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Nutrizione nella donna con carcinoma mammario

Maurizio Muscaritoli

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Nessun conflitto di interesse da dichiarare relativamente a questa presentazione

Department of Translational and Precision Medicine




Epidemiologia del carcinoma mammario

Estimated New Cases



Breast	266,120	30%
Lung & bronchus	112,350	13%
Colon & rectum	64,640	7%
Uterine corpus	63,230	7%
Thyroid	40,900	5%
Melanoma of the skin	36,120	4%
Non-Hodgkin lymphoma	32,950	4%
Pancreas	26,240	3%
Leukemia	25,270	3%
Kidney & renal pelvis	22,660	3%
All Sites	878,980	100%

Estimated Deaths



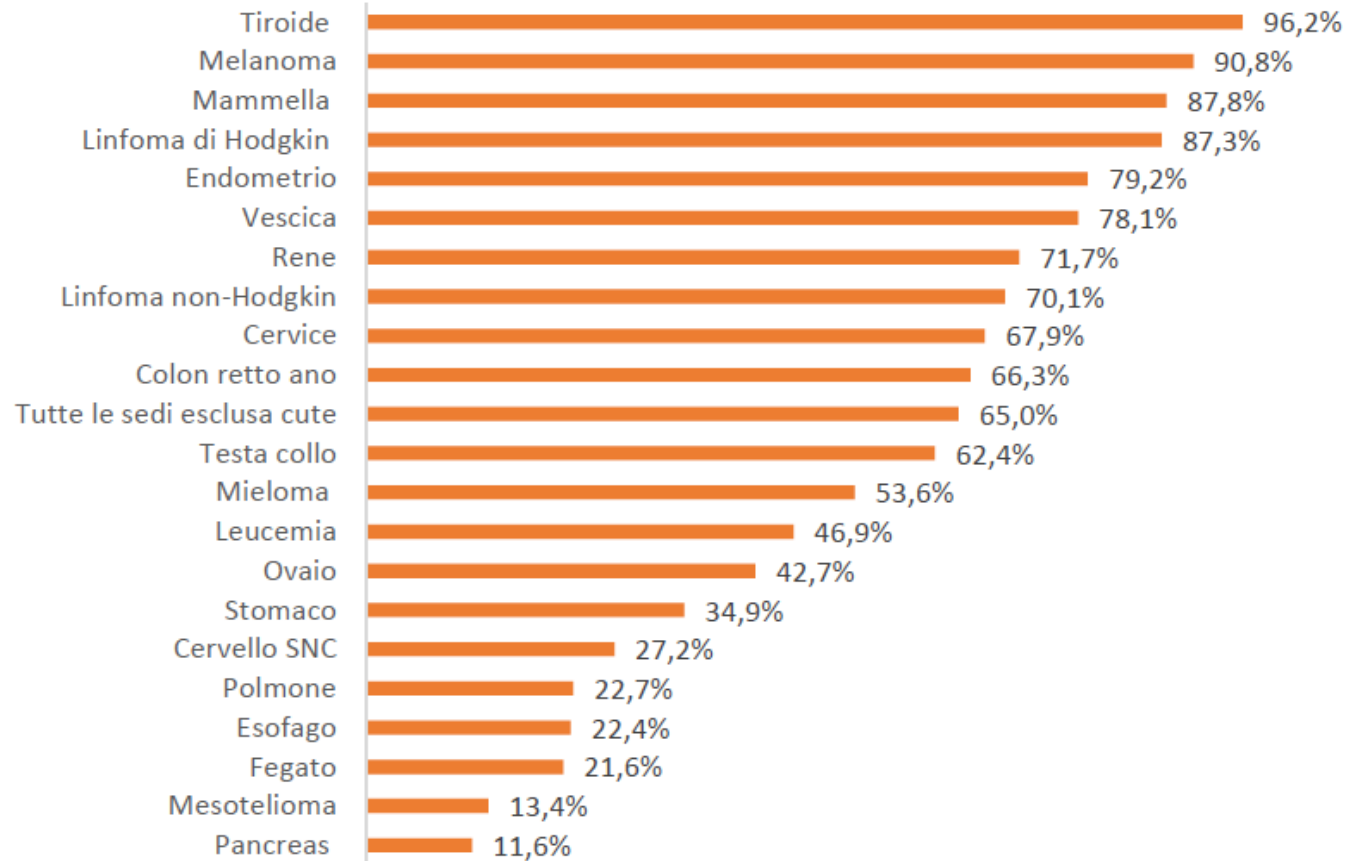
Lung & bronchus	70,500	25%
Breast	40,920	14%
Colon & rectum	23,240	8%
Pancreas	21,310	7%
Ovary	14,070	5%
Uterine corpus	11,350	4%
Leukemia	10,100	4%
Liver & intrahepatic bile duct	9,660	3%
Non-Hodgkin lymphoma	8,400	3%
Brain & other nervous system	7,340	3%
All Sites	286,010	100%

In Italia **nel 2020** 55.000 nuove diagnosi di carcinoma della mammella femminile



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Figura 2. Sopravvivenza netta a 5 anni dalla diagnosi.
Femmine, età ≥ 15 anni
Casi incidenti nel periodo 2010-2014, con follow-up al 2018
(fonte dei dati: "I numeri del cancro in Italia 2021")



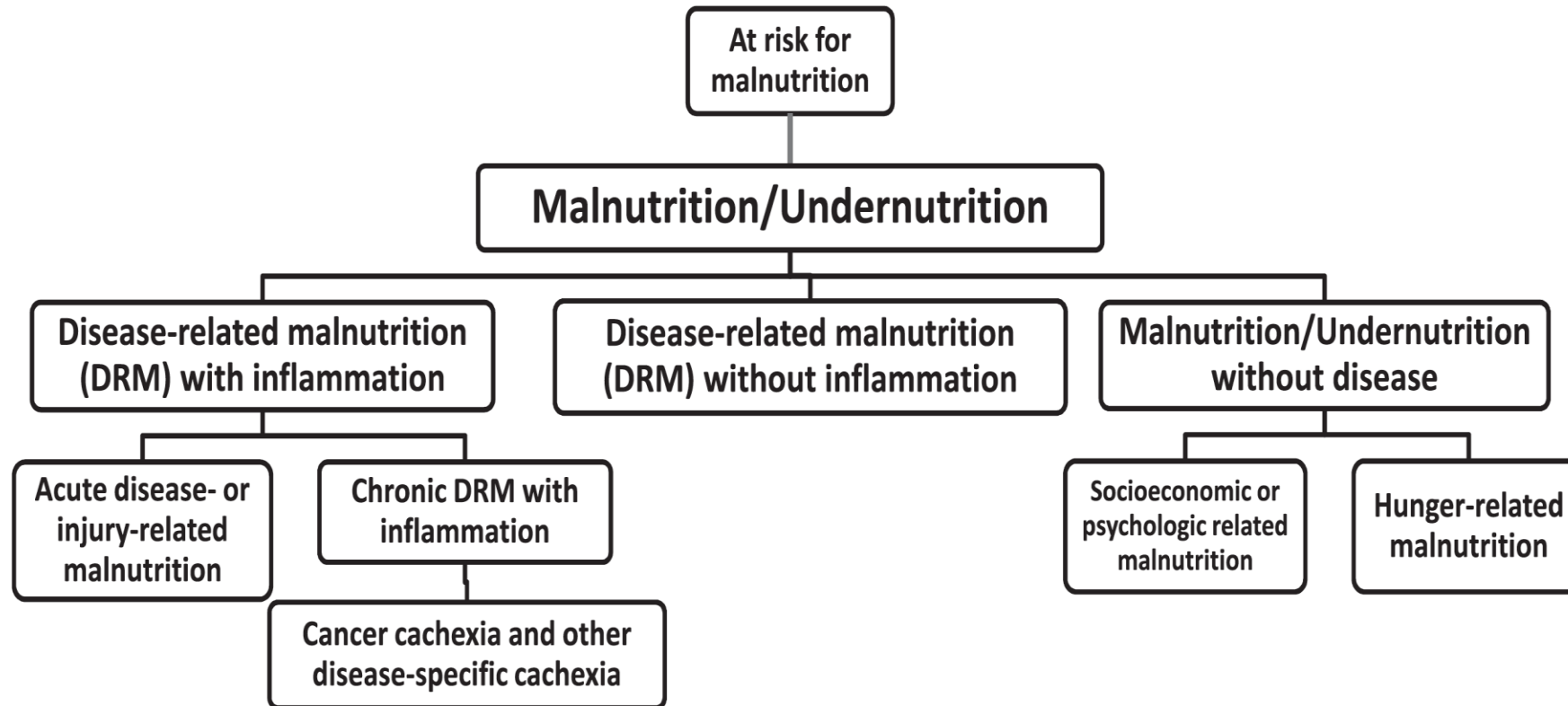
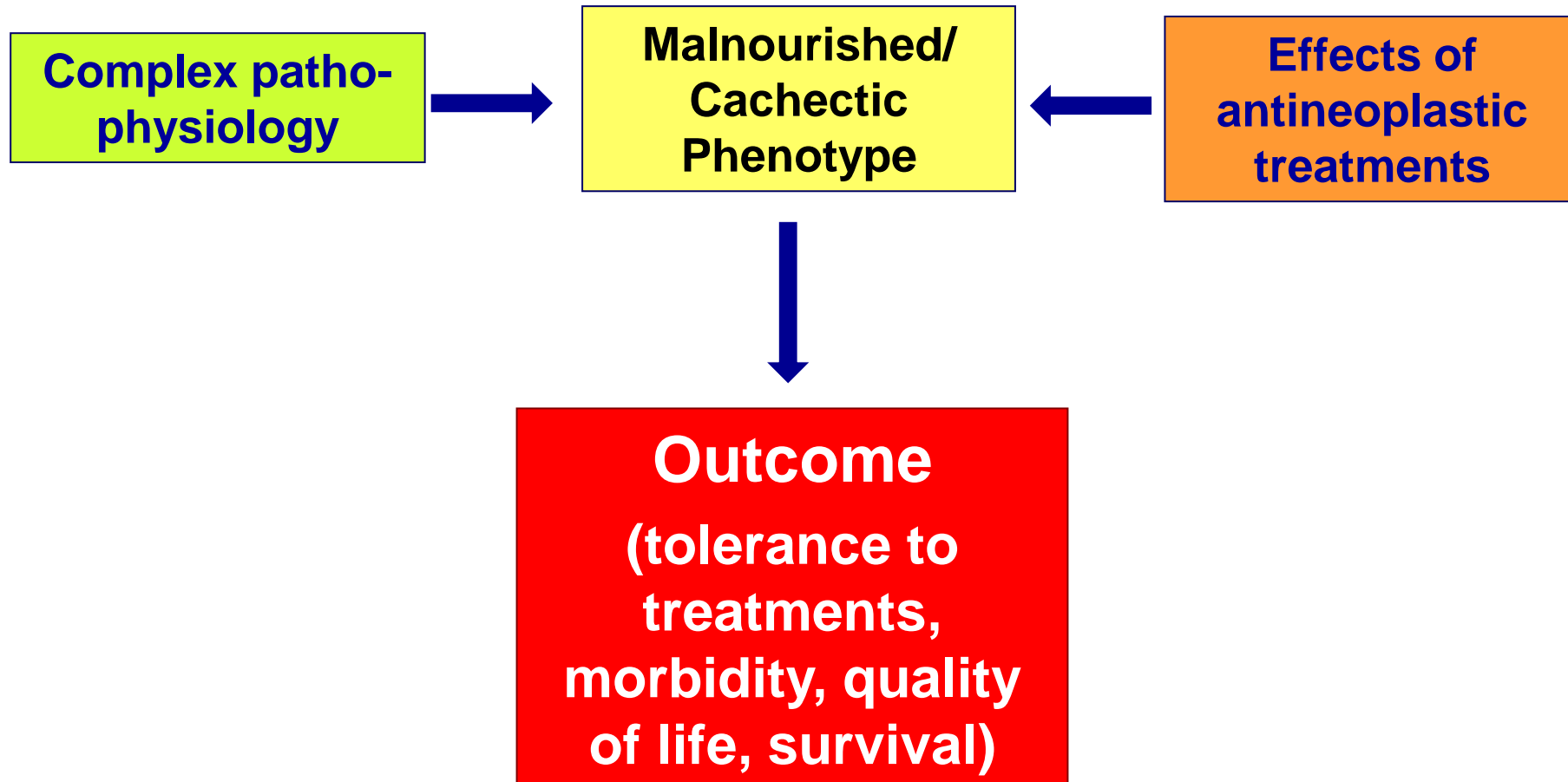


Fig. 2. Diagnoses tree of malnutrition; from at risk for malnutrition, basic definition of malnutrition to aetiology-based diagnoses

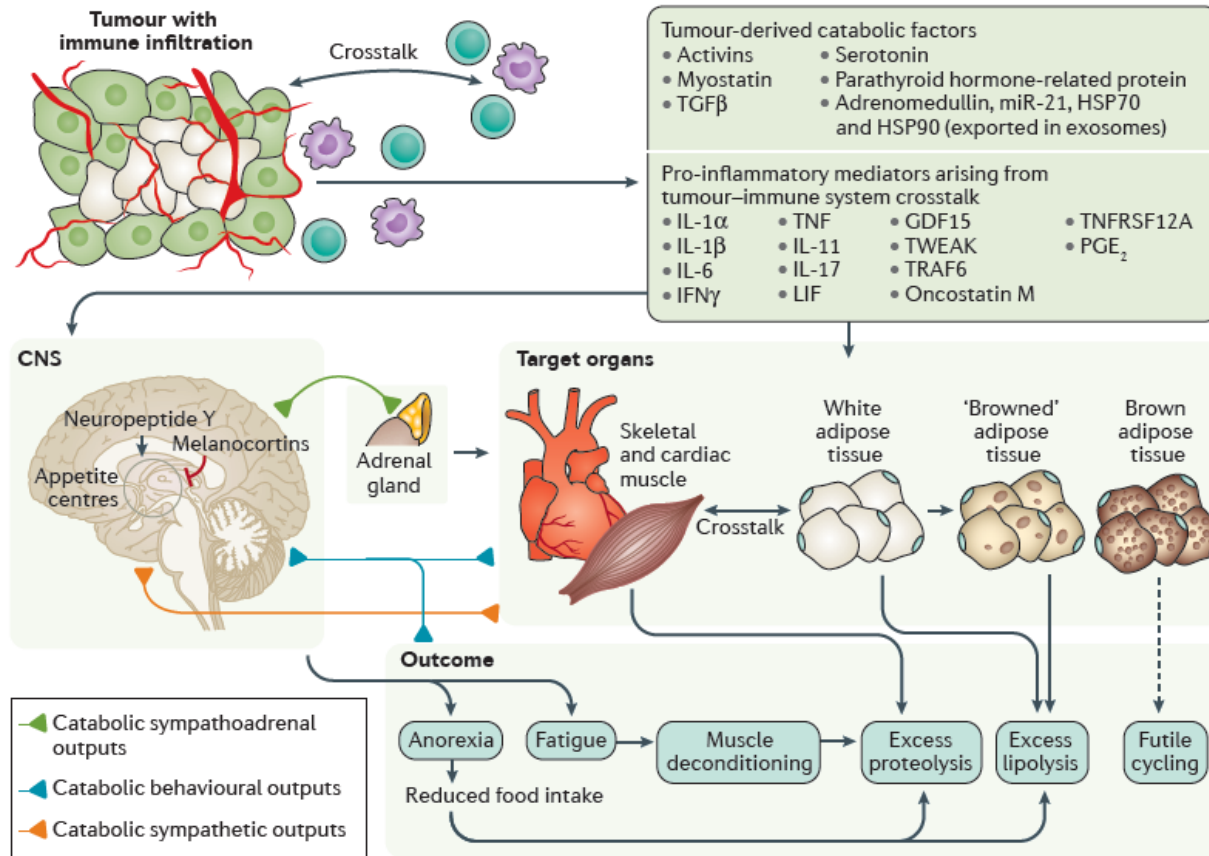


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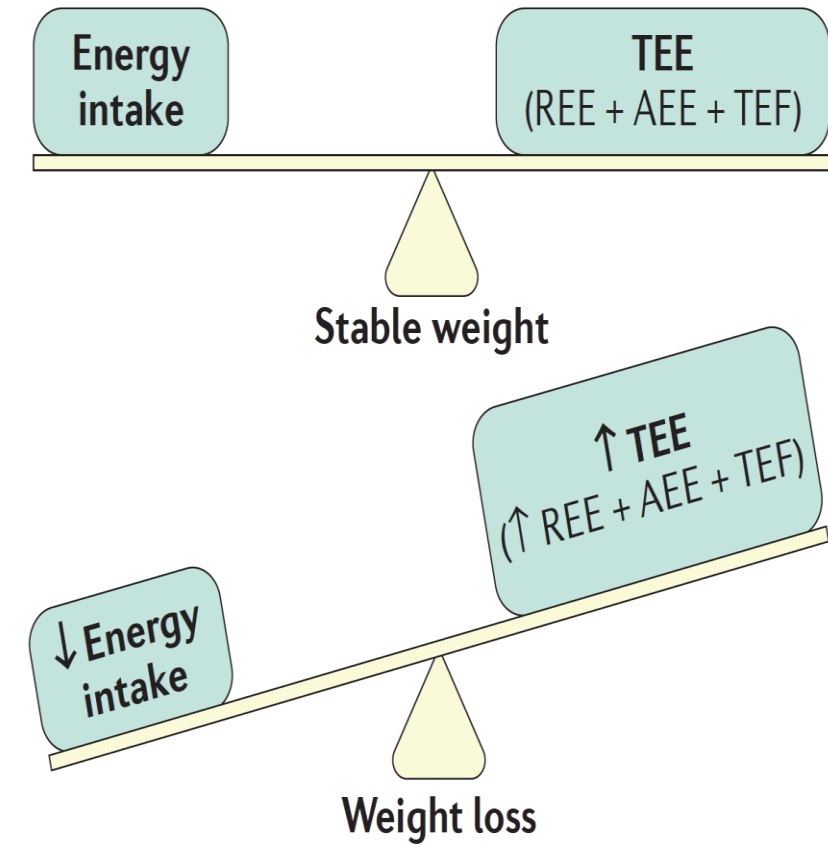
Cancer-related weight loss/malnutrition/cachexia



Mechanisms underlying cancer-related weight loss



Actors and targets in the pathophysiology of cancer cachexia



Negative energy balance is a driver of cancer-related malnutrition and cachexia

Baracos VE et al, 2018

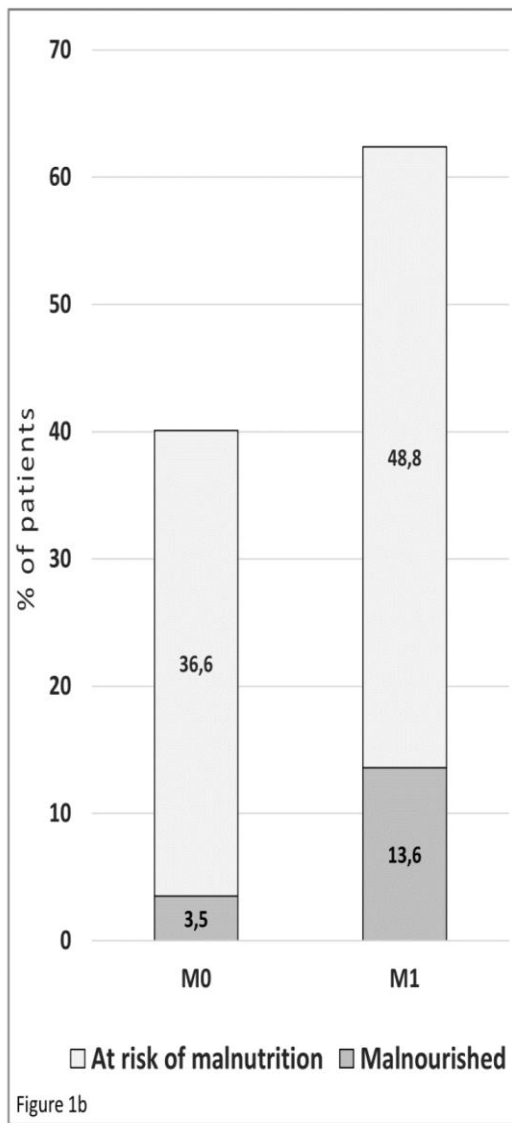
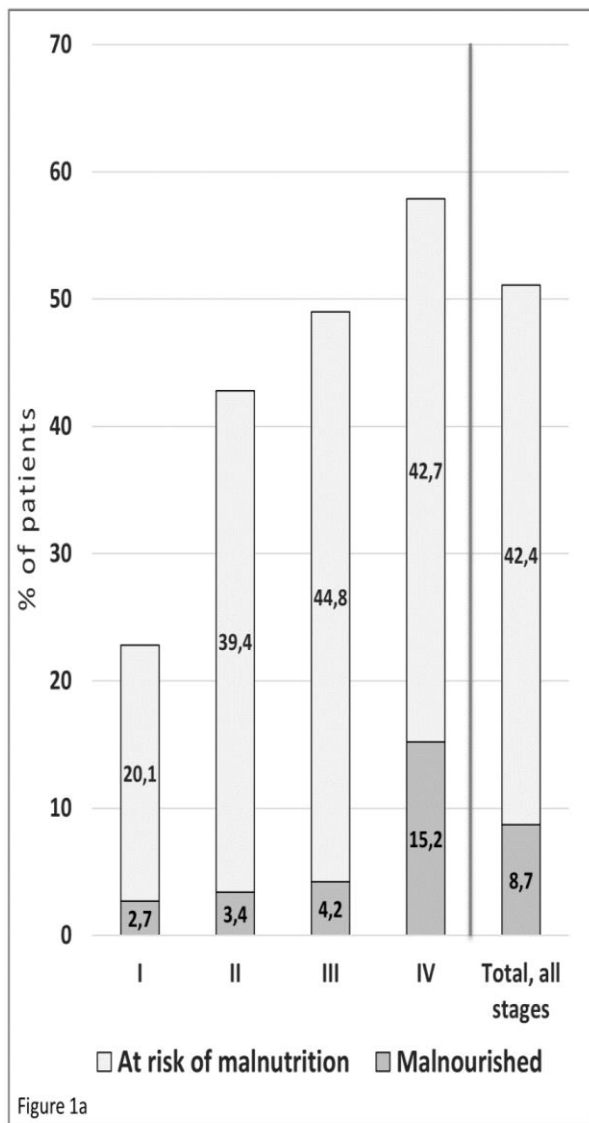
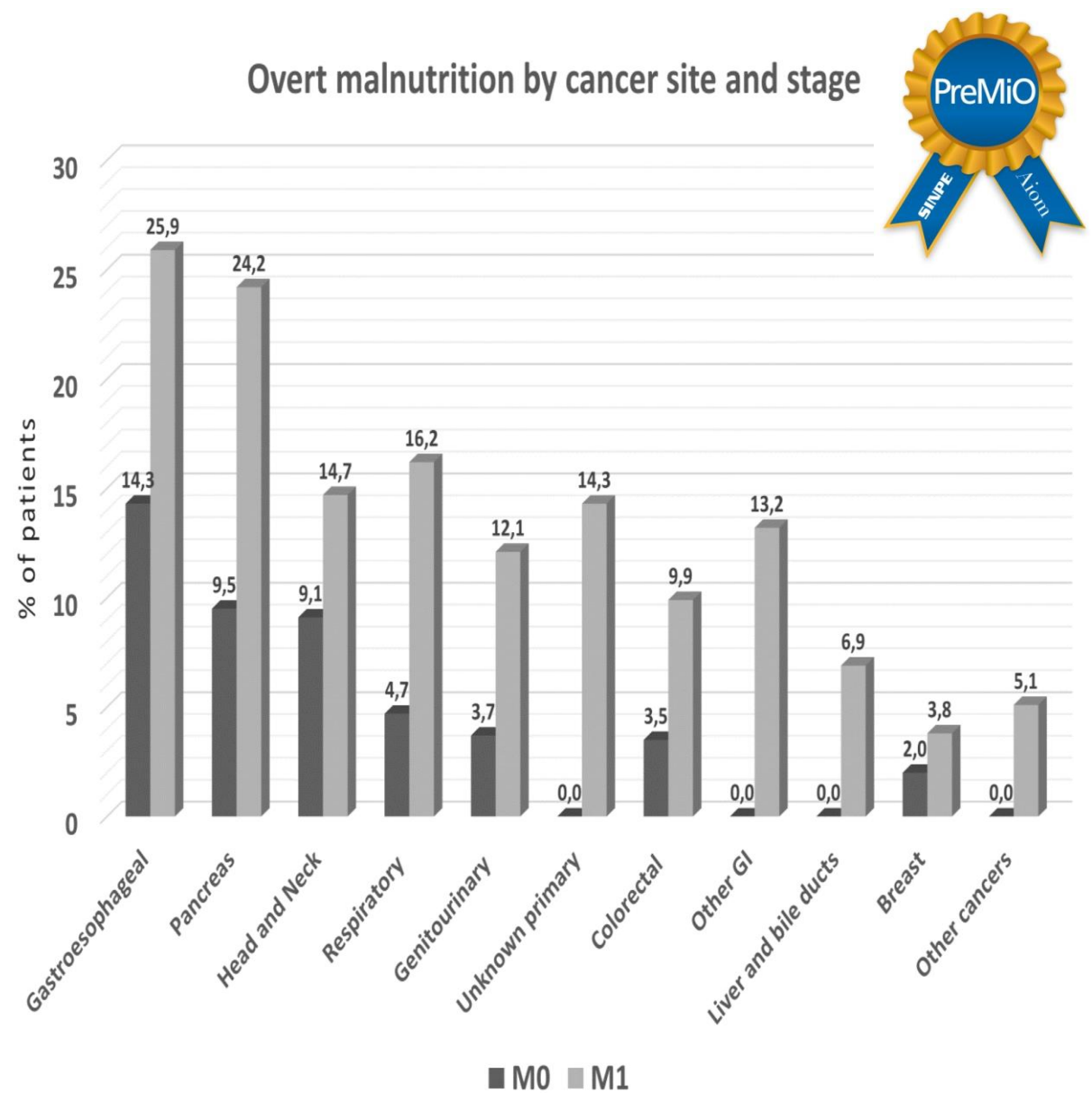


Figure 1. PreMiO patients with malnutrition or malnutrition risk using MNA scoring with results shown by tumor stage and for all tumors (Figure 1a) as well as classified in M0 and M1 groups (Figure 1b) (N = 1925) ($p < 0.001$ at ANOVA among cancer stage groups). Malnutrition was defined as MNA score < 17 , while risk of malnutrition was represented by a MNA scores of 17 to 23.5). M0 = stage I-III, M1 = stage IV



Prevalence of overt malnutrition by cancer site (% of patients with specified tumor type), with malnutrition defined as MNA score < 17 (N = 1925). M0 = stage I-III, M1 = stage IV ($p < 0.001$ at ANOVA among cancer site groups)



Prevalence of
cachexia by primary
tumor type in the
study population
(N = 1952).

M0 = stage I-III, M1 =
stage IV. ($p < 0.001$ at
ANOVA among
cancer site groups)

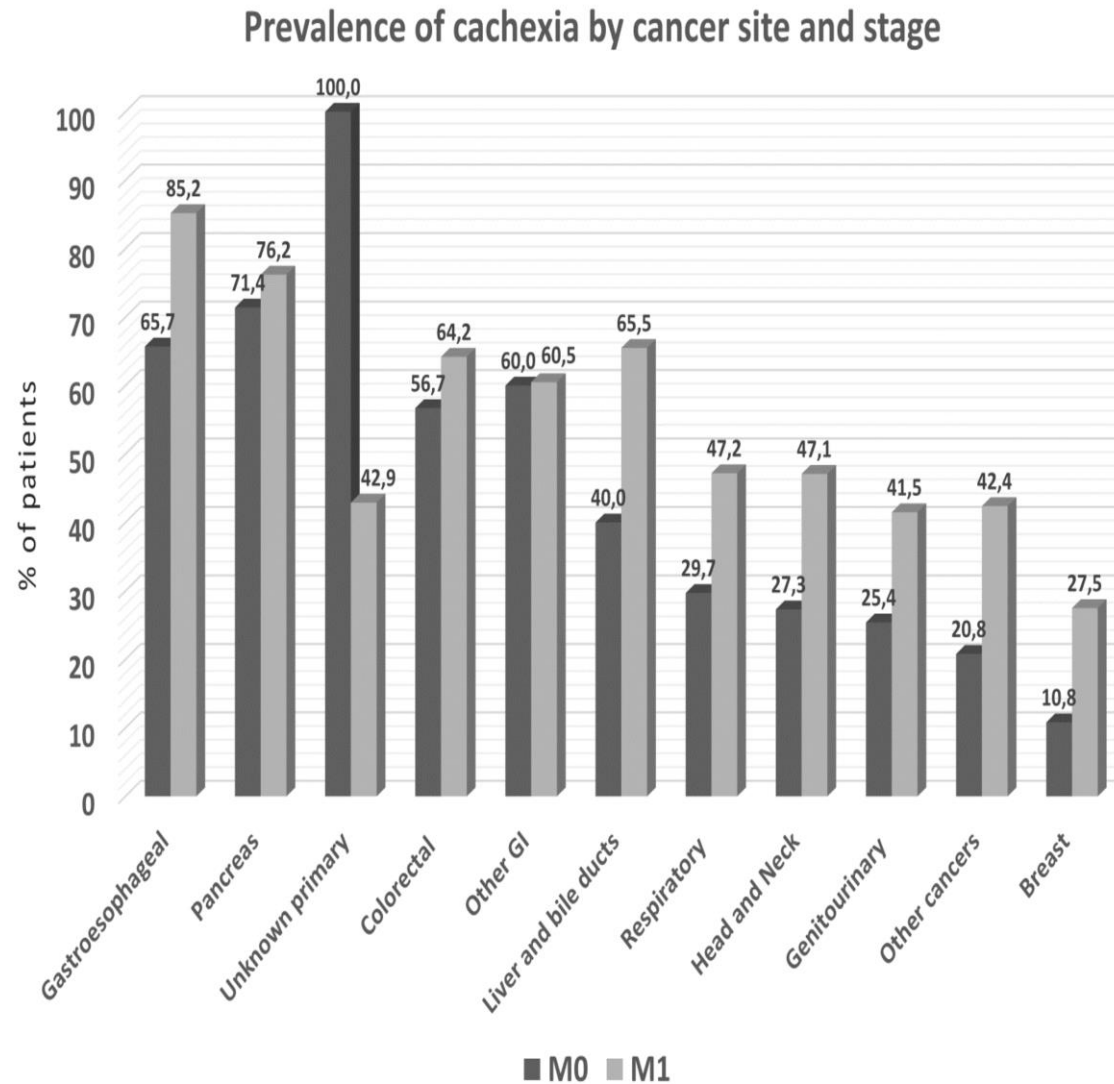


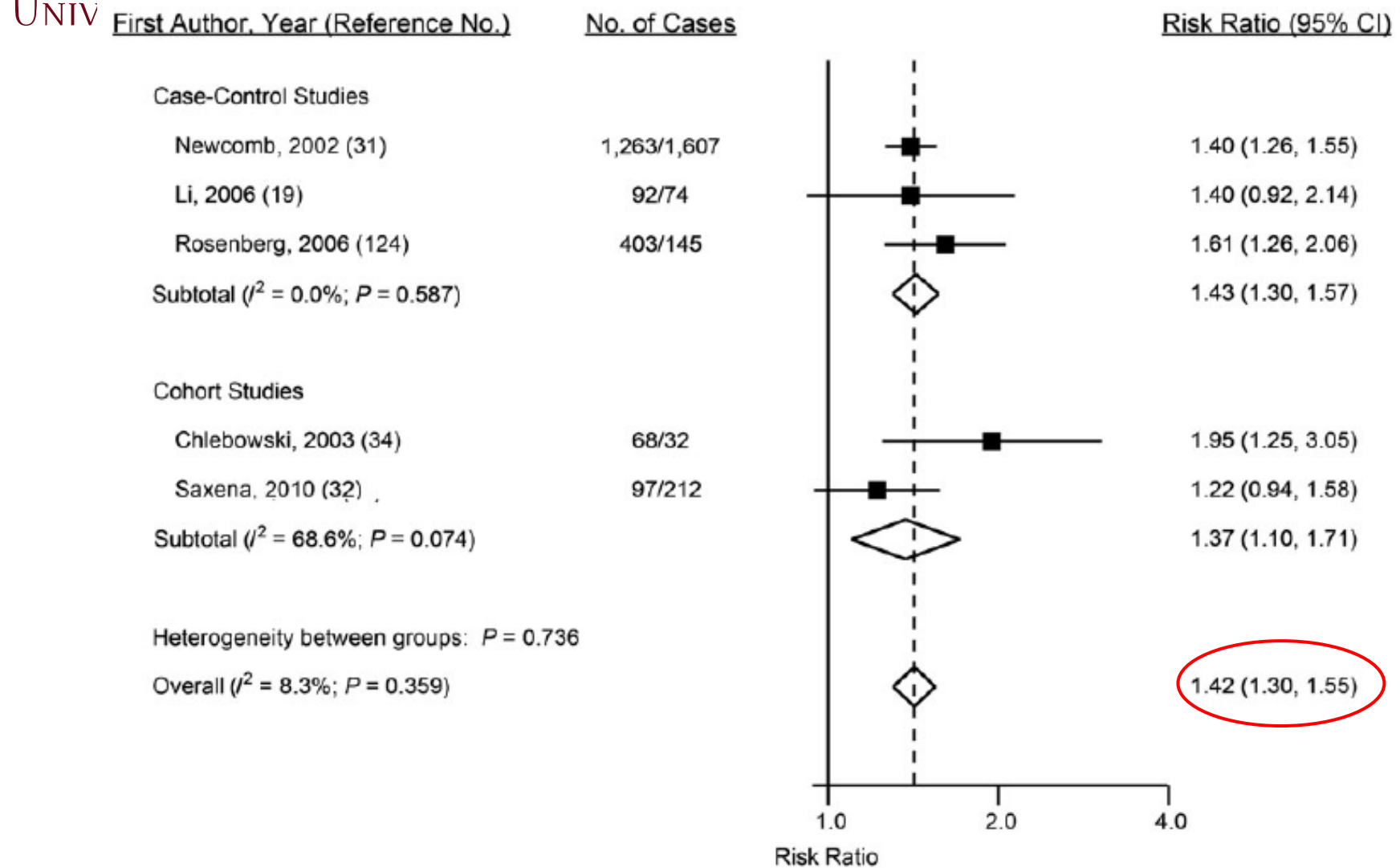


Table 2: Patient appetite scores by cancer site, based on FAACT (N=1949) and VAS scores (N=1857)

Cancer site	FAACT M0	FAACT M1	FAACT Total	VAS appetite M0	VAS appetite M1	VAS appetite total
Breast	33 ±5	29 ±5	32 ±5	73 ±20	69 ±19	73 ±20
Genitourinary tract	32 ±5	28 ±6	30 ±6	72 ±18	61 ±22	67 ±21
Colorectal	32 ±5	29 ±5	30 ±5	72 ±23	65 ±22	68 ±22
Lung	31 ±5	29 ±5	29 ±6	71 ±24	64 ±23	66 ±23
Other cancer ¹	33 ±6	29 ±6	32 ±6	78 ±22	69 ±21	75 ±22
Gastroesophageal	27 ±6	23 ±6	25 ±6	58 ±23	52 ±21	54 ±21
Pancreatic	28 ±4	24 ±7	25 ±6	62 ±22	48 ±27	53 ±26
Other GI	34 ±5	28 ±6	30 ±5	69 ±18	62 ±21	63 ±21
Liver/bile duct	33 ±2	26 ±5	28 ±5	82 ±8	62 ±21	65 ±20
Head and neck	33 ±6	30 ±5	31 ±5	75 ±23	64 ±19	68 ±22
Unknown primary site ²	26	25±6	28 ±6	20	45 ±17	55 ±24
ALL CANCERS	32 ±5	28 ±6	30 ±6	72 ±21	62 ±23	67 ±23



ASSOCIATION BETWEEN BODY MASS INDEX ≥ 30 kg/m² AND POSTMENOPAUSAL BREAST CANCER





STILI DI VITA NELLA PREVENZIONE DEL CARCINOMA MAMMARIO

TABLE 1. World Cancer Research Fund Prevention Guidelines for Breast Cancer and Carc

	Breast Cancer Risk Reduction
1. Be as lean as possible without becoming underweight	✓✓
2. Be physically active for at least 30 minutes every day	✓✓
3. Avoid sugary drinks and limit consumption of energy-dense foods	✓✓ (to achieve weight control)
4. Eat more vegetables, fruits	No effect
5. Eat more whole grains and legumes such as beans	✓✓
6. Limit red meats (i.e., beef, pork, and lamb) and avoid processed meats	Modest effect with processed meat ✓✓
7. Limit alcoholic drinks to 2 for men and 1 for women per day	✓✓
8. Limit consumption of salty foods and foods processed with salt	No effect
9. Do not use nutritional or vitamin supplements to reduce risk of disease	✓✓✓

Abbreviation: CVD, cardiovascular disease.
✓✓✓ = Supported by meta analyses of randomized trials or one or more randomized trials.
✓✓ = Association in three or more observational studies.
✓ = Association in one or two observational studies.



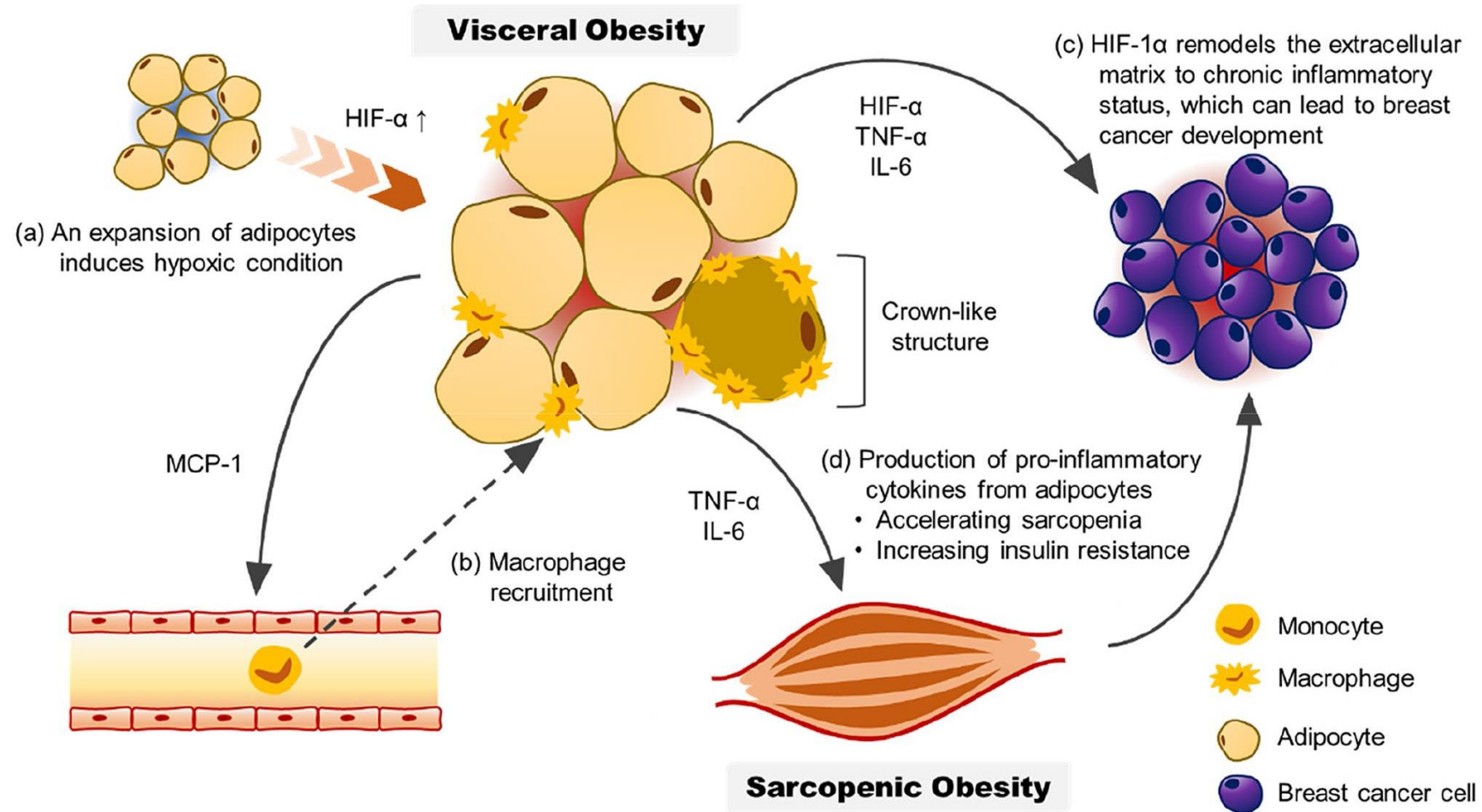
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WCRF/AICR recommendations





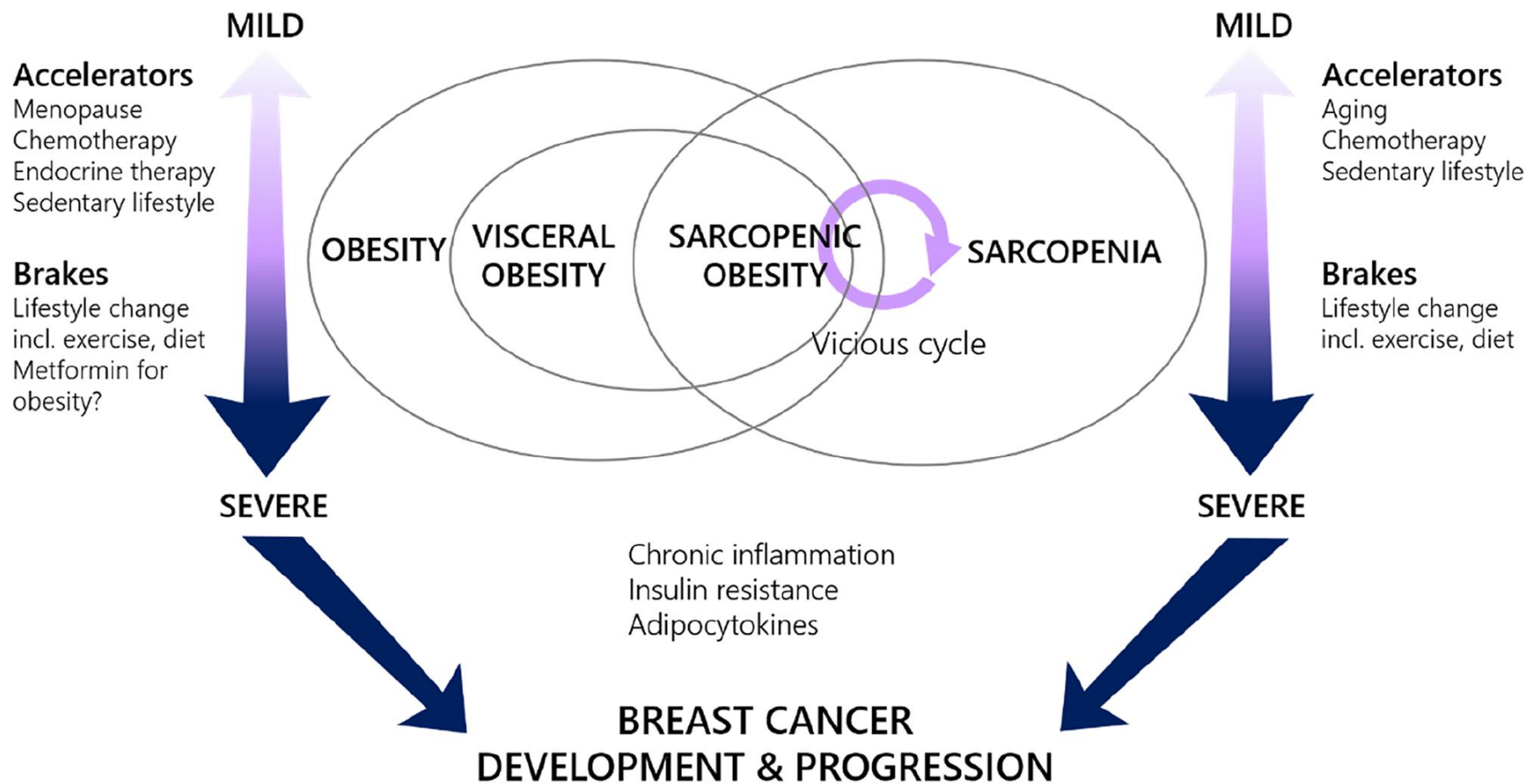
The physiologic roles of specific body composition phenotypes





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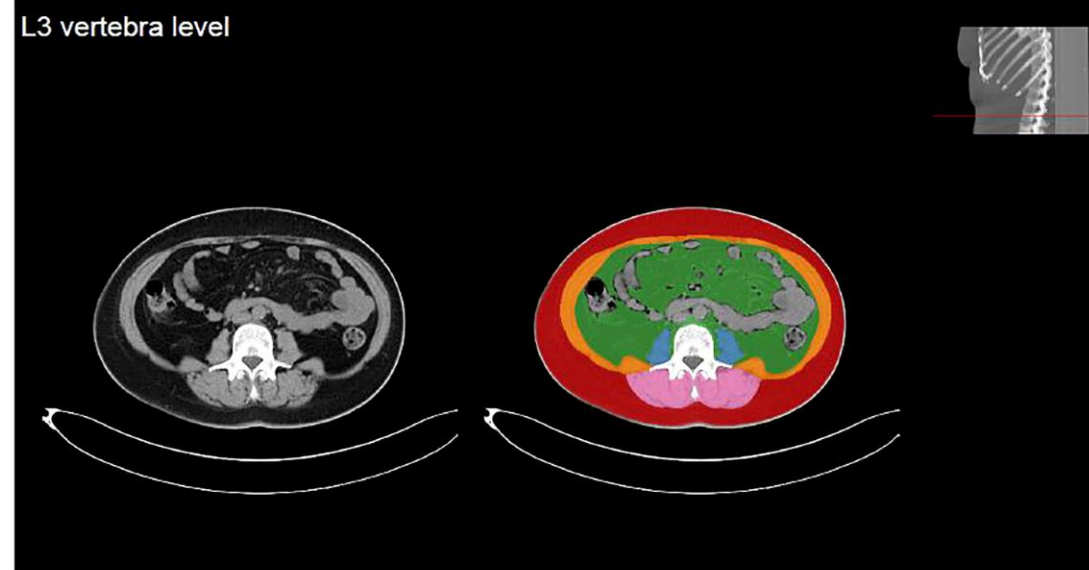
Proposed relationship between body composition phenotype and breast cancer





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A.I.- assisted body composition assessment by CT scan



Fat

Subcutaneous	Visceral
173.4 cm ² 31.0% -94.3 HU	163.0 cm ² 29.1% -92.2 HU

Muscle

Psoas	Abdominal	Long spine
13.1 cm ² 2.3% 33.2 HU	60.1 cm ² 10.7% 15.1 HU	42.5 cm ² 7.6% 22.3 HU

FIGURE 1

Screenshot generated by the software after performing an automatic segmentation at the level of the third lumbar vertebra and report of the measurements of SAT, VAT and SMA carried out. SAT values are indicated in red and VAT values in green. SMA is obtained by summing the areas of the psoas, the abdominal and the long spinal muscles indicated in blue, orange and pink respectively ($13.1 + 60.1 + 42.5 = 115.7 \text{ cm}^2$). SMI is then calculated as the ratio of SMA to the height squared of the patients.



A total of 2,948 Chinese female patients with breast cancer have been included in this retrospective study.



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Association of body composition with clinical outcome in Chinese women diagnosed with breast cancer

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Kaiwen Xi³ and Qiong Fang^{2*}

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Body composition and DFS in BC

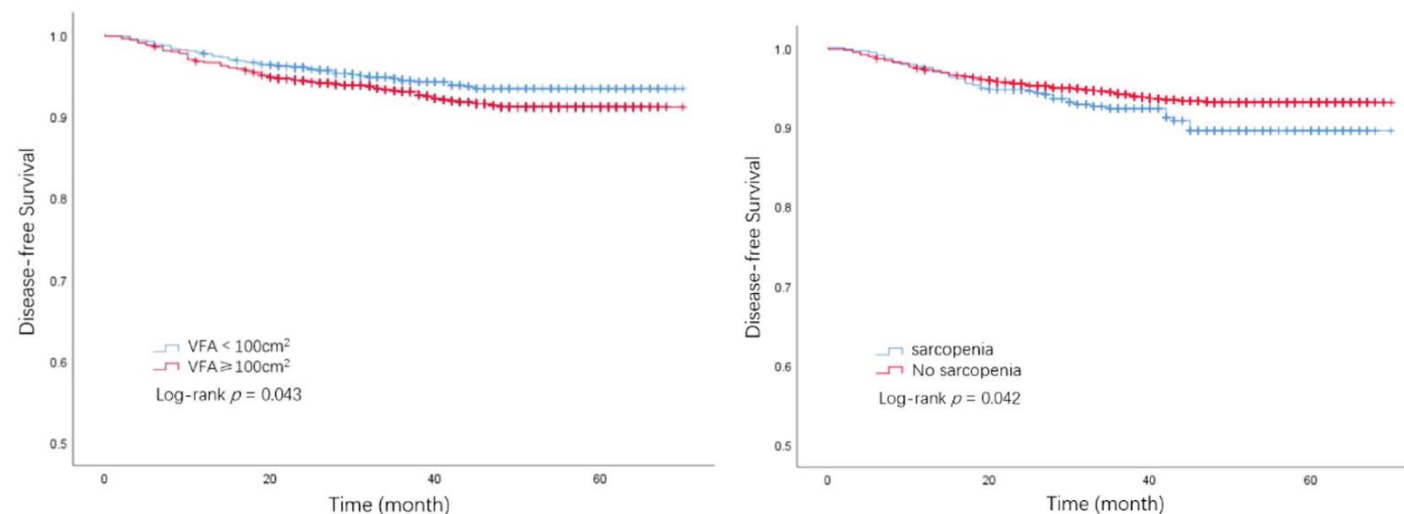


FIGURE 1

Kaplan–Meier plot of disease-free survival by VFA and ASMI. **(A)** VFA and disease-free survival. **(B)** ASMI and disease-free survival. VFA, visceral fat area; ASMI, appendicular skeletal muscle mass.



A total of 2,948 Chinese female patients with breast cancer have been included in this retrospective study.



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Body composition and OS in BC

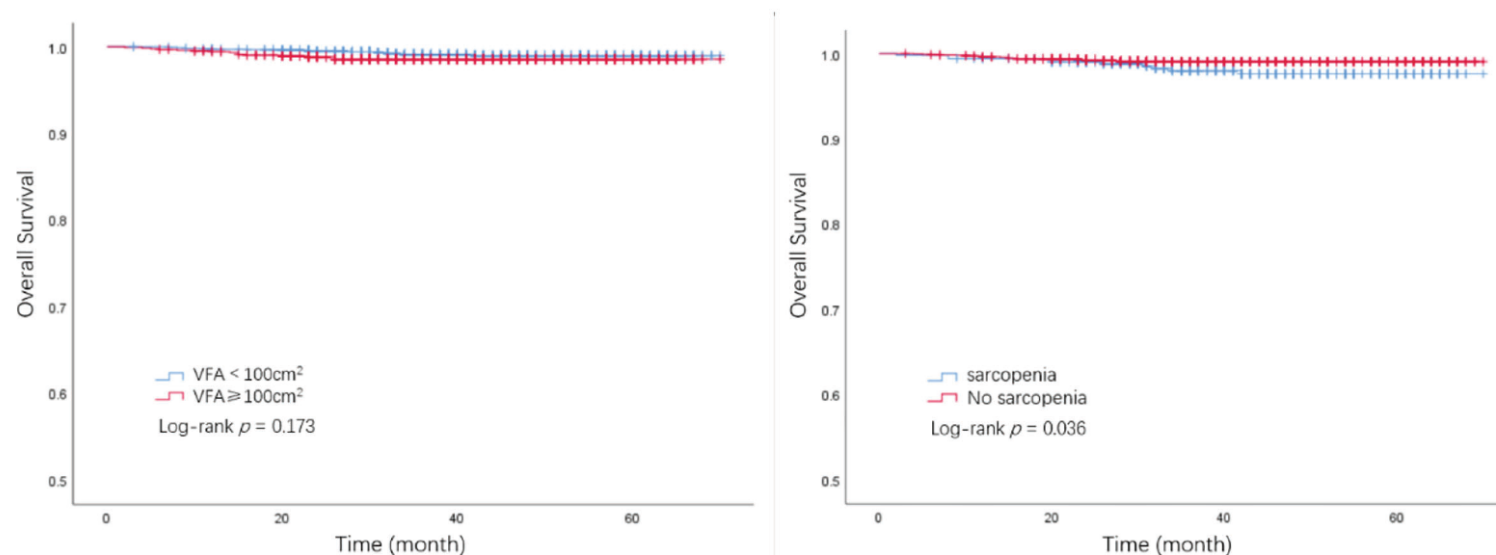


FIGURE 2

Kaplan–Meier plot of overall survival by VFA and ASMI. (A) VFA and disease-free survival. (B) ASMI and disease-free survival. VFA, visceral fat area; ASMI, appendicular skeletal muscle mass.



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frontiers | Frontiers in Oncology

TYPE Original Research
PUBLISHED 23 September 2022
DOI 10.3389/fonc.2022.987012



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SPECIALTY SECTION

This article was submitted to
Cancer Imaging and
Image-directed Interventions,
a section of the journal
Frontiers in Oncology

Do body composition parameters correlate with response to targeted therapy in ER+/HER2- metastatic breast cancer patients? Role of sarcopenia and obesity

Endi Kripa*, Veronica Rizzo, Francesca Galati, Giuliana Moffa, Federica Ciciarelli, Carlo Catalano and Federica Pediconi

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Association of Low-Fat Dietary Pattern With Breast Cancer Overall Survival

A Secondary Analysis of the Women's Health Initiative Randomized Clinical Trial

Rowan T. Chlebowski, MD, PhD; Aaron K. Aragaki, MS; Garnet L. Anderson, PhD; Michael S. Simon, MD, MPH; JoAnn E. Manson, MD, DrPH; Marian L. Neuhouser, PhD; Kathy Pan, MD; Marcia L. Stefanick, PhD; Thomas E. Rohan, MBBS, PhD; Dorothy Lane, PhD; Lihong Qi, PhD; Linda Snetselaar, PhD; Ross L. Prentice, PhD

This was a secondary analysis of the **Women's Health Initiative** randomized clinical trial that was conducted at 40 US clinical centers

Key Points

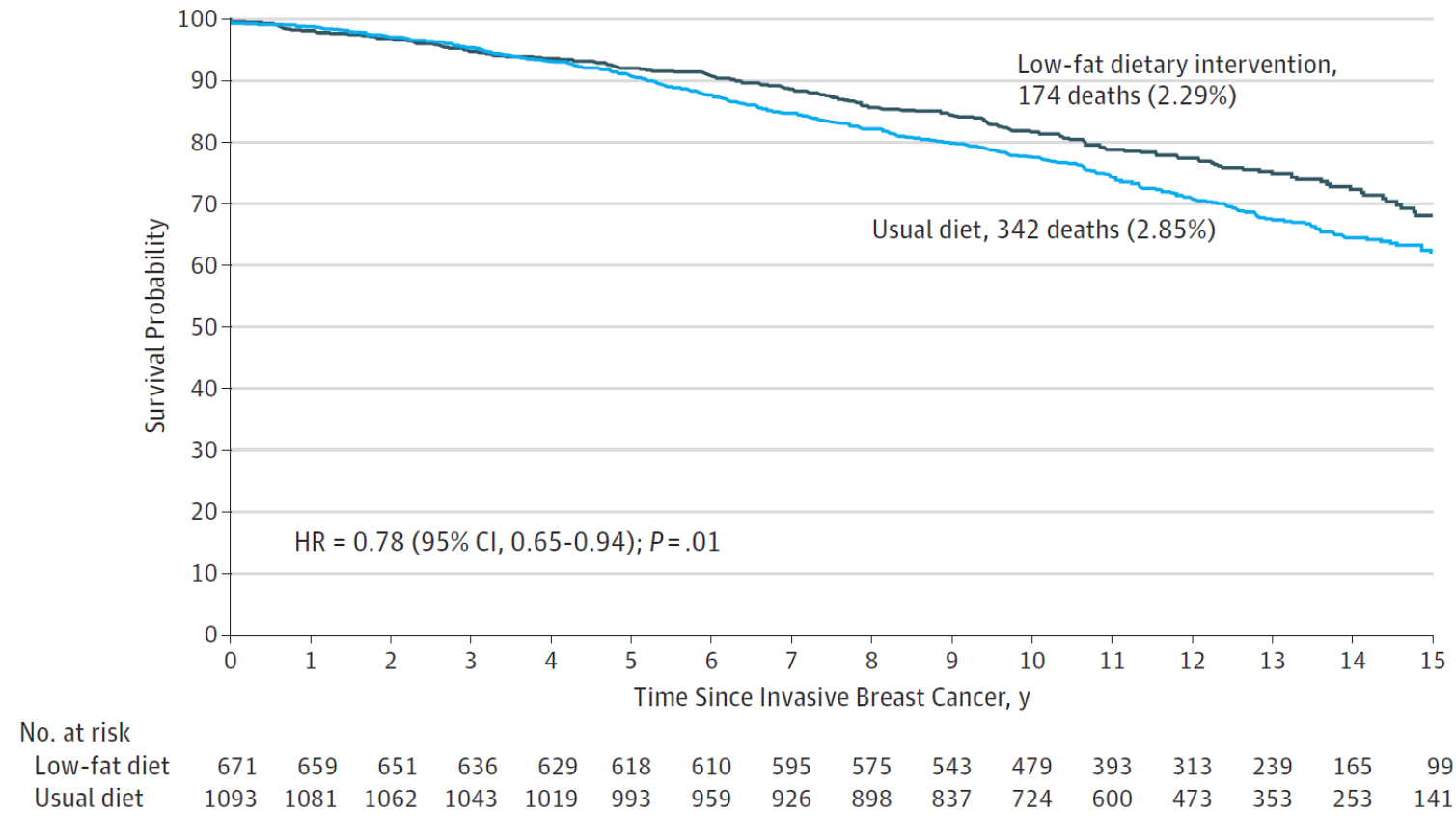
Question Is implementation of a low-fat eating pattern associated with breast cancer outcome?

Finding In this secondary analysis of the Women's Health Initiative randomized clinical trial that included 1764 postmenopausal women who received a diagnosis of breast cancer during the dietary intervention period, those randomized to a low-fat dietary pattern had increased breast cancer overall survival.

Meaning A dietary change may be able to influence breast cancer outcome.



Figure 2. Dietary Modification Association With Breast Cancer Overall Survival



Kaplan-Meier estimates for breast cancer overall survival (survival from diagnosis with death from any cause) among the 1764 breast cancer cases diagnosed during the dietary intervention period, measured from cancer diagnosis and observed through September 2013. Summary statistics are from a Cox model stratified by age at diagnosis, randomization status in the hormone therapy trials, and study period (intervention period, postintervention period extension 1, or postintervention period extension 2; time dependent). The P value corresponds to a 2-sided score (log-rank) test. Percentages are annualized. HR indicates hazard ratio.



CLINICAL TRIAL

Dietary intervention among breast cancer survivors increased adherence to a Mediterranean-style, anti-inflammatory dietary pattern: the Rx for Better Breast Health Randomized Controlled Trial


Krystle E. Zuniga¹  · Dorothy Long Parma² · Edgar Muñoz² · Mackenzie Spaniol² · Michael Wargovich³ · Amelie G. Ramirez²

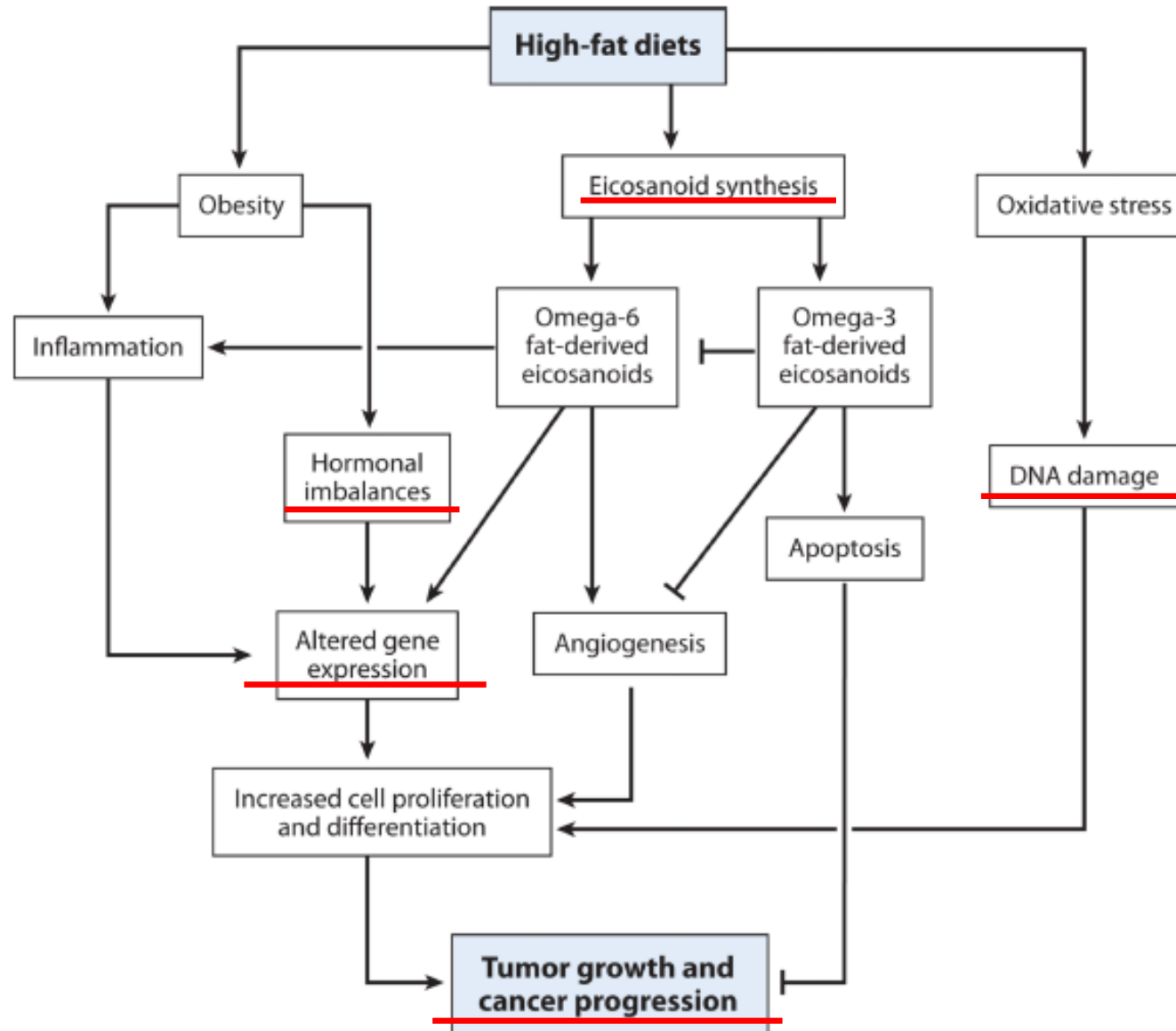
Table 5 Dietary changes between baseline and 6 months		Baseline	6 months	Change	<i>P</i> value ^a
Calories					0.045
Intervention		1779.3 (92.2)	1578.3 (85.6)	– 195.5 (83.0)	
Control		1765.8 (85.0)	1805.3 (79.0)	+ 34.8 (76.6)	

Table 4 Comparison of mediterranean diet and spices and herbs scores between intervention and control

	Baseline	6 months	Change	<i>P</i> value ^a
Mediterranean diet score				<0.001
Intervention	7.1 (0.3)	8.7 (0.3)	+ 1.6 (0.2)	
Control	7.4 (0.3)	7.6 (0.3)	+0.2 (0.2)	
Spices and herbs score				<0.001
Intervention	1.3 (0.2)	3.2 (0.3)	+ 1.9 (0.3)	
Control	1.2 (0.1)	1.6 (0.3)	+0.4 (0.2)	



ASSOCIATION BETWEEN DIETARY FATS AND BREAST CANCER





ω -3 and ω -6 Polyunsaturated Fatty Acid Intakes and the Risk of Breast Cancer in Mexican Women: Impact of Obesity Status

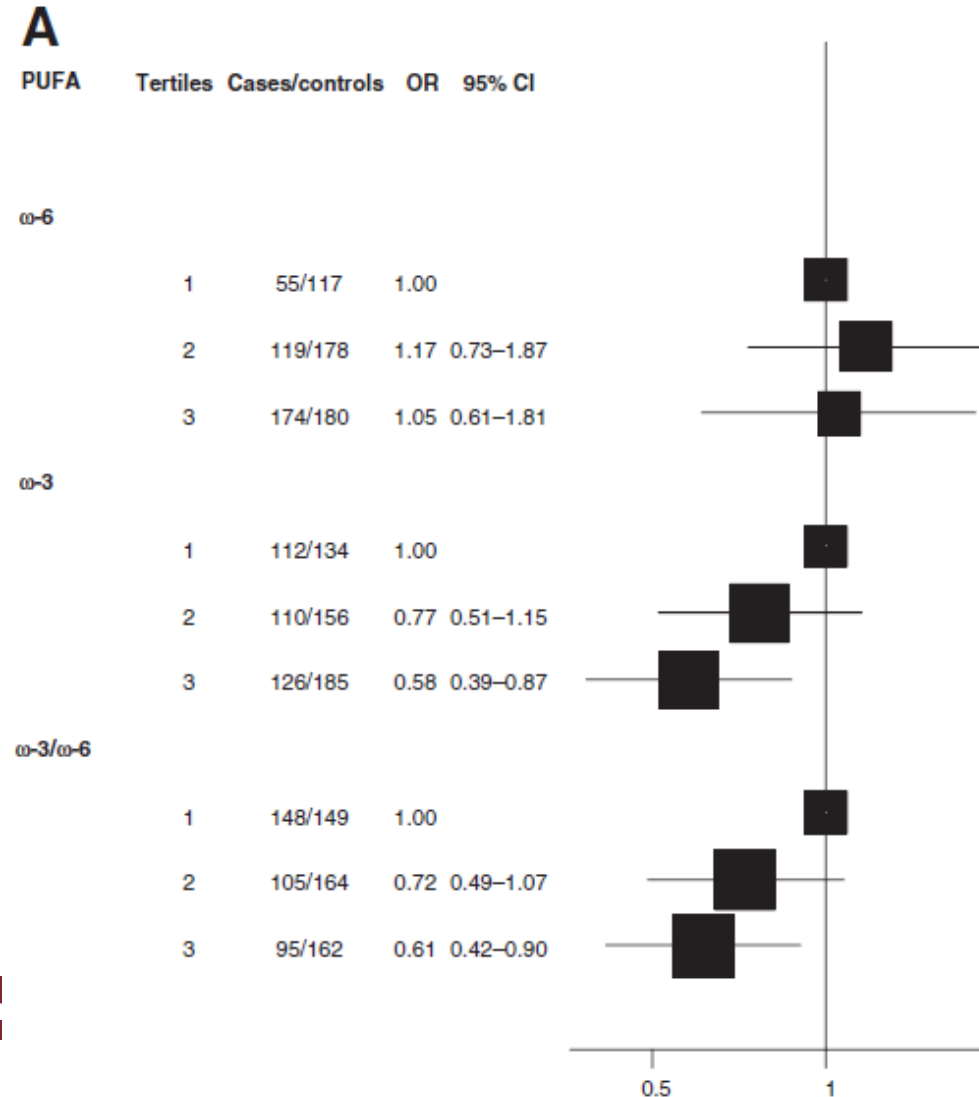
Véronique Chajès¹, Gabriela Torres-Mejía², Carine Biessy¹, Carolina Ortega-Olvera², Angélica Angeles-Llerenas², Pietro Ferrari¹, Eduardo Lazcano-Ponce², and Isabelle Romieu¹

Cancer Epidemiol Biomarkers Prev; 21(2); 319–26. ©2011

Effect of omega-3 PUFAs on breast cancer risk according to BMI tertiles

Associations between PUFA intake and breast cancer risk stratified by BMI:

- A decreased risk of breast cancer was significantly associated with increasing omega-3 PUFA intake in obese women
- but not in normal weight women



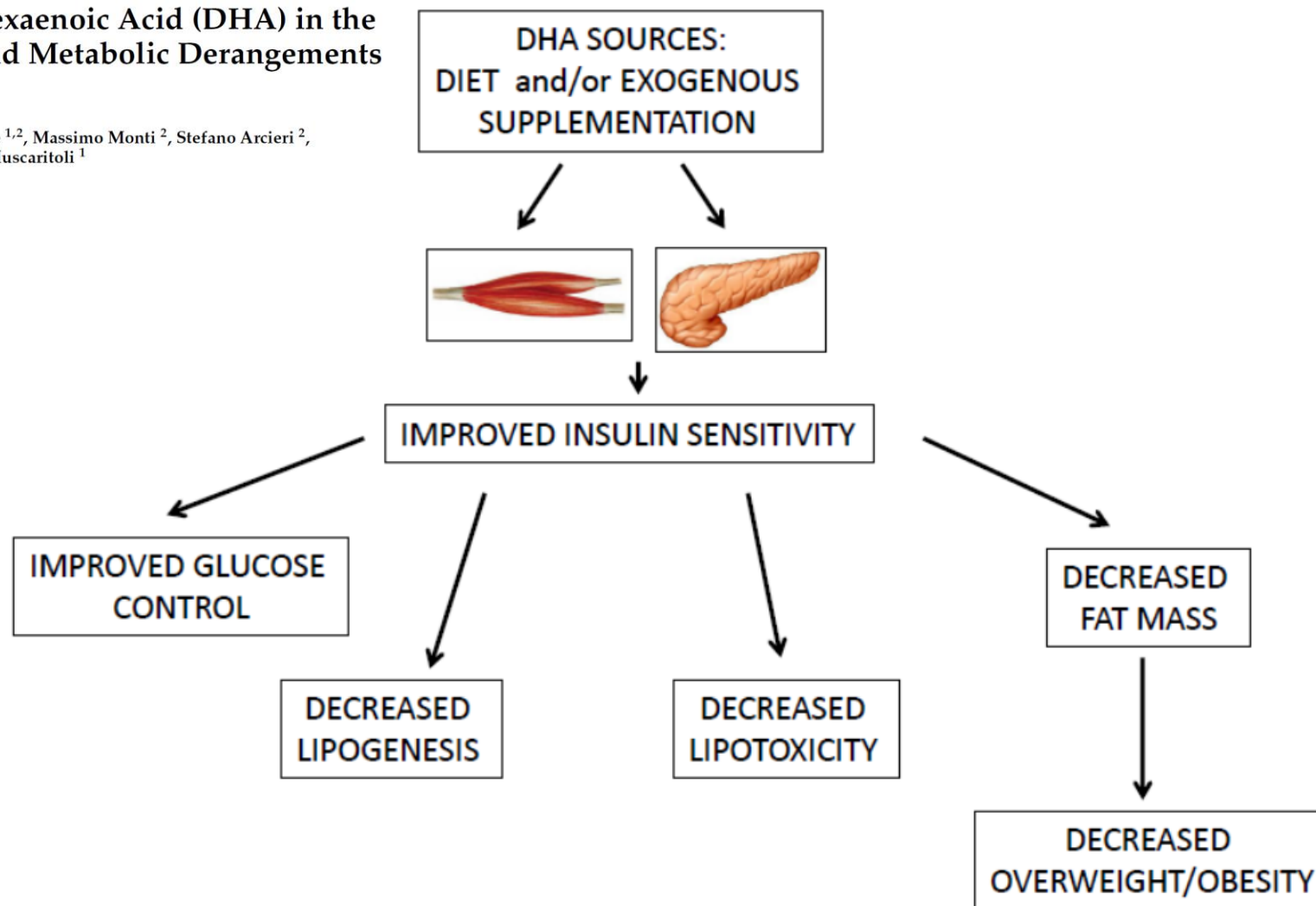


Review

The Role of Docosahexaenoic Acid (DHA) in the Control of Obesity and Metabolic Derangements in Breast Cancer

Alessio Molfino ^{1,*}, Maria Ida Amabile ^{1,2}, Massimo Monti ², Stefano Arcieri ²,
Filippo Rossi Fanelli ¹ and Maurizio Muscaritoli ¹

DHA AMELIORATES OBESITY



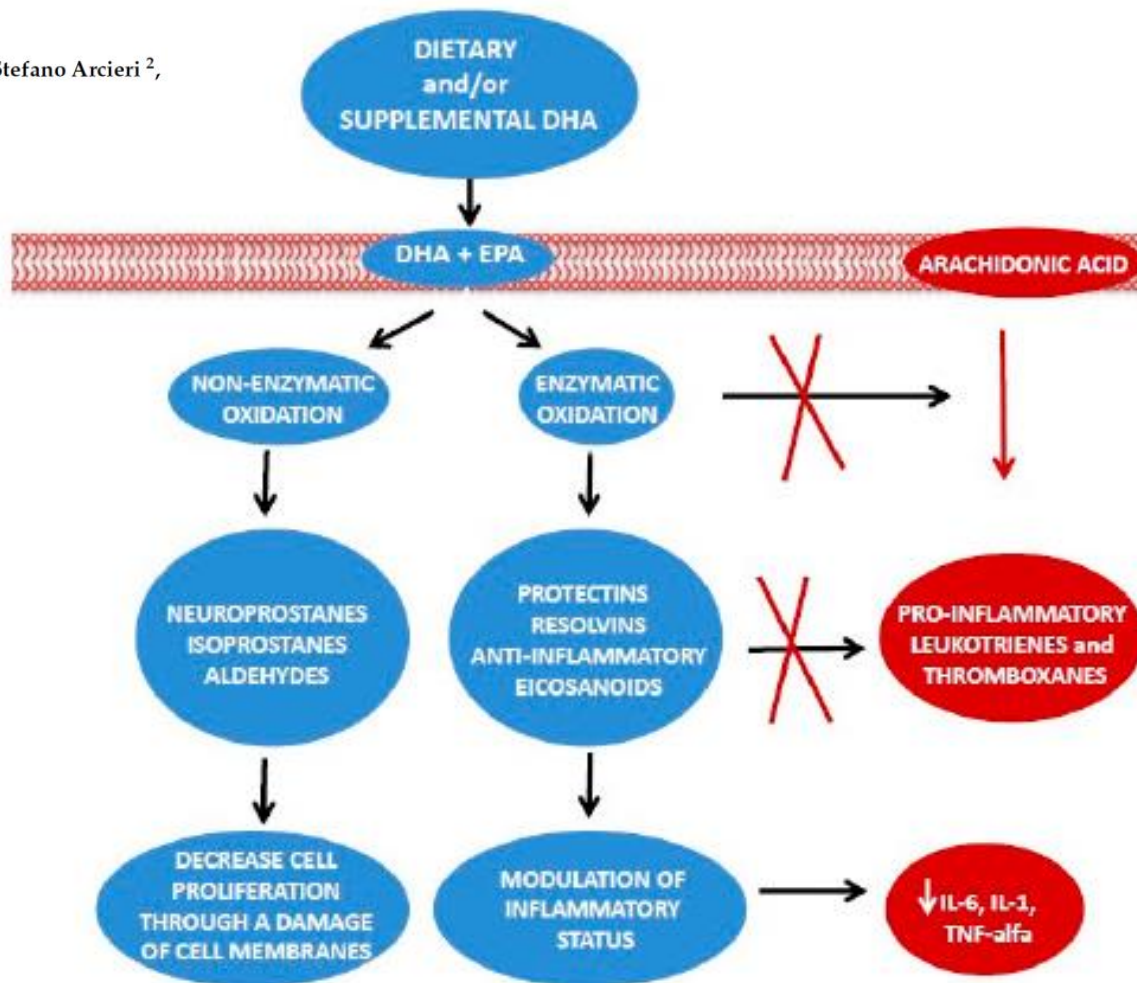


Review

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ENZYMATIC WAY AND NON-ENZYMATIC WAY FOR ω -3 PUFAS





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Valutazione degli effetti della supplementazione di acido docosaesaenoico (DHA) sul profilo metabolico lipidico in donne affette da neoplasia mammaria e sottoposte a chirurgia mammaria



ORIGINAL RESEARCH
published: 28 July 2017
doi: 10.3389/fphys.2017.00549



Effect of Oral Docosahexaenoic Acid (DHA) Supplementation on DHA Levels and Omega-3 Index in Red Blood Cell Membranes of Breast Cancer Patients

Alessio Molfino¹, Maria I. Amabile^{1,2}, Sara Mazzucco³, Gianni Biolo³, Alessio Farcomeni⁴, Cesarina Ramaccini¹, Simonetta Antonaroli⁵, Massimo Monti² and Maurizio Muscaritoli^{1*}

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DHA supplementation was associated with **increased DHA levels** and **omega-3 index** in RBC membranes of BC cancer patients, independent of the type of BC presentation, and in controls.

BRCA1/2 mutation, as well as **low seafood consuming habits** in both BC patients and healthy controls, seem to be **associated with greater ability of DHA incorporation**.

TABLE 3 | Multivariate regression models to predict variation of DHA levels in RBC membranes between baseline (T0) and after DHA supplementation (T1).

Clinical characteristics	Beta coefficient (95% CI)	P
TYPE OF BC PRESENTATION		
S group vs. C group	0.12 (−0.21, 0.44)	0.47
F group vs. C group	0.21 (−0.13, 0.55)	0.21
M group vs. C group	0.30 (0.05, 0.55)	0.02
DIETARY SEAFOOD HABITS		
Good vs. Low seafood consumer	−0.49 (−0.68, −0.30)	<0.001

DHA, docosahexaenoic acid; RBC, red blood cell; BC, breast cancer; S group, sporadic group; F group, familial group; M group, mutated group; C group, controls.

TABLE 2 | DHA levels, EPA levels, and omega-3 index in RBC membranes in the four groups of participants at baseline (T0) and after DHA supplementation (T1).

All participants (N = 43)	T0, Mean ± SD	T1, Mean ± SD	P
S GROUP (N = 10)			
DHA	6.14 ± 1.21	7.54 ± 0.97	0.002
EPA	0.53 ± 0.23	0.68 ± 0.22	0.014
Omega-3 Index	6.67 ± 1.33	8.21 ± 1.08	0.002
F GROUP (N = 12)			
DHA	6.09 ± 1.11	7.35 ± 1.12	0.003
EPA	0.62 ± 0.27	0.77 ± 0.33	0.004
Omega-3 Index	6.71 ± 1.32	8.12 ± 1.35	<0.001
M GROUP (N = 11)			
DHA	5.86 ± 1.51	7.4 ± 1.32	<0.001
EPA	0.60 ± 0.34	0.7 ± 0.27	0.018
Omega-3 Index	6.43 ± 1.75	8.1 ± 1.53	<0.001
C GROUP (N = 10)			
DHA	6.1 ± 0.88	7.23 ± 0.7	0.006
EPA	0.52 ± 0.11	0.6 ± 0.1	0.004
Omega-3 Index	6.62 ± 0.9	7.0 ± 0.74	0.002

Effect of Oral Docosahexaenoic Acid (DHA) Supplementation on DHA Levels and Omega-3 Index in Red Blood Cell Membranes of Breast Cancer Patients

Alessio Molfino¹, Maria I. Amabile^{1,2}, Sara Mazzucco³, Gianni Biolo³, Alessio Farcomeni⁴, Cesarina Ramaccini¹, Simonetta Antonaroli⁵, Massimo Monti² and Maurizio Muscaritoli^{1*}



A role for omega-3 fatty acid metabolites?

- Experimental studies have documented that specific epoxydocosapentaenoic acids (EDPs) - in particular 19,20-EDPs - are potent mediators in suppressing inflammation and inhibiting angiogenesis, endothelial cell migration, endothelial cell proliferation, and the growth and metastasis of human breast and prostate cancer



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Research Article

DHA Oral Supplementation Modulates Serum Epoxydocosapentaenoic Acid (EDP) Levels in Breast Cancer Patients

**Alessio Molfino,¹ Maria Ida Amabile,² Luana Lionetto,³ Alessandra Spagnoli,⁴
Cesarina Ramaccini,¹ Alessandro De Luca,² Maurizio Simmaco,³ Massimo Monti,²
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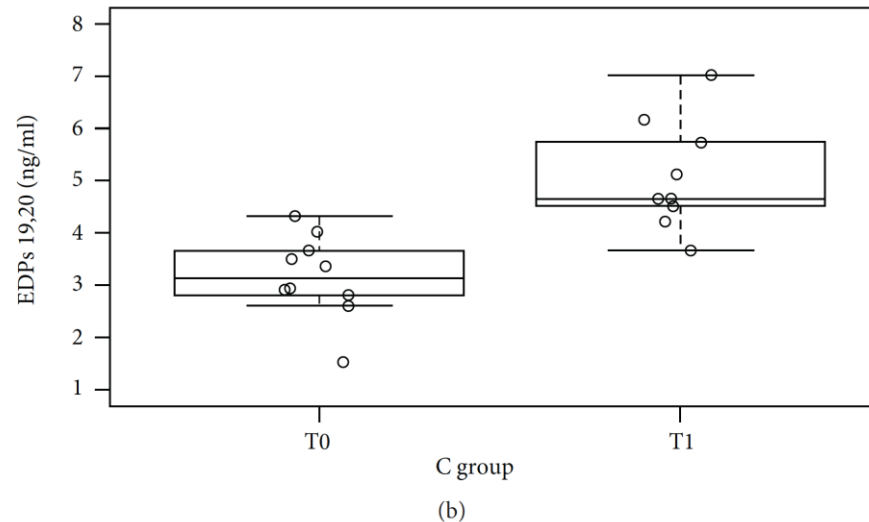
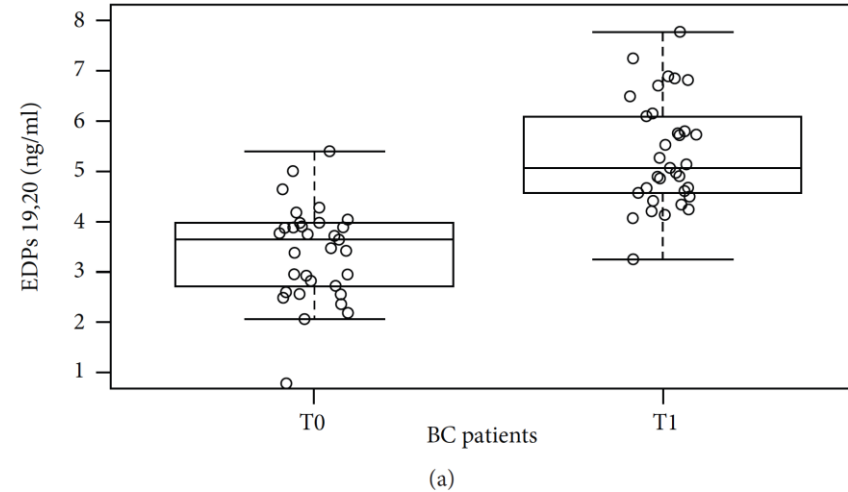
Received 13 November 2018; Revised 11 January 2019; Accepted 23 January 2019

Oxidative Medicine and Cellular Longevity

Department of Translational and Precision Medicine



Effect of oral DHA on 19,20EDP in BC patients



Oxidative Medicine and Cellular Longevity

DHA oral supplementation was associated with increased 19,20-EDP serum levels in BC patients, independent of the type of BC presentation.

19,20-EDP levels were higher at T1 with respect to T0 ($P < 0.001$)



Conclusioni

- In Italia, il tributo pagato ogni anno al tumore mammario femminile è ancora molto elevato, anche se la sopravvivenza a 5 anni è in miglioramento
- Abitudini alimentari e stili di vita scorretti, sovrappeso, obesità rappresentano fattori di aumentato rischio di ammalarsi di tumore mammario
- Quantità e qualità dei grassi alimentari hanno rilevanza sia nella prevenzione che nel trattamento del tumore mammario
- La composizione corporea influenza la sopravvivenza e la risposta ai trattamenti nelle donne con tumore mammario e deve entrare a fare parte della valutazione multidimensionale e multidisciplinare
- Studi prospettici e su ampie casistiche sono necessari per confermare le associazioni tra massa magra, massa grassa e prognosi di malattia