Supplementary material

Supplementary Table S1. Scoring method and mean scores (95%CI) of components of the modified Alternate Healthy Eating Index^{1,2} of adult childhood cancer survivors (CCSs) who filled in the food frequency questionnaire by sex

Components	Criteria for minimum score (0)	Criteria for maximum score (10)	Score in CCSs ¹³ mean (95%CI)		
			male (<i>n</i> =401)	female (<i>n</i> =401)	
Vegetables, excluding potatoes (servings/day) ³	0	≥ 5	4.6 (4.4; 4.9)	5.5 (5.3; 5.7)	
Fruit, excluding juice (servings/day) ⁴	0	≥ 4	2.9 (2.6; 3.2)	3.9 (3.6; 4.2)	
Whole grains (g/day) ⁵					
Males	0	≥ 90	3.3 (3.0; 3.7)		
Females	0	≥ 75	,	4.6 (4.3; 5.0)	
Sugar-sweetened beverage and fruit juice (servings/day) ⁶	≥ 1	0	3.8 (3.5; 4.2)	5.0 (4.6; 5.3)	
Nuts, seeds, legumes, and tofu (servings/day) ⁷	0	≥ 1	1.9 (1.6; 2.2)	2.5 (2.2; 2.8)	
Red and processed meat (servings/day)8	≥ 1.5	0	3.5 (3.2; 3.8)	5.7 (5.4; 6.0)	
Trans fat (% of total energy intake)9	≥ 4	≤ 0.5	9.4 (9.4; 9.4)	9.4 (9.3; 9.4)	
Fish, excluding processed products (g/day)	0	32.4	5.2 (4.8; 5.6)	5.0 (4.6; 5.3)	
PUFA (% of total energy intake) ¹⁰	≤2	≥10	4.8 (4.6; 5.1)	5.1 (4.8; 5.3)	
Sodium ¹¹ , mg/day	Highest decile	Lowest decile	5.3 (5.0; 5.6)	5.3 (5.0; 5.6)	
Alcohol (drinks/day) ¹²	•		,	, ,	
Males	≥ 3.5	0.5 - 2.0	5.4 (5.0; 5.7)		
Females	≥ 2.5	0.5 - 1.5	,	4.0 (3.7; 4.3)	
Total	0	100	50.3 (49.1; 51.5)	` '	

CCSs, childhood cancer survivors; Cl, confidence interval, ICCC-3, International Childhood Cancer Classification, 3rd edition; PUFA, polyunsaturated fatty acids

¹ Adapted from Chiuve et al. J Nutr 2012 142(6):1009-18.

² Intermediate food intake was scored proportionately between the minimum score 0 and the maximum score 10.

³ One serving was equal to 118.3 g of raw or cooked vegetables or 250 g of vegetable soup. Dried vegetables were not included.

⁴One serving was equal to 118.3 g of raw or cooked fruit. Dried fruit was not included.

⁵Whole grain products like whole grain bread and muesli were included.

⁶One serving was equal to 226.8 g.

⁷One serving was equal to 28.4 g.

⁸One serving was equal to 113.4 g of red meat or 42.5 g of processed meat.

⁹ Consumption was estimated based on the assumption that each food contains a maximum of 2 g of trans fat per 100 g of total fat, as defined in Swiss regulation.

¹⁰ The highest score was given to individuals with ≥10% of total energy intake from PUFA.

¹¹ Values in lowest decile were ≤1084 mg/d of sodium in females and ≤1168 mg/d in males and in highest decile were ≥2802 mg/d in females and ≥2969 mg/d in males.

¹² One drink was 113.4 g of wine, 340.2 g of beer or 42.5 g of liquor. A score of 2.5 was given to nondrinkers.

¹³ Adjusted for age at survey and ICCC-3 diagnosis.

Supplementary Table S2. Sociodemographic and lifestyle characteristics of adult childhood cancer survivors by participants and nonparticipants to the SCCSS-Nutrition study

		FFQ		Morning fasting spot urine				
	Participants (n = 802)	Non- participants ¹ (n = 797)	p-value ²	Participants (n = 111)	Non- participants ³ (n = 110)	p-value ²		
Characteristics n (%)								
Sex								
Females	401 (50)	354 (44)	0.025	62 (56)	57 (52)	0.547		
Males	401 (50)	443 (56)		49 (44)	53 (48)			
Age at follow-up survey ⁴ ,	, ,	, ,		, ,	` '			
median (IQR)	34.6 (28.8; 41.1)	32.0 (27.1; 38.7)	0.520	39.1 (30.7; 45.2)	33.4 (26.9; 40.4)	0.290		
≤30 y ` ´	248 (31)	328 (41) ^d	<0.001	26 (23)	44 (40)	0.010		
31-39 y	320 (40)	305 (38)		37 (33)	37 (34)			
≥40 y	234 (29)	164 (21)		48 (43)	29 (26)			
Country of birth	,	,		, ,	` ,			
Switzerland	763 (95)	737 (92)	0.027	101 (91)	98 (89)	0.637		
Other	39 (5)	60 (8)	0.02.	10 (9)	12 (11)			
Language region within	33 (3)	33 (3)		(0)	(,			
Switzerland								
German speaking	570 (71)	581 (73)	0.548	-	-	NA		
French speaking	214 (27)	195 (24)		111 (100)	110 (100)			
Italian speaking	18 (2)	21 (3)		-	-			
Education (highest degree) ⁵	(-)	_: (-)						
Lower than university	686 (86)	699 (88)	0.203	69 (62)	71 (65)	0.713		
University	116 (14)	98 (12)	0.203	42 (38)	39 (35)	0.7 13		
Living situation ⁵	110 (14)	30 (12)		72 (00)	00 (00)			
Alone	131 (16)	129 (16)	0.936	19 (17)	24 (22)	0.377		
With others	671 (84)	668 (84)	0.550	92 (83)	86 (78)	0.577		
Physical activity ^{5,6}	07 1 (04)	000 (04)		32 (00)	00 (10)			
Inactive	157 (20)	204 (26)	0.004	32 (29)	32 (29)	0.966		
Active	645 (80)	593 (74)	0.004	79 (72)	78 (71)	0.500		
Smoking status ⁵	0 10 (00)	000 (7-1)		70 (72)	70 (71)			
Current	151 (19)	207 (26)	0.002	25 (23)	21 (19)	0.261		
Former	97 (12)	79 (10)	0.002	15 (14)	24 (22)	0.201		
Never	554 (69)	511 (64)		71 (64)	65 (59)			
BMI at baseline survey ⁵	00 1 (00)	017 (0 1)		, , (0-7)	00 (00)			
Obese, ≥30 kg/m ²	46 (6)	59 (7)	0.548	10 (9)	14 (13)	0.134		
Overweight, ≥25 to <30 kg/m ²	146 (18)	141 (18)	0.040	15 (14)	25 (23)	J. 104		
Normal, ≥18.5 to <25 kg/m ²	546 (68)	540 (68)		79 (71)	68 (62)			
Underweight, <18.5 kg/m ²	64 (8)	57 (7)		7 (6)	3(3)			

BMI, body mass index; CCSs, childhood cancer survivors; FFQ, food frequency questionnaire; IQR, interquartile range; NA, not applicable; SCCSS, Swiss Childhood Cancer Survivor Study

¹This includes 635 CCSs who did not respond, 45 who refused, 11 who were pregnant or breastfeeding, 35 with missing dietary data, and 71 with unreliable dietary data.

² p-value calculated from chi-square statistics (categorical, 2-sided test) or Kruskal-Wallis test (continuous, nonparametric) to compare differences between

participants to nonparticipants.

This includes 9 CCSs with no valid address available anymore or being abroad, 15 who refused, and 86 who did not respond.

⁴ Age at follow-up survey is calculated for FFQ nonparticipants by taken the average participants' date of filling in the questionnaire.

⁵Based on information from the SCCSS baseline questionnaire filled in between 2007-2013.

⁶ Active: ≥150 minutes of moderate intense or 75 minutes of vigorous intense or a combination of moderate and vigorous intense physical activity per week.

Supplementary Table S3. Clinical characteristics of adult childhood cancer survivors by participants and nonparticipants to the SCCSS-Nutrition study

		FFQ		Morning f	asting spot u	rine
Characteristics, n (%)	Participants (n = 802)	Non- participants ¹ (n = 797)	p-value ²	Participants (n = 111)	Non- participants ³ (n = 110)	p-value ²
ICCC-3 diagnosis						
I: Leukemia	246 (31)	264 (33)	<0.001	30 (27)	27 (25)	0.607
II: Lymphoma	173 (22)	139 (17)		30 (27)	30 (27)	
III: CNS tumor	81 (10)	140 (18)		9 (8)	17 (15)	
IV: Neuroblastoma	28 (3)	31 (4)		4 (4)	3 (3)	
V: Retinoblastoma	12 (2)	22 (3)		2 (2)	3 (3)	
VI: Renal tumor	52 (6)	41 (5)		4 (4)	3 (3)	
VII: Hepatic tumor	6 (<1)	3 (<1)		1 (<1)	1 (<1)	
VIII: Bone tumor	50 (6)	29 (4)		11 (10)	5 (5)	
IX: Soft tissue sarcoma	66 (8)	32 (4)		7 (6)	10 (9)	
X: Germ cell tumor	43 (5)	42 (5)		8 (7)	4 (4)	
XI & XII: Other tumor	26 (3)	17 (2)		4 (4)	3 (3)	
Langerhans cell histiocytosis	19 (2)	37 (5)		1 (<1)	4 (4)	
Age at diagnosis,4 years	9.7 (3.9; 13.9)	8.0 (3.8; 13.1)	0.010	11.5 (5.0; 15.6)	10.7 (4.6; 15.8)	0.696
Time since diagnosis,4 years	26.1 (20.2; 31.7)	24.0 (19.5; 30.3)	0.186	28.3 (21.0; 32.7)	22.8 (18.5; 30.1)	0.004
Any irradiation	292 (36)	293 (37)	0.883	40 (36)	37 (34)	0.708
Chest irradiation	87 (11)	67 (8)	0.098	15 (14)	10 (9)	0.299
<30 Gy	31 (4)	21 (3)	0.218	3 (3)	3 (3)	0.498
≥30 Gy	56 (7)	46 (6)		12 (11)	7 (6)	
Any chemotherapy	642 (80)	606 (76)	0.052	80 (72)	78 (71)	0.848
Anthracyclines	298 (37)	298 (37)	0.923	39 (35)	26 (24)	0.061
HSCT	30 (4)	34 (4)	0.592	5 (5)	7 (6)	0.542
History of relapse	99 (12)	107 (13)	0.519	18 (16)	15 (14)	0.591

CCSs, childhood cancer survivor; CNS, central nervous system; Gy, gray; HSCT, hematopoietic stem cell transplantation; ICCC-3, International Childhood Cancer Classification, 3rd edition; SCCSS, Swiss Childhood Cancer Survivor Study

¹ This includes 635 CCSs who did not respond, 45 who refused, 11 who were pregnant or breastfeeding, 35 with missing dietary data, and 71 with

unreliable dietary data.

² p-value calculated from two-sample mean comparison test (t test) or chi-square statistics comparing participants excluded from analyses,

nonparticipants, and non-eligible CCS to participants (2-sided test).

³ This includes 9 CCSs with no valid address available anymore or being abroad, 15 who refused, and 86 who did not respond.

⁴ Values are medians with interquartile ranges (IQRs).

Supplementary Table S4. Dietary intake <u>based on self-reported food frequency questionnaire (FFQ) data</u> of adult childhood cancer survivors by participants to the SCCSS-Nutrition study

	DACH recommendations ¹		FFQ participants (n = 802)		FFQ but not participus n = 6	pants	Spot urine participants (n = 111)	
	male	female	mean ± SD	% DACH ²	mean ± SD	% DACH ²	mean ± SD	% DACH ²
Total energy, kcal ³	2300-3400	1800-2600	1639 ± 581	65	1654 ± 595	65	1543 ± 481	62
Macronutrients								
Protein, g	57	48	66.0 ± 29.3	126	67.0 ± 30.3	127	60.1 ± 21.2	116
% of energy ⁴	10-20	10-20	16.1 ± 3.6	107	16.2 ± 3.6	108	15.7 ± 3.2	105
Vegetal, g			18.2 ± 7.6		18.2 ± 7.8		17.7 ± 6.6	
Animal, g			47.9 ± 26.6		48.8 ± 27.6		42.4 ± 19.1	
Carbohydrates, g⁵	225-275	225-275	181 ± 74	72	182 ± 76	73	175 ± 65	70
% of energy ^{4, 5}	45-55	45-55	44.1 ± 8.9	88	44.0 ± 9.0	88	45.1 ± 8.0	90
Mono/disaccharides, g			86 ± 47		86 ± 47		86 ± 48	
Polysaccharides, g			94 ± 44		95 ± 46		88 ± 36	
Total fiber, g ⁵	≥30	≥30	12.2 ± 6.5	41	12.2 ± 6.5	41	12.1 ± 6.3	40
Total fat, g			67.9 ± 28.0		68.8 ± 28.4		62.7 ± 24.2	
Total fat, % of energy4	30-40	30-40	37.3 ± 6.9	107	37.4 ± 7.0	107	36.4 ± 6.5	104
Saturated fat, g			25.1 ± 11.6		25.5 ± 11.6		23.2 ± 11.0	
Saturated fat, % of energy6	<10	<10	13.7 ± 3.2	137	13.7 ± 3.1	137	13.3 ± 3.4	133
Monounsaturated fat, g			28.4 ± 12.6		28.8 ± 12.9		26.1 ± 10.4	
Polyunsaturated fat, g			8.6 ± 3.7		8.7 ± 3.8		7.9 ± 3.4	
Cholesterol, mg ⁴	300	300	341 ± 206	114	349 ± 215	116	288 ± 131	96
Micronutrients								
Vitamins								
A, mg-RE ⁷	1.0	0.8	0.7 ± 0.7	78	0.7 ± 0.7	78	0.7 ± 0.4	79
D, μg	20	20	2.4 ± 2.0	12	2.4 ± 2.0	12	2.4 ± 1.8	12
Minerals								
Calcium (Ca), mg	1000	1000	889 ± 467	89	901 ± 474	90	810 ± 416	81
Iron (Fe), mg	10 ⁴	15	9.0 ± 3.5	72	9.1 ± 3.7	73	8.4 ± 2.7	66
Phosphorus (P), mg	700	700	1227 ±448	175	1248 ± 462	178	1093 ± 320	156
Potassium (K), mg	4000	4000	2740 ± 1072	68	2759 ± 1080	68	2618 ± 1019	65
Sodium (Na), mg ⁸	1500	1500	2844 ± 659	190	2859 ± 656	190	2747 ± 674	183
Alcohol, g ⁹	20	10	5.7 ± 7.7	38	5.7 ± 8.0	38	5.7 ± 6.0	40

DACH, dietary recommendations for Germany (D), Austria (A) and Switzerland (CH); FFQ, food frequency questionnaire; SCCSS, Swiss Childhood Cancer Survivor Study; SD, standard deviation ¹ DACH recommendations for the general population age 20-50 years, excluding pregnant and lactating females, 2015.

³ Depending on physical activity level (PAL, 1.4-1.8) and age.

⁵ BLV (2009). Kohlenhydrate in unserer Ernährung – Empfehlungen des BLV. <u>www.blv.admin.ch</u>.

⁷ Total vitamin A = retinol + carotene/12, expressed in µg RAE.

⁹ Maximum tolerated dosage.

² Percentage of mean intake in relation to the DACH recommended intake level * 100. Recommended intake is estimated on the basis of the age-sex groups of the DACH guidelines, weighted by the age and sex distribution of the study population. For alcohol intake the maximum tolerated dosage was taken.

⁴ Federal Food Safety and Veterinary Office FSVO, www.naehrwertdaten.ch.

⁶ BLV (2012). Fett in unserer Ernährung – Empfehlung des BLV. www.blv.admin.chv, recommendations as a percentage of the daily energy demand and based on a daily energy intake of 2000 kcal.

⁸ Na intake based on FFQ data was calculated with the equation: (8.20 + 0.38*FFQ in males)/2.54 and (4.55 + 0.67*FFQ in females)/2.54, with 1 g Na = 2.54 g NaCl see Ref 35. Intake based on FFQ data only gives a mean daily intake of 1.8 g ± 0.7 for females and 1.9 g ± 0.8 for males.

Supplementary Table S5. Sodium (Na) and potassium (K) intake <u>based on food frequency questionnaire (FFQ) data</u> of adult childhood cancer survivors (CCSs) by language region within Switzerland

		CCSs									
	DACH ¹	FFQ participants n = 802						·			
		To : n = 8		German speaking n = 570		French/Italia n = 2					
		mean ± SD	% DACH ²	mean ± SD	% DACH ²	mean ± SD	% DACH ²	mean ± SD	% DACH ²	mean ± SD	% DACH ²
Na³, g	1.5	2.8 ± 0.7	190	2.9 ± 0.7	190	2.8 ± 0.7	188	2.7 ± 0.7	183	2.9 ± 0.7	191
K, g	4.0	2.7 ± 1.1	68	2.8 ± 1.1	69	2.6 ± 1.1	66	2.6 ± 1.0	65	2.8 ± 1.1	69
Na:K ratio⁴	-	1.2 ± 0.5	-	1.2 ± 0.5	-	1.2 ± 0.6	-	1.2 ± 0.5	-	1.2 ± 0.5	

CCS, childhood cancer survivors, DACH, dietary recommendations for Germany (D), Austria (A) and Switzerland (CH); FFQ, food frequency questionnaire, Na, sodium; K, potassium; SD, standard deviation

¹ DACH recommendations for the general population age 20-50 years, excluding pregnant and lactating females, 2015. Recommendations for Na and K are similar for males and females.

² Percentage of mean intake in relation to the DACH recommended intake level * 100. Recommended intake is estimated on the basis of the age-sex groups of the DACH guidelines, weighted by the age and sex distribution of the study population. For alcohol intake the maximum tolerated dosage was taken.

³ Na intake based on FFQ data was calculated with the equation: (8.20 + 0.38*FFQ in males)/2.54 and (4.55 + 0.67*FFQ in females)/2.54, with 1 g Na = 2.54 g NaCl see Ref 35. Intake based on FFQ data only gives a mean daily intake of 1.8 g ± 0.7 for females and 1.9 g ± 0.8 for males.

⁴ Na:K ratio was calculated by dividing the sodium intake (g/day) by the potassium intake (g/day).

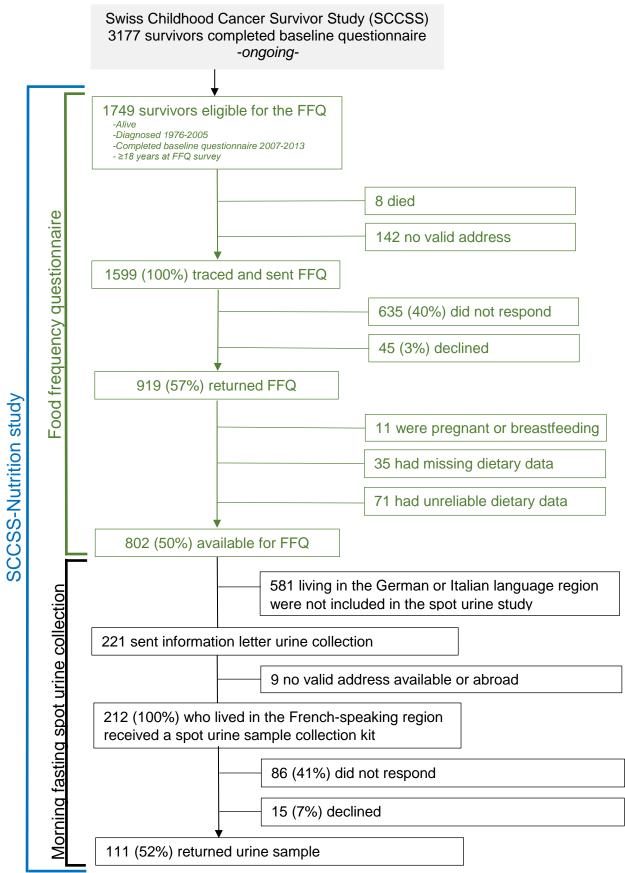


Figure 1. Response rates in the Swiss Childhood Cancer Survivor Study (SCCSS)-Nutrition study

FFQ, food frequency questionnaire

24. Système cardiovasculaire					
	Déjà eu?		Quand?	Enc prés	ore ent?
	Non	Oui		Non	Oui
Pression artérielle élevée (hypertension à répétition)			(année)		
Irrégularité du rythme cardiaque ou palpitations (arythmie), qui a-ont nécessité une investigation cardiologique			(année)		
Insuffisance cardiaque (insuffisance du muscle cardiaque)			(année)		
Maladie coronarienne (angine de poitrine: sensation de pression accompagné de douleurs au niveau de la poitrine à cause d'une moindre oxygénation du cœur)			(année)		
Infarctus du myocarde (crise cardiaque)			(année)		
Thrombose veineuse profonde (p. ex. thrombose d'une jambe)			(année)		
Embolie pulmonaire			(année)		
Attaque d'apoplexie (accident cérébrovasculaire etc.)			(année)		
Artériosclérose (Durcissement des artères des jambes, de la carotide ou des artères coronaires)			(année)		
Autres problèmes concernant le système cardiovasculaire Si oui, veuillez préciser			(année)		

Avez-vous eu des opérations du cœur ou du système cardiovasculaire depuis 2007? Si oui, veuillez indiquer toutes les opérations à la page 9, question 18.

Supplementary Figure 2A. Self-reported cardiovascular related medical history questions in <u>French</u> assessed within the Swiss Childhood Cancer Survivor Study (SCCSS)-Nutrition Study

	Irgendwann im Leben? Nein Ja		Seit wann?	Aktuell vorhai Nein	
Hoher Blutdruck (arterielle Hypertonie, wiederholt gemessen)			(Jahr)		
Herzrhythmusstörungen (Arrhythmien: unregelmässige Herzschläge), welche von einem Kardiologen (Herzspezialisten) abgeklärt werden mussten			(Jahr)		
Herzinsuffizienz (Herzmuskelschwäche)			(Jahr)		
Koronare Herzkrankheit (Angina pectoris: Engegefühl mit Schmerzen in der Brust, das durch einen Sauerstoffmangel im Herzen auftritt)			(Jahr)		
Herzinfarkt			(Jahr)		
Tiefe Venenthrombose (Blutgerinnsel z.B. im Bein)			(Jahr)		
Lungenembolie			(Jahr)		
Schlaganfall (Hirninfarkt oder Hinrblutung)			(Jahr)		
Arteriosklerose (Arterienverkalkung der Beine, Halsschlagader oder Herzkranzgefässe)			(Jahr)		
Andere Probleme, die das Herz- und Kreislaufsystem betreffen. Falls Ja, bitte beschreiben Sie die Probleme näher:			(Jahr)		

Wurden Sie seit 2007 am Herzen operiert oder hatten Sie Eingriffe am Herz- oder Kreislaufsystem? Falls Ja, bitte geben Sie alle Operationen auf Seite 9, Frage 18 an.

Supplementary Figure 2B. Self-reported cardiovascular related medical history questions in <u>German</u> assessed within the Swiss Childhood Cancer Survivor Study (SCCSS)-Nutrition Study