chances of Pregnancy Breast Cancer Survivors



RR 0.40 (95% CI 0.32-0.49); p<0.001



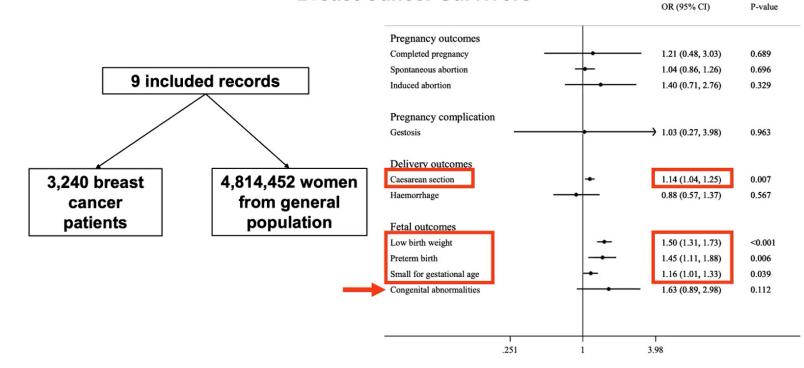
Chances of Pregnancy All Cancer Survivors

Diagnosis			RR (95% CI)	P-value
Cervical cancer	•		0.33 (0.31, 0.35)	<0.001
Breast cancer			0.40 (0.32, 0.49)	<0.001
Leukemia			0.40 (0.27, 0.58)	<0.001
Kidney cancer	< <u>∗</u>		0.42 (0.18, 0.99)	0.047
CNS cancer			0.52 (0.39, 0.69)	< 0.001
Bone cancer			0.56 (0.37, 0.86)	0.008
Ovarian cancer	-		0.56 (0.48, 0.65)	< 0.001
Hodgkin lymphoma	— <u>—</u>		0.62 (0.47, 0.82)	0.001
All cancers			0.65 (0.55, 0.77)	<0.001
Liver cancer			0.65 (0.19, 2.26)	0.500
Non-Hodgkin lymphoma			0.66 (0.53, 0.82)	< 0.001
Colon cancer		-	0.70 (0.41, 1.17)	0.171
Thyroid cancer			0.82 (0.65, 1.03)	0.094
Skin cancer	-	+	0.97 (0.87, 1.09)	0.636
			1	
	.18	5.	.56	





Reproductive Outcomes Breast Cancer Survivors







Reproductive Outcomes Breast Cancer Survivors

Subgroup analysis	Low birth weight	Preterm birth	Small for gestational age				
	PR (95%CI)	PR (95%CI)	PR (95%CI)				
Treatment received							
Chemotherapy	1.62 (1.08-2.42)	1.60 (0.84-3.05)	1.51 (1.22-2.88)				
No chemotherapy	1.05 (0.77-1.43)	1.16 (0.90-1.49)	0.85 (0.53-1.36)				
Pregnancy interval*							
Early pregnancy	1.47 (0.66-3.28)	1.59 (0.60-4.24)	1.13 (0.75-1.69)				
Late pregnancy	1.25 (0.97-1.62)	1.21 (0.96-1.53)	1.34 (1.05-1.71)				

*The cut-off used in the studies ranged between 2 years and 5 years





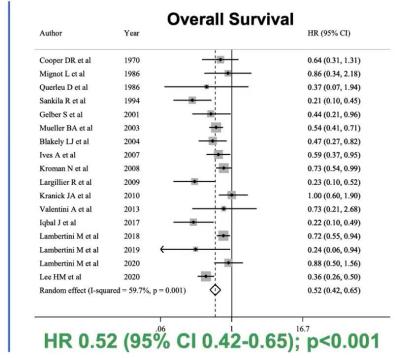
Maternal Safety

Studies Adjusting for the Potential «Guarantee-time Bias»

Author Year HR (95% CI) Dow KH et al 1994 0.60 (0.33, 1.08) von Schoultz E et al 1995 0.48 (0.18, 1.29) Blakely LJ et al 2004 0.70 (0.25, 1.95) Largillier R et al 2009 0.65 (0.36, 1.17) Kranick JA et al 2010 1.20 (0.80, 2.00) Lambertini M et al 2018 0.85 (0.68, 1.06) Lambertini M et al 2019 1.12 (0.52, 2.42) Lambertini M et al 2020 0.87 (0.61, 1.23) Lee HM et al 2020 0.49 (0.40, 0.60) Random effect (I-squared = 66.6%, p = 0.002) 0.74 (0.58, 0.96) .18 5.56

Disease-free Survival

HR 0.74 (95% CI 0.58-0.96); p=0.023







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Glycemic Index, Glycemic Load and breast cancer risk: results from the prospective NutriNet-Santé cohort

Charlotte Debras, Eloi Chazelas, Bernard Srour, Chantal Julia, Emmanuelle Kesse-Guyot, Laurent Zelek, Cédric Agaësse, Nathalie Druesne-Pecollo, Valentina A. Andreeva, Pilar Galan, Serge Hercberg, Paule Latino-Martel, Mélanie Deschasaux-Tanguy, Mathilde Touvier

Nutritional Epidemiology Research Team (EREN)

UMR U1153 Inserm / U1125 Inrae / Cnam / Sorbonne Paris Nord Center of Research in Epidemiology and Statistics University of Paris

Funding: Institut National du Cancer (INCa) 2019 (Doctoral scholarship n°2019-158 – C Debras)







le cnam











- NutriNet-Santé cohort study
 - E-cohort France since 2009 (open recruitment) > 171,000 participants
 - Online self-administered questionnaires repeated every 6 months : sociodemographic, lifestyle, health status, physical activity, three 24h-dietary records
 - Cancer cases validated by an expert medical committee
- Study population
 - Exclusion criteria:
 - Prevalent cancer or T1D/T2D at baseline
 - <3 24h-dietary records over the first 2 years of follow-up
 - Null follow-up



	Cancer all sites (3131 cases)			Breast (927 cases)		
	HR Q5 vs. Q1	95% Cl	P	HR Q5 vs. Q1	95% CI	P
Average dietary GI	1.12	0.99-1.27	0.09	1.21	0.96-1.54	0.1
Dietary GL	1.25	1.03-1.52	0.008	1.29	0.90-1.84	2 0.1
Contribution of low GI foods to caloric intake	0.83	0.74-0.94	0.005	0.78	0.62-0.99	0.07
Contribution of medium-to-high GI foods to caloric intake	1.21	1.07-1.38	0.002	1.34	1.05-1.71	0.04
Contribution of low GI foods to carbohydrate intake	0.80	0.71-0.91	0.0006	0.74	0.59-0.92	2 0.02
Contribution of medium to high GI foods to carbohydrate intake	1.27	1.13-1.44	0.0001	1.48	1.18-1.86	2 20.002

Significant for premenopausal breast cancer (297 cases)

Significant for postmenopausal breast cancer (525 cases)

GI = glycemic index; GL = glycemic load

Association independent of BMI and weight gain : inflammation, oxidative stress, increased insulin levels, ...



Diabetes Risk Reduction Diet and Survival Following Breast Cancer

Tengteng Wang, PhD, MSPH, MBBS Postdoctoral Research Fellow Channing Division of Network Medicine, Brigham and Women's Hospital Harvard Medical School & Harvard School of Public Health







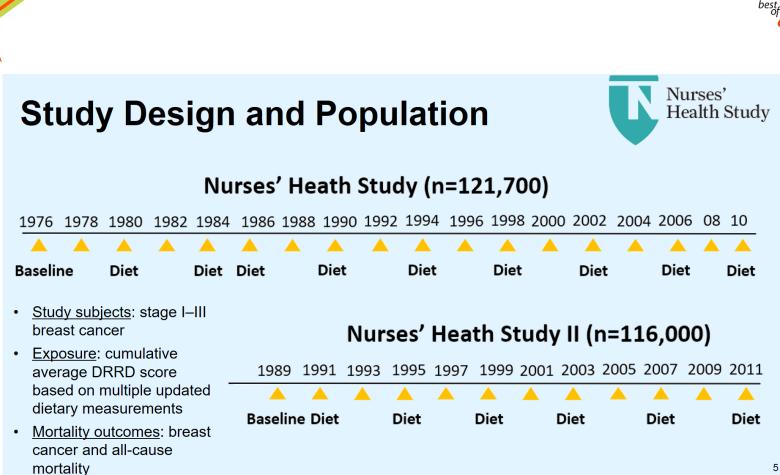
GS2-09





Diabetes risk reduction diet (DRRD) score

- Emphasizes nine different dietary factors:
 - Higher intakes of cereal fiber, coffee, nuts, polyunsaturated:saturated fat ratio, and whole fruits
 - Lower glycemic index, lower intakes of trans fat, sugar-sweetened beverages/fruit juices, and red meat

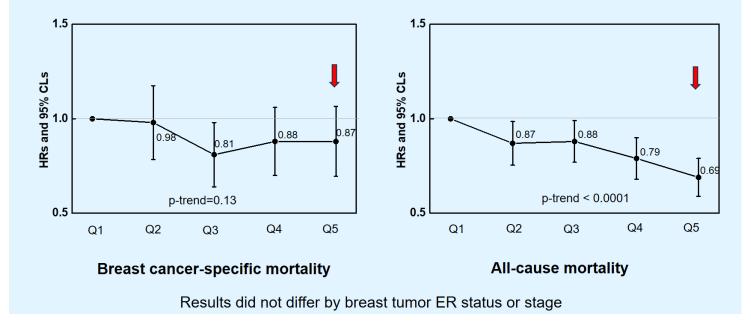


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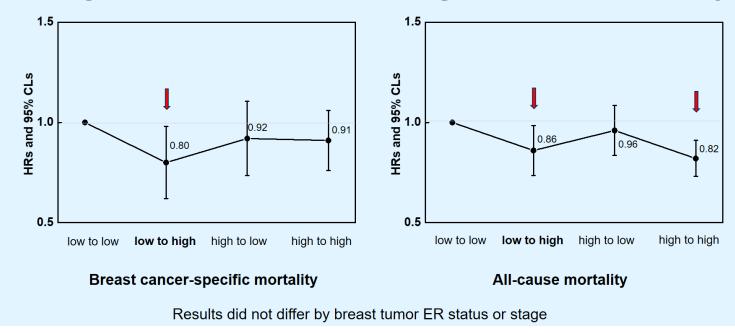
Results: Post-diagnosis DRRD and mortality



Results: Changes from pre- to post-diagnosis and mortality

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→ Promoting dietary changes consistent with prevention of T2D may be important for breast cancer survivors.

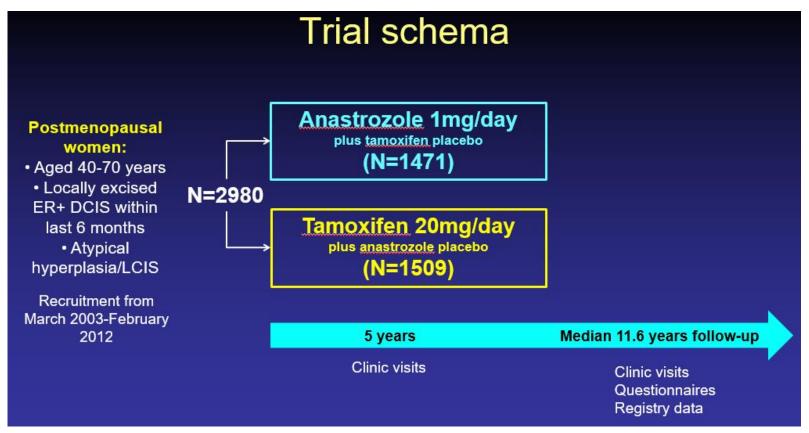
Updated long-term results of anastrozole versus tamoxifen for the prevention of breast cancer in postmenopausal women with locally excised Ductal Carcinoma In-Situ (IBIS-II DCIS) OFFICIAL

Ivana Sestak¹, Jack Cuzick¹, Bernardo Bonanni², Nigel Bundred³, <u>Christelle</u> Levy⁴, Sibylle Loibl⁵, Patrick Neven⁶, Michael Stierer⁷, Chris Holcombe⁸, Robert E. Coleman⁹, John F. Forbes¹⁰, Anthony Howell¹¹ on behalf of the IBIS-II investigators

Wolfson Institute of Preventive Medicine, Queen Mary University London, London, UK
Division of Chemoprevention and Genetics, European Institute of Oncology, Milan, Italy
South Manchester University Hospital, Manchester, UK
Centre François Baclesse, Caen, France
German Breast Group, Frankfurt, Germany
University of Leuven, Leuven, Belgium
Austrian Breast and Colorectal Cancer Study Group, Vienna, Austria
Royal Liverpool University Hospital, Liverpool, UK
Weston Park Hospital, Sheffield, UK
University of Newcastle, Newcastle, Australia
Nightingale Breast Screening Centre, Manchester University HS Foundation Trust, Manchester, UK

GS2-02







Breast cancer recurrence 10 9.7% (7.9-11.8) HR (95% CI) P-Recurrences 8.5% (6.9-10.5) value A=103 0.89 (0.69-1.16) 0.401 T=118 Recurrence (%) S. 0 8 10 Follow-up time [years] 12 14 16 2 6 0 4 Number at risk Tamoxifen 1489 1466 1436 1403 1367 1087 674 275 24 1449 1434 1375 1342 1072 641 272 24 Anastrozole 1399 Tamoxifen Anastrozole

	Invasive			DCIS		
	A vs. T	HR (95% CI)	P-value	A vs. T	HR (95% CI)	P-value
All	66 vs. 76	0.89 (0.64-1.24)	0.48	36 vs. 42	0.88 (0.56-1.37)	0.56
Ipsilateral	37 vs. 39	0.97 (0.62-1.52)	0.9	23 vs. 25	0.94 (0.53-1.66)	0.83
Contralateral	28 vs. 33	0.87 (0.52-1.43)	0.57	12 vs. 15	0.82 (0.38-1.75)	0.61



San Antonio Breast Cancer Symposium® - December 8-11, 2020 Influence of physician's lifestyle on the prescription of healthy habits to breast cancer patients (LACOG 1218)

Renata Cangussú¹, Eldsamira Mascarenhas¹, Taiane F Rebelatto², Paulo R Nunes², Rafaela G de Jesus², Facundo Zaffaroni² and Gustavo Werutsky². ¹Oncologia D`or - Hospital Cardiopulmonar, Salvador, Brazil ² Latin American Cooperative Oncology Group, Porto Alegre, Brazil

PS7-91





MULTIVARIABLE ANALYSIS

Table 1. Physicians characteristics and advice on the importance of lifestyle modification for BC patients

Parameter	Levels	Do not advice about life style (%)	Relative Risk	95% confidence interval	p-value
Do you personall	y practice physical activity?				0.0265
	Yes	31 (13.1)	1.00		
	No	11 (35.5)	2.48	1.28 to 4.82	

Physicians characteristics and reference to dietitian or endocrinologist on obesity management for BC patients



More likely to refer if :

Parameter	Levels	Do not refer to a dietitian or endocrinologist (%)	Relative Risk	95% confidence interval	p-value
Gender					0.0296
	Female	66 (42.3)	1.00		
	Male	53 (47.7)	1.35	1.03 to 1.76	
Age (years)					0.0005
	30 - 49	105 (49.3)	1.00		
	≥ 50	14 (25.9)	0.46	0.28 to 0.75	
Do you treat l	breast cancer?				0.0692
	Yes	114 (43.8)	1.00		
	No	5 (71.4)	2.38	1.33 to 4.26	
Do you perso	nally follow the recomn	nendations of eating habits	17		0.4311
	Yes	100 (43.9)	1.00		
	No	19 (48.7)	1.15	0.83 to 1.60	
Do you perso	nally practice physical a	ctivity?			0.3851
	Yes	103 (43.6)	1.00		
	No	16 (51.6)	1.18	0.82 to 1.71	
Do you consu	me alcohol?				0.1320
	Yes	101 (47.6)	1.00		
	No	18 (32.7)	0.75	0.50 to 1.11	
Do you smoke	e?				0.1488
	Yes	2 (25.0)	1.00		
	No	117 (45.2)	1.84	0.69 to 4.94	
Avarage sleep	time per day				0.4283
	< 5h	6 (54.5)	1.00		
	Between 5 - 7h	16 (36.4)	0.59	0.30 to 1.17	
	>7h	97 (45.7)	0.66	0.38 to 1.17	
What is your I	BMI?				0.1031
	<25	83 (47.7)	1.00		
	≥25	36 (38.7)	0.80	0.61 to 1.05	

MULTIVARIABLE ANALYSIS