Supplemental Online Content

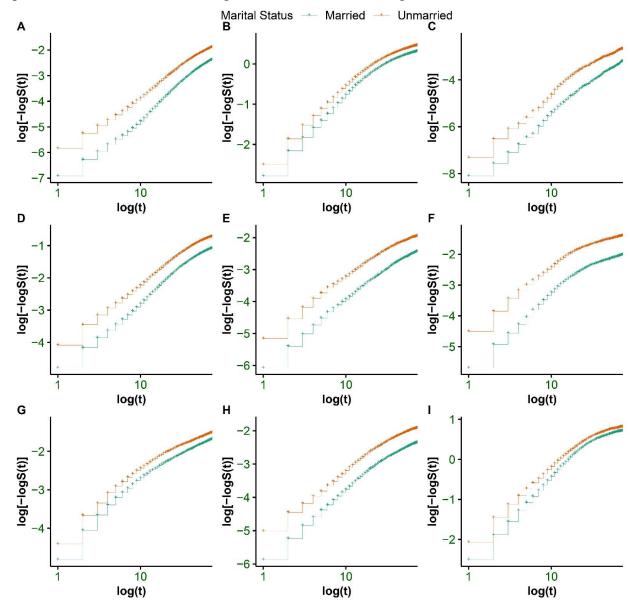
Chen ZH, Yang KB, Zhang YZ, et al. Assessment of modifiable factors for the association of marital status with cancer-specific survival. *JAMA Netw Open*. 2021;4(5):e2111813. doi:10.1001/jamanetworkopen.2021.11813

eFigure 1. Evaluation of the Appropriateness of Weibull Distribution in AFT Models Assessing the Association Between Marital Status and Cancer-Specific Survival for the Following Cancer Sites: Breast, Lung, Prostate, Colorectum, Melanoma, Bladder, Kidney, Endometrium, and Pancreas eTable 1. Patient Demographic and Clinical Characteristics by Marital Status and Cancer Site **eFigure 2.** Survival Curves of Cancer-Specific Survival of Cancer of the Lung, Colorectum, Melanoma, Bladder, Kidney, Pancreas, and Breast by Marital Status and Sex eTable 2. Associations Between Cancer-Specific Survival and Marital Status (Married vs Unmarried), Stage at Diagnosis, and Treatment eTable 3. Associations Between Cancer-Specific Survival and Marital Status (Married vs Unmarried), Stage at Diagnosis, and Treatment by Cancer Site and Histological Subtype eTable 4. Mediation Analyses of Direct and Indirect Effects of Marital Status on Cancer-Specific Survival Mediated by Stage at Diagnosis and Treatment and Receiving Radiation Therapy **eTable 5.** Mediation Analyses of Direct and Indirect Effects of Marital Status on Cancer-Specific Survival Mediated by Stage at Diagnosis and Treatment by Histological Subtype eTable 6. Mediation Analyses of Direct and Indirect Effects of Marital Status on Cancer-Specific Survival Mediated by Stage at Diagnosis and Treatment by Sex

This supplemental material has been provided by the authors to give readers additional information about their work.

eTable 7. Demographic and Clinical Characteristics of Individuals Included and Excluded

eFigure 1. Evaluation of the Appropriateness of Weibull Distribution in AFT Models Assessing the Association Between Marital Status and Cancer-Specific Survival for the Following Cancer Sites: Breast, Lung, Prostate, Colorectum, Melanoma, Bladder, Kidney, Endometrium, and Pancreas



eTable 1. Patient Demographic and Clinical Characteristics by Marital Status and Cancer Site

		Total Sample			Breast			Lung			Prostate	
Characteristics	Married	Unmarried	_	Married	Unmarried	_	Married	Unmarried	_	Married	Unmarried	_
	1067726	666180	P	259555	186601	P	146391	127669	P	287143	96782	P
A 4 . 1°	63.05	64.89	<0.001	58.25	63.16	<0.001	67.26	67.87	<0.001	65.46	65.14	<0.001
Age at diagnosis ^a	(11.74)	(13.79)	<0.001	(12.24)	(14.56)	< 0.001	(10.28)	(11.43)	<0.001	(8.71)	(9.50)	< 0.001
Race ^b			< 0.001			< 0.001			< 0.001			< 0.001
White	883655	509551		212617	140483		121240	99304		230879	67121	
winte	(82.8)	(76.5)		(81.9)	(75.3)		(82.8)	(77.8)		(80.4)	(69.4)	
Dlaala	88389	112906		18009	31463		11267	21225		36093	24908	
Black	(8.3)	(16.9)		(6.9)	(16.9)		(7.7)	(16.6)		(12.6)	(25.7)	
Others	87585	39835		27613	13714		13626	6917		16621	3580	
Others ^c	(8.2)	(6.0)		(10.6)	(7.3)		(9.3)	(5.4)		(5.8)	(3.7)	
TT.I	8097	3888		1316	941		258	223		3550	1173	
Unknown	(0.8)	(0.6)		(0.5)	(0.5)		(0.2)	(0.2)		(1.2)	(1.2)	
Sex ^b			< 0.001			< 0.001			< 0.001			< 0.001
	478681	415698		257331	185593		56406	73394				
Female	(44.8)	(62.4)		(99.1)	(99.5)		(38.5)	(57.5)				
M.I.	589045	250482		2224	1008		89985	54275		287143	96782	
Male	(55.2)	(37.6)		(0.9)	(0.5)		(61.5)	(42.5)		(100.0)	(100.0)	
Grade ^b			< 0.001			< 0.001			< 0.001			< 0.001
Codel	126551	80027		56790	38231		8312	6221		22102	7612	
Grade I	(11.9)	(12.0)		(21.9)	(20.5)		(5.7)	(4.9)		(7.7)	(7.9)	
C 1 H	383913	228578		108196	76796		26540	21324		119954	37760	
Grade II	(36.0)	(34.3)		(41.7)	(41.2)		(18.1)	(16.7)		(41.8)	(39.0)	
C - 1 III	309987	178684		81178	60111		38553	32959		137400	47277	
Grade III	(29.0)	(26.8)		(31.3)	(32.2)		(26.3)	(25.8)		(47.9)	(48.8)	
Cuada IV	35305	25483		1192	939		5083	4560		472	240	
Grade IV	(3.3)	(3.8)		(0.5)	(0.5)		(3.5)	(3.6)		(0.2)	(0.2)	
TT1	211970	153408		12199	10524		67903	62605		7215	3893	
Unknown	(19.9)	(23.0)		(4.7)	(5.6)		(46.4)	(49.0)		(2.5)	(4.0)	

eTable 1 cont.

Characteristics		Total Sample			Breast		Lung			Prostate		
	Married	Unmarried	_	Married	Unmarried	_	Married	Unmarried	_	Married	Unmarried	_
	1067726	666180	P	259555	186601	P	146391	127669	P	287143	96782	P
Surgery ^b			< 0.001			< 0.001			< 0.001			< 0.001
	317376	219538		11755	16310		107078	100184		149111	59887	
No	(29.7)	(33.0)		(4.5)	(8.7)		(73.1)	(78.5)		(51.9)	(61.9)	
*7	750350	446642		247800	170291		39313	27485		138032	36895	
Yes	(70.3)	(67.0)		(95.5)	(91.3)		(26.9)	(21.5)		(48.1)	(38.1)	
Radiation ^b			< 0.001			< 0.001			< 0.001			< 0.001
N. (F)	711213	448228		117187	96243		79872	71315		187570	61289	
No/Unknown	(66.6)	(67.3)		(45.1)	(51.6)		(54.6)	(55.9)		(65.3)	(63.3)	
T 7	356513	217952		142368	90358		66519	56354		99573	35493	
Yes	(33.4)	(32.7)		(54.9)	(48.4)		(45.4)	(44.1)		(34.7)	(36.7)	
Chemotherapy ^b			< 0.001			< 0.001			< 0.001			< 0.001
N. (F)	751265	456478		142721	114570		62616	65693		284576	95550	
No/Unknown	(70.4)	(68.5)		(55.0)	(61.4)		(42.8)	(51.5)		(99.1)	(98.7)	
\$ 7	316461	209702		116834	72031		83775	61976		2567	1232	
Yes	(29.6)	(31.5)		(45.0)	(38.6)		(57.2)	(48.5)		(0.9)	(1.3)	
Stage ^b			< 0.001			< 0.001			< 0.001			< 0.001
Paula.	760931	429026		222976	152277		42196	35925		242680	79180	
Early	(71.3)	(64.4)		(85.9)	(81.6)		(28.8)	(28.1)		(84.5)	(81.8)	
A d	306795	237154		36579	34324		104195	91744		44463	17602	
Advanced	(28.7)	(35.6)		(14.1)	(18.4)		(71.2)	(71.9)		(15.5)	(18.2)	
Insurance ^b			< 0.001			< 0.001			< 0.001			< 0.001
Insured	960622	514788		235173	146260		128801	93827		261955	78997	
insurea	(90.0)	(77.3)		(90.6)	(78.4)		(88.0)	(73.5)		(91.2)	(81.6)	
Medicaid	64888	116159		18208	34021		12855	27039		9962	10932	
Medicald	(6.1)	(17.4)		(7.0)	(18.2)		(8.8)	(21.2)		(3.5)	(11.3)	
Uninsured	17243	22719		3280	4151		3134	5139		3031	2844	
Uninsurea	(1.6)	(3.4)		(1.3)	(2.2)		(2.1)	(4.0)		(1.1)	(2.9)	
Unknown	24973	12514		2894	2169		1601	1664		12195	4009	
UHKHOWH	(2.3)	(1.9)		(1.1)	(1.2)		(1.1)	(1.3)		(4.2)	(4.1)	

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Characteristics		Total Sample			Breast			Lung		Prostate		
	Married	Unmarried	n	Married	Unmarried	n	Married	Unmarried	n	Married	Unmarried	P
	1067726	666180	P	259555	186601	P	146391	127669	P	287143	96782	1
Dovontvil	32.02	33.00	< 0.001	31.64	32.70	< 0.001	32.97	33.60	<0.001	31.79	33.17	<0.001
Poverty ^a	(9.57)	(9.50)	\0.001	(9.39)	(9.33)	\0.001	(9.92)	(9.77)	<0.001	(9.53)	(9.53)	<0.001
Education	14.71	15.12	< 0.001	14.54	15.01	<0.001	15.08	15.21	<0.001	14.60	15.20	<0.001
Education ^a	(6.20)	(6.10)	<0.001	(6.14)	(6.05)	<0.001	(6.35)	(6.20)	<0.001	(6.13)	(6.01)	\0.001
Residenceb			< 0.001			< 0.001			< 0.001			< 0.001
Matuanalitan	939181	591974		232835	168658		123404	110042		255010	86575	
Metropolitan	(88.0)	(88.9)		(89.7)	(90.4)		(84.3)	(86.2)		(88.8)	(89.5)	
	127647	73611		26467	17759		22817	17483		32045	10167	
Nonmetropolitan	(12.0)	(11.0)		(10.2)	(9.5)		(15.6)	(13.7)		(11.2)	(10.5)	
Unknown	898	595		253	184		170	144		88	40	
Unknown	(0.1)	(0.1)		(0.1)	(0.1)		(0.1)	(0.1)		(0.0)	(0.0)	

Characteristics		Colorectum			Melanoma		Bladder			Kidney		
	Married	Unmarried	_	Married	Unmarried	_	Married	Unmarried	_	Married	Unmarried	_
	135567	104409	P	67135	31666	P	31189	20564	P	57392	32936	P
Age at diagnosis ^a	63.70	66.69	< 0.001	59.68	58.03	< 0.001	70.30	71.56	< 0.001	61.10	60.54	< 0.001
Age at diagnosis	(12.82)	(14.91)	<0.001	(14.37)	(19.45)	<0.001	(11.02)	(13.10)	<0.001	(11.89)	(14.47)	<0.001
Raceb			< 0.001			< 0.001			< 0.001			< 0.001
White	108889	78895		65035	30506		27826	17461		48239	25338	
winte	(80.3)	(75.6)		(96.9)	(96.3)		(89.2)	(84.9)		(84.1)	(76.9)	
Black	11268	17295		216 (0.3)	305		1383	2153		4624	5543	
Diack	(8.3)	(16.6)		210 (0.3)	(1.0)		(4.4)	(10.5)		(8.1)	(16.8)	
Othersc	14716	7793		609 (0.9)	363		1846	861		4058	1826	
Others	(10.9)	(7.5)		009 (0.9)	(1.1)		(5.9)	(4.2)		(7.1)	(5.5)	
Unknown	694 (0.5)	426		1275	492		134 (0.4)	89		471 (0.8)	229	
UIKIIOWII	094 (0.3)	(0.4)		(1.9)	(1.6)		134 (0.4)	(0.4)		4/1 (0.8)	(0.7)	
Sex ^b			< 0.001			< 0.001			< 0.001			< 0.001
Female	53558	60503		26003	16537		4890	7651		17506	15534	
remaie	(39.5)	(57.9)		(38.7)	(52.2)		(15.7)	(37.2)		(30.5)	(47.2)	
Male	82009	43906		41132	15129		26299	12913		39886	17402	
Male	(60.5)	(42.1)		(61.3)	(47.8)		(84.3)	(62.8)		(69.5)	(52.8)	
$\mathbf{Grade}^{\mathbf{b}}$			< 0.001			< 0.001			< 0.001			< 0.001
Grade I	12291	9002		108 (0.2)	37		986 (3.2)	655		5218	3037	
Grader	(9.1)	(8.6)		108 (0.2)	(0.1)		980 (3.2)	(3.2)		(9.1)	(9.2)	
Grade II	86670	65058		170 (0.3)	81		2921	1885		22688	12572	
Grade II	(63.9)	(62.3)		170 (0.3)	(0.3)		(9.4)	(9.2)		(39.5)	(38.2)	
Grade III	19407	15809		293 (0.4)	162		6989	4734		13885	7440	
Grade III	(14.3)	(15.1)		293 (0.4)	(0.5)		(22.4)	(23.0)		(24.2)	(22.6)	
Grade IV	3171	2730		103 (0.2)	73		17519	11224		4960	2878	
Grautiv	(2.3)	(2.6)		103 (0.2)	(0.2)		(56.2)	(54.6)		(8.6)	(8.7)	
Unknown	14028	11810		66461	31313		2774	2066		10641	7009	
UHKHUWH	(10.3)	(11.3)		(99.0)	(98.9)		(8.9)	(10.0)		(18.5)	(21.3)	

eTable 1 cont.

Characteristics		Colorectum			Melanoma			Bladder			Kidney		
	Married	Unmarried		Married	Unmarried		Married	Unmarried		Married	Unmarried		
	135567	104409	P	67135	31666	P	31189	20564	P	57392	32936	P	
Surgeryb			< 0.001			< 0.001			< 0.001			< 0.001	
> T	14010	14956		2025 (4.5)	1905		1125	1132		4986	4141		
No	(10.3)	(14.3)		3035 (4.5)	(6.0)		(3.6)	(5.5)		(8.7)	(12.6)		
***	121557	89453		64100	29761		30064	19432		52406	28795		
Yes	(89.7)	(85.7)		(95.5)	(94.0)		(96.4)	(94.5)		(91.3)	(87.4)		
Radiation ^b			< 0.001			< 0.001			< 0.001			< 0.001	
N. (T.)	114023	89232		65172	30417		28403	18109		54692	31247		
No/Unknown	(84.1)	(85.5)		(97.1)	(96.1)		(91.1)	(88.1)		(95.3)	(94.9)		
**	21544	15177		10(2(20)	1249		2786	2455		2700	1689		
Yes	(15.9)	(14.5)		1963 (2.9)	(3.9)		(8.9)	(11.9)		(4.7)	(5.1)		
Chemotherapy ^b			< 0.001			< 0.001			< 0.001			< 0.001	
N. (T.)	73477	62568		65513	30626		20094	13974		51284	29516		
No/Unknown	(54.2)	(59.9)		(97.6)	(96.7)		(64.4)	(68.0)		(89.4)	(89.6)		
***	62090	41841		1(22 (2.4)	1040		11095	6590		6108	3420		
Yes	(45.8)	(40.1)		1622 (2.4)	(3.3)		(35.6)	(32.0)		(10.6)	(10.4)		
Stage ^b			< 0.001			< 0.001			< 0.001			< 0.001	
E. J	74593	55235		59267	26438		24515	15274		40604	22940		
Early	(55.0)	(52.9)		(88.3)	(83.5)		(78.6)	(74.3)		(70.7)	(69.7)		
A.1 1	60974	49174		7868	5228		6674	5290		16788	9996		
Advanced	(45.0)	(47.1)		(11.7)	(16.5)		(21.4)	(25.7)		(29.3)	(30.3)		
Insurance ^b			< 0.001			< 0.001			< 0.001			< 0.001	
I	119381	78474		60430	25927		28470	16365		51640	24320		
Insured	(88.1)	(75.2)		(90.0)	(81.9)		(91.3)	(79.6)		(90.0)	(73.8)		
Maria	100(7 (0.1)	19840		1442 (2.1)	2955		1776	3165		3996	6721		
Medicaid	10967 (8.1)	(19.0)		1442 (2.1)	(9.3)		(5.7)	(15.4)		(7.0)	(20.4)		
II	2245 (2.5)	4614		0(5 (1.4)	1152		125 (1.4)	699		1140	1506		
Uninsured	3345 (2.5)	(4.4)		965 (1.4)	(3.6)		435 (1.4)	(3.4)		(2.0)	(4.6)		
II.	1074 (1.4)	1481		4209 (6.4)	1632		500 (1.6)	335		616 (1.1)	389		
Unknown	1874 (1.4)	(1.4)		4298 (6.4)	(5.2)		508 (1.6)	(1.6)		616 (1.1)	(1.2)		

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Characteristics		Colorectum			Melanoma			Bladder		Kidney		
	Married	Unmarried	n	Married	Unmarried	n	Married	Unmarried	n	Married	Unmarried	n
	135567	104409	P	67135	31666	P	31189	20564	P	57392	32936	P
Poverty ^a	32.67 (9.64)	33.31	< 0.001	30.80	31.47	< 0.001	31.98	32.70	< 0.001	32.80	33.54	< 0.001
roverty.	32.07 (9.04)	(9.51)	<0.001	(9.17)	(9.01)	<u> </u>	(9.53)	(9.52)	\0.001	(9.69)	(9.48)	\0.001
Ed	15 20 (6 22)	15.40	<0.001	13.85	14.38	<0.001	14.72	15.08	< 0.001	15.14	15.41	< 0.001
Education ^a	15.20 (6.33)	(6.17)	< 0.001	(5.90)	(5.91)	<0.001	(6.22)	(6.16)	<0.001	(6.38)	(6.25)	٠٥.001
Residence ^b			< 0.001			< 0.001			< 0.001			< 0.001
Maturalitan	117408	91984		59427	28367		27222	18088		50057	29148	
Metropolitan	(86.6)	(88.1)		(88.5)	(89.6)		(87.3)	(88.0)		(87.2)	(88.5)	
	17930	12278		7694	3292		3952	2467		7274	3757	
Nonmetropolitan	(13.2)	(11.8)		(11.5)	(10.4)		(12.7)	(12.0)		(12.7)	(11.4)	
I labanas	220 (0.2)	147		14	7		15	9		61	31	
Unknown	229 (0.2)	(0.1)		(0.0)	(0.0)		(0.0)	(0.0)		(0.1)	(0.1)	

Characteristic	Eı	ndometrium			Pancreas	
	Married	Unmarried	_	Married	Unmarried	
	49528	42479	P	33826	23074	P
Age at diagnosis ^a	60.47	62.72	< 0.001	65.78	66.91 (12.95)	< 0.001
Age at ulagilosis	(10.40)	(12.70)	\0.001	(10.97)	00.91 (12.93)	\0.001
Raceb			< 0.001			< 0.001
White	41197 (83.2)	33210		27733	17233 (74.7)	
white	41197 (65.2)	(78.2)		(82.0)	17233 (74.7)	
Dlask	2774 (5.6)	5836		2755	4178	
Black	2774 (5.6)	(13.7)		(8.1)	(18.1)	
Othors	5225 (10.6)	3169		3261	1612	
Others ^c	5235 (10.6)	(7.5)		(9.6)	(7.0)	
TI.1	222 (0.7)	264		77	51	
Unknown	322 (0.7)	(0.6)		(0.2)	(0.2)	
Sexb			< 0.001			< 0.001
F I.	49528	42479		13459	14007 ((0.7)	
Female	(100.0)	(100.0)		(39.8)	14007 (60.7)	
Mala				20367	9067	
Male				(60.2)	(39.3)	
Grade ^b			< 0.001			< 0.001
Grade I	18481 (37.3)	13873		2263	1359	
Graue I	10401 (37.3)	(32.7)		(6.7)	(5.9)	
Coole II	1007((22.2)	9521		5798	3581	
Grade II	10976 (22.2)	(22.4)		(17.1)	(15.5)	
C. A. III	(077 (12.0)	6891		5405	3301	
Grade III	6877 (13.9)	(16.2)		(16.0)	(14.3)	
Coods IV	2426 (4.0)	2621		369	218	
Grade IV	2436 (4.9)	(6.2)		(1.1)	(0.9)	
II	10759 (21.7)	9573		19991	14(15 ((2.2)	
Unknown	10758 (21.7)	(22.5)		(59.1)	14615 (63.3)	

eTable 1 cont.

Characteristic	En	dometrium			Pancreas	
	Married	Unmarried		Married	Unmarried	
	49528	42479	P	33826	23074	P
Surgeryb			< 0.001			< 0.001
NI.	1720 (2.5)	3012		24547	10011 (70.1)	
No	1729 (3.5)	(7.1)		(72.6)	18011 (78.1)	
Yes	47799 (96.5)	39467		9279	5063	
1 65	47799 (90.3)	(92.9)		(27.4)	(21.9)	
Radiation ^b			< 0.001			< 0.001
No/Unknown	37063 (74.8)	31061		27231	19315 (83.7)	
140/ CHKHOWH	37003 (74.8)	(73.1)		(80.5)	19313 (63.7)	
Yes	12465 (25.2)	11418		6595	3759	
103	12403 (23.2)	(26.9)		(19.5)	(16.3)	
Chemotherapyb			< 0.001			< 0.001
No/Unknown	40174 (81.1)	33713		10810	10268 (44.5)	
140/ CHKHOWH	40174 (01.1)	(79.4)		(32.0)	10268 (44.5)	
Yes	9354 (18.9)	8766		23016	12806 (55.5)	
103	7554 (10.7)	(20.6)		(68.0)	12000 (33.3)	
Stage ^b			< 0.001			< 0.001
Early	40991 (82.8)	32955		13109	8802	
Larry	40771 (62.6)	(77.6)		(38.8)	(38.1)	
Advanced	8537 (17.2)	9524		20717	14272 (61.9)	
7 tu vanecu	0337 (17.2)	(22.4)		(61.2)	11272 (01.5)	
Insurance ^b			< 0.001			< 0.001
Insured	44571 (90.0)	33131		30201	17487 (75.8)	
Insured	11371 (30.0)	(78.0)		(89.3)	17 107 (73.0)	
Medicaid	3161 (6.4)	7142		2521	4344	
Modicala	3101 (0.1)	(16.8)		(7.5)	(18.8)	
Uninsured	1194 (2.4)	1684		719 (2.1)	930	
Ciniisui cu	1171 (2.7)	(4.0)		(2.1)	(4.0)	
Unknown	602 (1.2)	522		385 (1.1)	313	
CHRIIOWII	002 (1.2)	(1.2)		303 (1.1)	(1.4)	

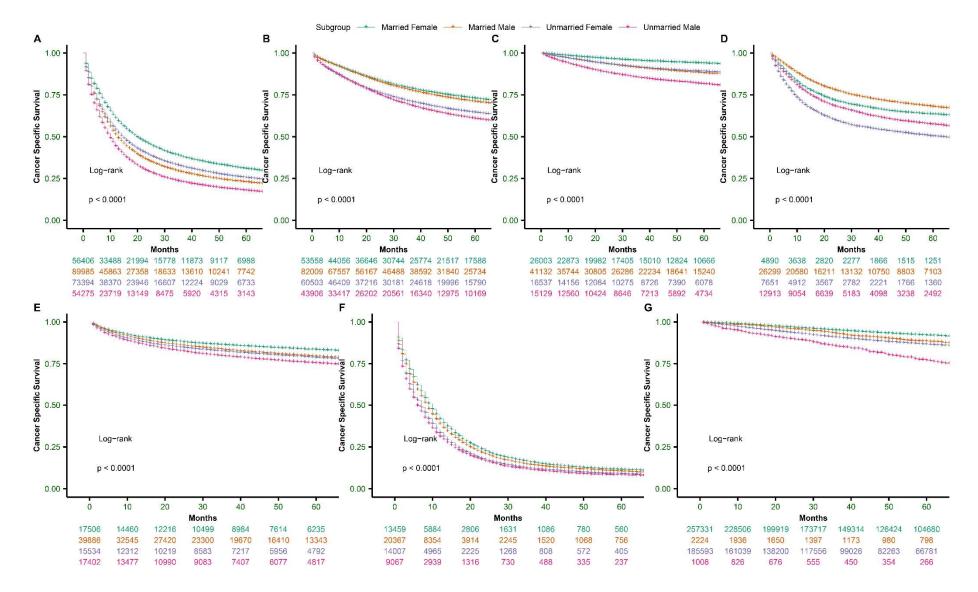
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eTable 1 cont.

Characteristic	En	dometrium			Pancreas	
	Married	Unmarried	n	Married	Unmarried	n
	49528	42479	P	33826	23074	P
Poverty ^a	31.65 (9.52)	32.34	< 0.001	32.00	32.75 (9.51)	<0.001
roverty	31.03 (9.32)	(9.38)	\0.001	(9.52)	32.73 (9.31)	<0.001
Education ^a	14 50 (6 10)	14.88	< 0.001	14.72	14.95 (6.07)	<0.001
Education"	14.50 (6.19)	(6.07)	<0.001	(6.18)	14.93 (0.07)	<0.001
Residence ^b			< 0.001			< 0.001
Matuanalitan	12027 (00.5)	38337		29981	20775 (00.0)	
Metropolitan	43837 (88.5)	(90.2)		(88.6)	20775 (90.0)	
	5650 (11.4)	4124		3810	2284	
Nonmetropolitan	5658 (11.4)	(9.7)		(11.3)	(9.9)	
Unknow	33 (0.1)	18		35	15	
Unknown		(0.0)		(0.1)	(0.1)	

^amean (SD); ^bN (%); ^cinclude American Indian, AK Native, Asian and Pacific Islander

eFigure 2. Survival Curves of Cancer-Specific Survival of Cancer of the Lung, Colorectum, Melanoma, Bladder, Kidney, Pancreas, and Breast by Marital Status and Sex



eTable 2. Associations Between Cancer-Specific Survival and Marital Status (Married vs Unmarried), Stage at Diagnosis, and Treatment

D: 1 6 4	Total Sample	;	Breast		Lung		Prostate	
Risk factors	TRe (95%CI)	P	TRe (95%CI)	P	TRe (95%CI)	P	TRe (95%CI)	P
Stage ^a								
Early	Reference		Reference		Reference		Reference	
Advanced	0.051 (0.051, 0.052)	< 0.001	0.125 (0.122, 0.128)	< 0.001	0.158 (0.155, 0.161)	< 0.001	0.139 (0.134, 0.144)	< 0.001
Surgeryb								
No	Reference		Reference		Reference		Reference	
Yes	3.442 (3.417, 3.467)	< 0.001	3.824 (3.729, 3.922)	< 0.001	4.735 (4.638, 4.834)	< 0.001	4.137 (3.979, 4.301)	< 0.001
Radiotherapy ^c								
No	Reference		Reference		Reference		Reference	
Yes	1.446 (1.436, 1.456)	< 0.001	1.251 (1.226, 1.276)	0.015	1.277 (1.263, 1.291)	< 0.001	1.587 (1.539, 1.635)	< 0.001
Chemotherapy ^d								
No	Reference		Reference		Reference		Reference	1
Yes	0.727 (0.722, 0.733)	< 0.001	0.973 (0.952, 0.995)	< 0.001	1.837 (1.815, 1.86)	< 0.001	0.418 (0.396, 0.441)	< 0.001
	Colorectum		Melanoma		Bladder		Kidney	
	TRe (95%CI)	P	TR ^e (95%CI)	P	TR ^e (95%CI)	P	TRe (95%CI)	P
Stage ^a	Reference		Reference		Reference		Reference	
Early	0.100 (0.097, 0.103)	< 0.001	0.047 (0.044, 0.05)	< 0.001	0.138 (0.131, 0.144)	< 0.001	0.042 (0.039, 0.045)	< 0.001
Advanced								
Surgeryb	Reference		Reference		Reference		Reference	
No	6.593 (6.447, 6.743)	< 0.001	2.263 (2.128, 2.408)	< 0.001	2.335 (2.16, 2.525)	< 0.001	6.976 (6.606, 7.367)	< 0.001
Yes								
Radiotherapy ^c	Reference		Reference		Reference		Reference	
No	1.25 (1.22, 1.282)	< 0.001	0.461 (0.432, 0.492)	< 0.001	0.525 (0.497, 0.556)	< 0.001	0.591 (0.563, 0.62)	< 0.001
Yes								
Chemotherapy ^d	Reference	•	Reference		Reference		Reference	•
No	1.33 (1.302, 1.359)	< 0.001	0.485 (0.454, 0.518)	< 0.001	1.039 (0.994, 1.086)	0.094	0.662 (0.633, 0.692)	< 0.001
Yes								

Risk factors	Endometriun	n	Pancreas	
	TR ^e (95%CI)	P	TR ^e (95%CI)	P
Stage ^a				
Early	Reference		Reference	
Advanced	0.093 (0.088, 0.099)	< 0.001	0.384 (0.375, 0.393)	< 0.001
Surgery ^b				
No	Reference		Reference	
Yes	6.582 (6.225, 6.959)	< 0.001	3.156 (3.056, 3.259)	< 0.001
Radiotherapy ^c				
No	Reference		Reference	
Yes	1.323 (1.266, 1.383)	< 0.001	1.152 (1.123, 1.182)	< 0.001
Chemotherapy ^d				
No	Reference		Reference	
Yes	0.954 (0.907, 1.003)	0.068	1.548 (1.516, 1.581)	< 0.001

^aAfter adjusted for age, race, marital status, sex, grade, residence, poverty level and educational level; ^bAfter adjusted for age, race, marital status, sex, grade, residence, poverty level, educational level, stage at diagnosis, radiotherapy, and chemotherapy; ^cAfter adjusted for age, race, marital status, sex, grade, residence, poverty level, educational level, stage at diagnosis, surgery, and chemotherapy; ^dAfter adjusted for age, race, marital status, sex, grade, residence, poverty level, educational level, stage at diagnosis, surgery, and radiotherapy; ^cTime ratio

eTable 3. Associations Between Cancer-Specific Survival and Marital Status (Married vs Unmarried), Stage at Diagnosis, and Treatment by Cancer Site and Histological Subtype

	CSS	1	Diagnosed Advanced		Receiving S	Surgery	Receiv Chemoth	_	Receivin	g RT ^f
Cancer and Sex	TR ^b for Marriage (95% CI)	P	OR ^c for Marriage (95% CI)	P	OR ^d for Marriage (95% CI)	P	OR ^e for Marriage (95% CI)	P	ORg for Marriage(9 5% CI)	P
Lung										
SCLC ^h	1.072(1.046, 1.099)	< 0.001	1.019(0.945, 1.100)	0.621	1.374(1.197, 1.58)	<0.001	1.36(1.284, 1.441)	<0.001	1.127(1.076, 1.179)	<0.001
NSCLC ⁱ	1.105(1.091, 1.118)	<0.001	1.003(0.983, 1.023)	0.774	1.437(1.397, 1.477)	< 0.001	1.478(1.45, 1.507)	<0.001	1.046(1.027, 1.066)	<0.001
Pancreas										
Adenocarci noma	1.088(1.067, 1.109)	<0.001	1.030(0.990, 1.072)	0.147	1.292(1.212, 1.379)	<0.001	1.615(1.551 ,1.682)	<0.001	1.025(0.975, 1.079)	0.332
pNET ^j	1.340(1.154, 1.556)	<0.001	0.884(0.738, 1.058)	0.180	1.253(0.984, 1.596)	0.067	1.078(0.878	0.472	1.111(0.804, 1.548)	0.529
Breast	,		,				,			
Triple Negative	1.090(1.023, 1.161)	0.008	0.819(0.762, 0.879)	<0.001	1.255(1.112, 1.416)	<0.001	1.117(1.026 ,1.217)	0.011	1.072(1.01,1	0.023
HER2 Enriched	1.171(1.042, 1.316)	<0.001	0.779(0.708, 0.857)	<0.001	1.369(1.172, 1.599)	<0.001	0.986(0.87, 1.116)	0.821	1.029(0.939, 1.127)	0.544
Luminal A	1.110(1.065- 1.157)	<0.001	0.835(0.806, 0.865)	<0.001	1.298(1.214, 1.387)	<0.001	1.1(1.069,1. 131)	<0.001	1.052(1.025, 1.078)	<0.001
Luminal B	1.093(0.995- 1.201)	0.041	0.865(0.808, 0.926)	<0.001	1.447(1.293, 1.62)	<0.001	1.118(1.037 ,1.204)	0.004	1.025(0.967, 1.086)	0.409

^aCancer-specific survival; ^bTime ratio, adjusted for patient demographics (age, race, sex, insurance status, residence, poverty level and educational level), stage, grade, receiving surgery, receiving chemotherapy and receiving radiation therapy; ^cOdds ratio, adjusted for patient demographics, and grade; ^dadjusted for patient demographics, stage, grade, receiving surgery and receiving radiation therapy; ^eadjusted for patient demographics, stage, grade, receiving surgery and receiving surgery and receiving radiation therapy; ^fRadiation therapy; ^gadjusted for patient demographics, stage, grade, receiving surgery; ^hSCLC, small cell lung cancer; ⁱNSCLC, non-small cell lung cancer; ⁱpNET, pancreatic neuroendocrine tumor.

eTable 4. Mediation Analyses of Direct and Indirect Effects of Marital Status on Cancer-Specific Survival Mediated by Stage at Diagnosis and Treatment and Receiving Radiation Therapy

N. 1	Total Sample		Breast		Lung		Prostate	
Mediators	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P
Stage								
TE ^a	0.387 (0.378,0.396)	< 0.001	0.275 (0.253,0.296)	< 0.001	0.177 (0.163,0.191)	< 0.001	0.451 (0.421,0.48)	< 0.001
NIEb	0.065 (0.063,0.067)	< 0.001	0.028 (0.026,0.03)	< 0.001	-0.002 (-0.01,0.006)	< 0.001	0.013 (0.011,0.015)	< 0.001
NDEc	0.322 (0.313,0.331)	< 0.001	0.247 (0.226,0.268)	< 0.001	0.179 (0.167,0.191)	< 0.001	0.437 (0.408,0.467)	< 0.001
PM ^d	0.196 (0.195,0.197)	< 0.001	0.114 (0.112,0.116)	< 0.001	-0.013 (-0.015, -0.01)	< 0.001	0.036 (0.034,0.038)	< 0.001
Surgery								
TE	0.341 (0.335,0.348)	< 0.001	0.191 (0.171,0.21)	< 0.001	0.19 (0.177,0.203)	< 0.001	0.474 (0.446,0.503)	< 0.001
NIE	0.034 (0.033,0.036)	< 0.001	0.005 (0.004,0.006)	< 0.001	0.095 (0.087,0.102)	< 0.001	0.161 (0.153,0.168)	< 0.001
NDE	0.307 (0.301,0.314)	< 0.001	0.186 (0.166,0.205)	< 0.001	0.096 (0.084,0.107)	< 0.001	0.314 (0.286,0.341)	< 0.001
PM	0.117 (0.116,0.117)	< 0.001	0.029 (0.027,0.03)	< 0.001	0.522 (0.519,0.524)	< 0.001	0.393 (0.39,0.396)	< 0.001
Radiotherapy								
TE	0.318 (0.312,0.325)	< 0.001	0.194 (0.175,0.214)	< 0.001	0.099 (0.088,0.11)	< 0.001	0.333 (0.305,0.361)	< 0.001
NIE	0.011 (0.01,0.012)	< 0.001	0.009 (0.008,0.01)	< 0.001	0.004 (0.002,0.005)	< 0.001	0.019 (0.017,0.021)	< 0.001
NDE	0.307 (0.301,0.314)	< 0.001	0.186 (0.166,0.205)	< 0.001	0.096 (0.084,0.107)	< 0.001	0.314 (0.286,0.341)	< 0.001
PM	0.04 (0.04,0.041)	< 0.001	0.05 (0.048,0.051)	< 0.001	0.037 (0.036,0.038)	< 0.001	0.067 (0.065,0.069)	< 0.001
Chemotherapy								
TE	0.298 (0.292,0.305)	< 0.001	0.185 (0.165,0.204)	< 0.001	0.149 (0.138,0.161)	< 0.001	0.313 (0.286,0.341)	< 0.001
NIE	-0.009 (-0.009,-0.008)	< 0.001	-0.001 (-0.002,0)	0.016	0.054 (0.051,0.057)	< 0.001	0 (0,0)	0.051
NDE	0.307 (0.301,0.314)	< 0.001	0.186 (0.166,0.205)	< 0.001	0.096 (0.084,0.107)	< 0.001	0.314 (0.286,0.341)	< 0.001
PM	-0.034 (-0.035,-0.034)	< 0.001	-0.006 (-0.008,-0.004)	< 0.001	0.377 (0.376,0.379)	< 0.001	0 (-0.002,0.002)	0.707

Mediators	Colorectum		Melanoma		Bladder		Kidney	
	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P
Stage								
TEª	0.351 (0.329,0.372)	< 0.001	0.386 (0.336,0.437)	< 0.001	0.442 (0.398,0.486)	< 0.001	0.248 (0.204,0.292)	< 0.001
NIEb	0.033 (0.026,0.04)	< 0.001	0.039 (0.034,0.044)	< 0.001	0.029 (0.02,0.037)	< 0.001	0.019 (0.01,0.027)	< 0.001
NDEc	0.318 (0.298,0.338)	< 0.001	0.347 (0.297,0.397)	< 0.001	0.413 (0.37,0.456)	< 0.001	0.229 (0.186,0.273)	< 0.001
PM ^d	0.109 (0.107,0.112)	< 0.001	0.12 (0.117,0.124)	< 0.001	0.079 (0.075,0.082)	< 0.001	0.085 (0.081,0.09)	< 0.001
Surgery								
TE	0.244 (0.226,0.263)	< 0.001	0.338 (0.29,0.385)	< 0.001	0.389 (0.347,0.432)	< 0.001	0.159 (0.122,0.197)	< 0.001
NIE	0.02 (0.018,0.021)	< 0.001	0 (0,0.001)	0.269	0.003 (0.002,0.005)	< 0.001	0.011 (0.01,0.013)	< 0.001
NDE	0.225 (0.207,0.243)	< 0.001	0.337 (0.29,0.385)	< 0.001	0.386 (0.344,0.429)	< 0.001	0.148 (0.111,0.185)	< 0.001
PM	0.09 (0.088,0.091)	< 0.001	0.001 (-0.002,0.005)	0.387	0.009 (0.007,0.012)	< 0.001	0.076 (0.072,0.079)	< 0.001
Radiotherapy								
TE	0.223 (0.205,0.242)	< 0.001	0.338 (0.29,0.385)	< 0.001	0.395 (0.353,0.437)	< 0.001	0.148 (0.111,0.185)	< 0.001
NIE	-0.002 (-0.002,-0.001)	0.004	0 (0,0.001)	0.12	0.009 (0.006,0.011)	< 0.001	0 (0,0)	< 0.001
NDE	0.225 (0.207,0.243)	< 0.001	0.337 (0.29,0.385)	< 0.001	0.386 (0.344,0.429)	< 0.001	0.148 (0.111,0.185)	< 0.001
PM	-0.008 (-0.009,-0.006)	< 0.001	0.001 (-0.002,0.004)	0.474	0.026 (0.024,0.029)	< 0.001	0 (-0.004,0.003)	< 0.001
Chemotherapy								
TE	0.245 (0.227,0.264)	< 0.001	0.337 (0.29,0.385)	< 0.001	0.388 (0.345,0.43)	< 0.001	0.146 (0.108,0.183)	< 0.001
NIE	0.021 (0.018,0.023)	< 0.001	0 (0,0)	0.309	0.002 (0,0.003)	0.102	-0.002 (-0.003,-0.002)	< 0.001
NDE	0.225 (0.207,0.243)	< 0.001	0.337 (0.29,0.385)	< 0.001	0.386 (0.344,0.429)	< 0.001	0.148 (0.111,0.185)	< 0.001
PM	0.094 (0.092,0.095)	< 0.001	-0.001 (-0.004,0.003)	0.762	0.005 (0.002,0.008)	0.001	-0.015 (-0.019,-0.012)	< 0.001

Mediators	Endometrium	1	Pancreas	
	Effect size (95%CI)	P	Effect size (95%CI)	P
Stage				
TEa	0.274 (0.226,0.322)	< 0.001	0.166 (0.143,0.189)	< 0.001
NIEb	0.031 (0.026,0.037)	< 0.001	-0.006 (-0.015,0.004)	< 0.001
NDEc	0.243 (0.195,0.29)	< 0.001	0.172 (0.15,0.193)	< 0.001
PM ^d	0.129 (0.125,0.133)	< 0.001	-0.037 (-0.04,-0.033)	< 0.001
Surgery				
TE	0.173 (0.13,0.217)	< 0.001	0.153 (0.131,0.176)	< 0.001
NIE	0.014 (0.012,0.016)	< 0.001	0.042 (0.032,0.052)	< 0.001
NDE	0.16 (0.116,0.203)	< 0.001	0.111 (0.091,0.131)	< 0.001
PM	0.086 (0.082,0.09)	< 0.001	0.289 (0.286,0.293)	< 0.001
Radiotherapy				
TE	0.16 (0.116,0.203)	< 0.001	0.112 (0.091,0.132)	< 0.001
NIE	0 (-0.002,0.002)	< 0.001	0 (0,0.001)	0.392
NDE	0.16 (0.116,0.203)	< 0.001	0.111 (0.091,0.131)	< 0.001
PM	-0.001 (-0.005,0.003)	< 0.001	0.004 (0.002,0.006)	< 0.001
Chemotherapy				
TE	0.159 (0.115,0.203)	< 0.001	0.153 (0.132,0.173)	< 0.001
NIE	-0.001 (-0.001,0)	0.076	0.041 (0.037,0.045)	< 0.001
NDE	0.16 (0.116,0.203)	< 0.001	0.111 (0.091,0.131)	< 0.001
PM	-0.004 (-0.008,0)	0.059	0.286 (0.284,0.289)	< 0.001

^aTotal effect; ^bNatural indirect effect; ^cNatural direct effect; ^dProportion mediated

eTable 5. Mediation Analyses of Direct and Indirect Effects of Marital Status on Cancer-Specific Survival Mediated by Stage at Diagnosis and Treatment by

Histological Subtype

		Lı	ung	Pancreas					
Mediators	NSCLC ^e		SCLCf		Adenocarcinom	ıa	pNETg		
	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	
Stage									
TEª	0.191 (0.175,0.207)	< 0.001	0.107 (0.078,0.135)	< 0.001	0.156 (0.133,0.178)	< 0.001	0.359 (0.189,0.53)	< 0.001	
NIEb	-0.001 (-0.01,0.007)	0.774	-0.003 (-0.014,0.009)	0.621	-0.007 (-0.016,0.002)	0.147	0.052 (-0.024,0.127)	0.183	
NDEc	0.193 (0.179,0.206)	< 0.001	0.11 (0.083,0.136)	< 0.001	0.163 (0.142,0.183)	< 0.001	0.308 (0.155,0.461)	< 0.001	
PM ^d	-0.007 (-0.01,-0.004)	< 0.001	-0.029 (-0.033,-0.024)	< 0.001	-0.048 (-0.051,-0.044)	< 0.001	0.166 (0.14,0.192)	< 0.001	
Surgery									
TE	0.208 (0.193,0.222)	< 0.001	0.077 (0.052,0.102)	< 0.001	0.118 (0.096,0.139)	< 0.001	0.351 (0.189,0.514)	< 0.001	
NIE	0.108 (0.1,0.116)	< 0.001	0.007 (0.002,0.012)	0.004	0.034 (0.025,0.042)	< 0.001	0.059 (-0.005,0.122)	0.073	
NDE	0.1 (0.087,0.112)	< 0.001	0.07 (0.045,0.094)	< 0.001	0.084 (0.064,0.103)	< 0.001	0.293 (0.143,0.442)	< 0.001	
PM	0.546 (0.543,0.549)	< 0.001	0.096 (0.093,0.099)	< 0.001	0.299 (0.296,0.302)	< 0.001	0.192 (0.169,0.214)	< 0.001	
Radiotherapy									
TE	0.102 (0.089,0.114)	< 0.001	0.086 (0.06,0.111)	< 0.001	0.084 (0.065,0.104)	< 0.001	0.292 (0.143,0.441)	< 0.001	
NIE	0.002 (0.001,0.003)	< 0.001	0.016 (0.01,0.022)	< 0.001	0.001 (-0.001,0.002)	0.330	-0.001 (-0.003,0.002)	0.666	
NDE	0.1 (0.087,0.112)	< 0.001	0.07 (0.045,0.094)	< 0.001	0.084 (0.064,0.103)	< 0.001	0.293 (0.143,0.442)	< 0.001	
PM	0.02 (0.019,0.022)	< 0.001	0.19 (0.187,0.193)	< 0.001	0.007 (0.005,0.009)	0.036	-0.002 (-0.014,0.009))	0.669	
Chemotherapy									
TE	0.159 (0.146,0.171)	< 0.001	0.094 (0.069,0.119)	< 0.001	0.139 (0.119,0.159)	< 0.001	0.291 (0.141,0.44)	< 0.001	
NIE	0.059 (0.056,0.062)	< 0.001	0.024 (0.019,0.029)	< 0.001	0.055 (0.05,0.06)	< 0.001	-0.002 (-0.007,0.003)	0.496	
NDE	0.1 (0.087,0.112)	< 0.001	0.07 (0.045,0.094)	< 0.001	0.084 (0.064,0.103)	< 0.001	0.293 (0.143,0.442)	< 0.001	
PM	0.391 (0.389,0.392)	< 0.001	0.265 (0.262,0.268)	< 0.001	0.411 (0.409,0.413)	< 0.001	-0.007 (-0.018,0.004)	0.219	

^aTotal effect; ^bNatural indirect effect; ^cNatural direct effect; ^dProportion mediated; ^eNSCLC, non-small cell lung cancer; ^fSCLC, small cell lung cancer; ^gpNET, pancreatic neuroendocrine tumor.

				Br	east			
Mediators	Triple Negativ	'e	HER2 Enriche	d	Luminal A		Luminal B	
	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P
Stage								
TEª	0.163 (0.094,0.231)	< 0.001	0.272 (0.145,0.399)	0.002	0.181 (0.136,0.225)	< 0.001	0.232 (0.132,0.333)	< 0.001
NIEb	0.032 (0.02,0.044)	< 0.001	0.054 (0.032,0.075)	< 0.001	0.018 (0.014,0.022)	< 0.001	0.025 (0.013,0.036)	< 0.001
NDEc	0.131 (0.063,0.198)	< 0.001	0.219 (0.094,0.344)	0.008	0.163 (0.118,0.207)	< 0.001	0.208 (0.108,0.307)	< 0.001
PM^d	0.21 (0.202,0.217)	< 0.001	0.219 (0.207,0.231)	< 0.001	0.108 (0.103,0.112)	< 0.001	0.117 (0.108,0.126)	< 0.001
Surgery								
TE	0.091 (0.028,0.155)	< 0.001	0.169 (0.052,0.286)	0.004	0.107 (0.066,0.148)	< 0.001	0.098 (0.004,0.193)	0.040
NIE	0.005 (0.002,0.008)	< 0.001	0.012 (0.003,0.02)	0.007	0.003 (0.002,0.003)	< 0.001	0.009 (0.006,0.012)	< 0.001
NDE	0.086 (0.022,0.149)	0.007	0.158 (0.041,0.274)	0.008	0.104 (0.063,0.146)	< 0.001	0.089 (-0.005,0.183)	0.063
PM	0.06 (0.053,0.066)	< 0.001	0.074 (0.063,0.085)	< 0.001	0.026 (0.022,0.03)	< 0.001	0.1 (0.09,0.109)	0.136
Radiotherapy								
TE	0.089 (0.025,0.152)	0.006	0.157 (0.04,0.274)	0.008	0.106 (0.064,0.147)	< 0.001	0.09 (-0.004,0.184)	0.061
NIE	0.003 (0,0.006)	0.039	-0.001 (-0.002,0.001)	0.592	0.001 (0,0.002)	0.003	0.001 (-0.001,0.002)	0.437
NDE	0.086 (0.022,0.149)	< 0.001	0.158 (0.041,0.274)	0.008	0.104 (0.063,0.146)	< 0.001	0.089 (-0.005,0.183)	0.063
PM	0.033 (0.027,0.04)	0.008	-0.003 (-0.014,0.007)	0.529	0.012 (0.007,0.016)	0.745	0.008 (-0.002,0.018)	0.101
Chemotherapy								
TE	0.087 (0.023,0.15)	0.007	0.157 (0.04,0.274)	0.009	0.103 (0.061,0.144)	< 0.001	0.096 (0.001,0.19)	0.047
NIE	0.001 (0,0.002)	0.177	-0.001 (-0.006,0.005)	0.822	-0.002 (-0.003,-0.001)	0.002	0.007 (0.002,0.011)	0.005
NDE	0.086 (0.022,0.149)	0.008	0.158 (0.041,0.274)	0.008	0.104 (0.063,0.146)	< 0.001	0.089 (-0.005,0.183)	0.063
PM	0.01 (0.003,0.016)	0.003	-0.004 (-0.015,0.007)	0.449	-0.019 (-0.023,-0.015)	< 0.001	0.072 (0.062,0.081)	< 0.001

eTable 6. Mediation Analyses of Direct and Indirect Effects of Marital Status on Cancer-Specific Survival Mediated by Stage at Diagnosis and Treatment by Sex

		Total S	Sample			Lu	ing			Color	ectum	
N	Female)	Male		Femal	e	Male		Femal	e	Male	
Mediators	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P
Stage												
TE ^a	0.261 (0.248,0.274)	<0.001	0.372 (0.358,0.386)	<0.001	0.129 (0.107,0.15)	<0.001	0.21 (0.19,0.229)	<0.001	0.261 (0.231,0.291)	<0.001	0.405 (0.377,0.434)	<0.001
NIEb	0.048 (0.045,0.051)	<0.001	0.07 (0.064,0.075)	<0.001	-0.01 (-0.022,0.002)	0.001	-0.001 (- 0.012,0.01)	<0.001	0.061 (0.05,0.072)	<0.001	0.035 (0.025,0.044)	<0.001
NDEc	0.213 (0.201,0.226)	<0.001	0.302 (0.29,0.315)	<0.001	0.139 (0.121,0.157)	<0.001	0.211 (0.195,0.227)	<0.001	0.2 (0.172,0.228)	<0.001	0.371 (0.343,0.398)	<0.001
PM^d	0.203 (0.202,0.204)	<0.001	0.217 (0.215,0.219)	<0.001	-0.084 (-0.089, - 0.08)	<0.001	-0.006 (-0.01,-0.002)	<0.001	0.257 (0.253,0.261)	<0.001	0.103 (0.099,0.106)	<0.001
Surgery												
TE	0.203 (0.194,0.213)	<0.001	0.308 (0.297,0.318)	<0.001	0.161 (0.141,0.182)	<0.001	0.211 (0.194,0.229)	<0.001	0.21 (0.182,0.239)	<0.001	0.278 (0.254,0.303)	<0.001
NIE	0.021 (0.019,0.022)	<0.001	0.073 (0.069,0.077)	<0.001	0.085 (0.073,0.097)	<0.001	0.096 (0.088,0.105)	<0.001	0.01 (0.008,0.013)	<0.001	0.029 (0.026,0.032)	<0.001
NDE	0.183 (0.173,0.192)	<0.001	0.235 (0.225,0.244)	<0.001	0.076 (0.059,0.093)	<0.001	0.115 (0.1,0.13)	<0.001	0.2 (0.172,0.228)	<0.001	0.249 (0.225,0.274)	< 0.001
PM	0.111 (0.11,0.112)	<0.001	0.266 (0.264,0.267)	<0.001	0.547 (0.544,0.551)	<0.001	0.483 (0.48,0.485)	<0.001	0.054 (0.051,0.056)	<0.001	0.117 (0.115,0.119)	<0.001

		Total S	Sample			Lu	ing			Color	ectum	
Mediators	Female	2	Male		Femal	e	Male		Femal	e	Male	
Mediators	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P
Radiotherapy	,											
TE	0.193 (0.184,0.203)	<0.001	0.233 (0.223,0.242)	<0.001	0.079 (0.062,0.096)	<0.001	0.119 (0.103,0.134)	<0.001	0.199 (0.171,0.227)	<0.001	0.247 (0.223,0.272)	<0.001
NIE	0.01 (0.01,0.011)	<0.001	-0.002 (- 0.003,-0.002)	<0.001	0.003 (0.001,0.004)	<0.001	0.004 (0.002,0.005)	<0.001	-0.001 (-0.002,0)	<0.001	-0.002 (-0.003,- 0.001)	<0.001
NDE	0.183 (0.173,0.192)	<0.001	0.235 (0.225,0.244)	<0.001	0.076 (0.059,0.093)	<0.001	0.115 (0.1,0.13)	<0.001	0.2 (0.172,0.228)	<0.001	0.249 (0.225,0.274)	<0.001
PM	0.059 (0.058,0.06)	<0.001	-0.01 (-0.011,- 0.009)	<0.001	0.038 (0.037,0.04)	<0.001	0.033 (0.032,0.035)	<0.001	-0.006 (-0.008,- 0.003)	<0.001	-0.01 (-0.012,- 0.008)	<0.001
Chemotherap	y		ı									
TE	0.192 (0.182,0.201)	<0.001	0.25 (0.24,0.26)	<0.001	0.117 (0.1,0.134)	<0.001	0.178 (0.163,0.194)	<0.001	0.219 (0.191,0.248)	<0.001	0.269 (0.245,0.294)	<0.001
NIE	0.009 (0.008,0.01)	<0.001	0.015 (0.014,0.016)	<0.001	0.041 (0.037,0.045)	<0.001	0.063 (0.059,0.067)	<0.001	0.019 (0.016,0.022)	<0.001	0.02 (0.017,0.023)	<0.001
NDE	0.183 (0.173,0.192)	<0.001	0.235 (0.225,0.244)	<0.001	0.076 (0.059,0.093)	<0.001	0.115 (0.1,0.13)	<0.001	0.2 (0.172,0.228)	<0.001	0.249 (0.225,0.274)	<0.001
PM	0.051 (0.05,0.052)	<0.001	0.068 (0.067,0.068)	<0.001	0.363 (0.361,0.365)	<0.001	0.375 (0.373,0.377)	<0.001	0.096 (0.093,0.099)	<0.001	0.083 (0.081,0.086)	<0.001

		Mela	noma			Bla	dder		Kidney			
N. T	Female	.	Male		Female	e	Male	;	Femal	le	Male	
Mediators	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P
Stage												
TE ^a	0.313 (0.224,0.402)	<0.001	0.409 (0.347,0.471)	<0.001	0.373 (0.281,0.464)	<0.001	0.469 (0.419,0.518)	<0.001	0.117 (0.042,0.192	<0.001	0.294 (0.24,0.349)	<0.001
NIEb	0.023 (0.017,0.029)	<0.001	0.05 (0.042,0.057)	<0.001	0.009 (-0.009,0.028)	0.281	0.038 (0.029,0.048)	<0.001	0.006 (- 0.005,0.018)	<0.001	0.025 (0.013,0.036)	0.115
NDEc	0.29 (0.201,0.379)	<0.001	0.359 (0.298,0.421)	<0.001	0.363 (0.274,0.453)	<0.001	0.431 (0.382,0.479)		0.111 (0.036,0.185	<0.001	0.27 (0.216,0.323)	<0.001
PM^d	0.084 (0.077,0.091)	<0.001	0.144 (0.139,0.149)	<0.001	0.03 (0.022,0.039)	<0.001	0.1 (0.096,0.105)	<0.001	0.057 (0.049,0.065)	<0.001	0.095 (0.09,0.101)	<0.001
Surgery												
TE	0.299 (0.215,0.383)	<0.001	0.35 (0.292,0.408)	<0.001	0.341 (0.253,0.429)	<0.001	0.405 (0.358,0.453)	<0.001	0.173 (0.13,0.217)	<0.001	0.177 (0.13,0.224)	<0.001
NIE	0 (-0.001,0.001)	0.604	0.001 (0,0.002)	0.103	0.003 (-0.001,0.006)	0.147	0.003 (0.001,0.005)	<0.001	0.014 (0.012,0.016)	<0.001	0.014 (0.012,0.017)	<0.001
NDE	0.299 (0.215,0.383)	<0.001	0.349 (0.291,0.407)	<0.001	0.338 (0.25,0.426)	<0.001	0.402 (0.354,0.45)	<0.001	0.16 (0.116,0.203)	<0.001	0.162 (0.116,0.209)	<0.001
PM	-0.001 (-0.007,0.005)	0.736	0.003 (-0.001,0.007)	0.136	0.009 (0.003,0.016)	0.003	0.009 (0.006,0.012)	<0.001	0.086 (0.082,0.09)	<0.001	0.088 (0.084,0.093)	<0.001

		Mela	noma			Bla	ıdder			Kie	dney	
3.6 11 4	Female	2	Male		Femal	e	Male	,	Fema	le	Male	
Mediators	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P
Radiotherapy	7											
TE	0.299 (0.215,0.383)	<0.001	0.349 (0.291,0.408)	<0.001	0.347 (0.259,0.435)	<0.001	0.411 (0.363,0.459)	<0.001	0.16 (0.116,0.203	<0.001	0.162 (0.116,0.209)	<0.001
NIE	0 (-0.002,0.002)	0.81	0 (0,0.001)	0.296	0.009 (0.004,0.014)	<0.001	0.009 (0.006,0.011)	<0.001	0 (- 0.002,0.002)	<0.001	0 (0,0)	0.487
NDE	0.299 (0.215,0.383)	<0.001	0.349 (0.291,0.407)	<0.001	0.338 (0.25,0.426)	<0.001	0.402 (0.354,0.45)	<0.001	0.16 (0.116,0.203	<0.001	0.162 (0.116,0.209)	<0.001
PM	0.001 (-0.005,0.007)	0.745	0.001 (-0.003,0.005)	0.528	0.031 (0.024,0.037)	<0.001	0.026 (0.023,0.029)	<0.001	-0.001 (- 0.005,0.003)	<0.001	-0.001 (-0.005,0.003)	0.695
Chemotherap	ру					1			1	1		
TE	0.299 (0.215,0.383)	<0.001	0.349 (0.291,0.407)	<0.001	0.343 (0.255,0.431)	<0.001	0.403 (0.355,0.451)	<0.001	0.159 (0.115,0.203	<0.001	0.16 (0.113,0.206)	<0.001
NIE	0 (-0.001,0)	0.7135	0 (0,0)	0.657	0.005 (0,0.01)	0.047	0.001 (-0.001,0.003)	0.412	-0.001 (-0.001,0)	0.076	-0.003 (-0.003,- 0.002)	<0.001
NDE	0.299 (0.215,0.383)	<0.001	0.349 (0.291,0.407)	<0.001	0.338 (0.25,0.426)	<0.001	0.402 (0.354,0.45)	<0.001	0.16 (0.116,0.203	<0.001	0.162 (0.116,0.209)	<0.001
PM	-0.001 (-0.007,0.005)	0.699	0 (-0.004,0.004)	0.88	0.017 (0.01,0.023)	<0.001	0.002 (-0.001,0.005)	0.139	-0.004 (-0.008,0)	0.059	-0.018 (-0.022,- 0.014)	<0.001

		Pano	creas		Breast					
Mallatan	Female	:	Male		Femal	e	Male	;		
Mediators	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P		
Stage										
TEª	0.106 (0.074,0.138)	<0.001	0.213 (0.179,0.247)	<0.001	0.271 (0.25,0.293)	< 0.001	0.546 (0.344,0.748)	0.003		
NIEb	-0.012 (- 0.025,0)	<0.001	-0.002 (- 0.016,0.012)	<0.001	0.028 (0.026,0.03)	<0.001	0.032 (0,0.064)	0.277		
NDE°	0.119 (0.089,0.148)	<0.001	0.215 (0.184,0.246)	<0.001	0.244 (0.223,0.265)	<0.001	0.514 (0.314,0.713)	0.005		
PM^d	-0.123 (-0.128, -0.118)	<0.001	-0.011 (- 0.016,-0.006)	<0.001	0.115 (0.113,0.117)	<0.001	0.076 (0.061,0.09)	<0.001		
Surgery										
TE	0.109 (0.078,0.141)	<0.001	0.188 (0.156,0.221)	<0.001	0.103 (0.07,0.135)	<0.001	0.567 (0.175,0.959)	0.005		
NIE	0.035 (0.021,0.049)	<0.001	0.042 (0.029,0.056)	<0.001	0.004 (0.003,0.005)	<0.001	0 (-0.006,0.007)	0.899		
NDE	0.074 (0.046,0.102)	<0.001	0.146 (0.116,0.175)	<0.001	0.098 (0.066,0.131)	<0.001	0.567 (0.175,0.958)	0.005		
PM	0.331 (0.326,0.336)	<0.001	0.242 (0.238,0.247)	<0.001	0.042 (0.038,0.045)	< 0.001	0.001 (- 0.017,0.019)	0.914		

		Pan	creas		Breast					
3.6 11 4	Female	,	Male		Femal	e	Male			
Mediators	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P	Effect size (95%CI)	P		
Radiotherapy	7									
TE	0.074 (0.046,0.102)	<0.001	0.147 (0.117,0.176)	<0.001	0.1 (0.067,0.132)	<0.001	0.567 (0.175,0.958)	0.005		
NIE	0 (-0.001,0.001)	0.71	0.001 (0,0.002)	0.109	0.001 (0.001,0.002)	0.06731 2613	0 (- 0.001,0.001)	0.995		
NDE	0.074 (0.046,0.102)	<0.001	0.146 (0.116,0.175)	<0.001	0.098 (0.066,0.131)	<0.001	0.567 (0.175,0.958)	0.005		
PM	-0.003 (-0.006,0)	0.036	0.008 (0.005,0.011)	<0.001	0.014 (0.011,0.018)	<0.001	0 (- 0.018,0.018)	0.999		
Chemotherap	ру									
TE	0.112 (0.083,0.14)	<0.001	0.191 (0.161,0.22)	<0.001	0.098 (0.065,0.13)	<0.001	0.576 (0.183,0.969)	0.004		
NIE	0.037 (0.032,0.043)	<0.001	0.045 (0.039,0.051)	<0.001	-0.001 (- 0.002,0)	<0.001	0.009 (- 0.026,0.045)	0.604		
NDE	0.074 (0.046,0.102)	<0.001	0.146 (0.116,0.175)	<0.001	0.098 (0.066,0.131)	<0.001	0.567 (0.175,0.958)	0.005		
PM	0.345 (0.342,0.348)	<0.001	0.252 (0.249,0.256)	<0.001	-0.009 (- 0.013,-0.006)	<0.001	0.021 (0,0.043)	0.054		

^aTotal effect; ^bNatural indirect effect; ^cNatural direct effect; ^dProportion mediated

eTable 7. Demographic and Clinical Characteristics of Individuals Included and Excluded

	Included	Excluded	
Characteristics	1733906 (81.3)	397632 (18.7)	P
Age at diagnosis ^a	63.76 (12.60)	67.79 (14.90)	< 0.001
Race ^b			< 0.001
White	1393206 (80.4)	304445 (76.6)	
Black	201295 (11.6)	42183 (10.6)	
Other	127420 (7.3)	23254 (5.8)	
Unknown	11985 (0.7)	27750 (7.0)	
Sex ^b			< 0.001
Female	894379 (51.6)	163721 (41.2)	
Male	839527 (48.4)	233911 (58.8)	
Grade ^b			< 0.001
Grade I	206578 (11.9)	34179 (8.6)	
Grade II	612491 (35.3)	92795 (23.3)	
Grade III	488671 (28.2)	67334 (16.9)	
Grade IV	60788 (3.5)	7827 (2.0)	
Unknown	365378 (21.1)	195497 (49.2)	
Surgery ^b			< 0.001
No	536914 (31.0)	182464 (45.9)	
Yes	1196992 (69.0)	154303 (38.8)	
Unknown	0 (0.0)	60865 (15.3)	
Radiation ^b			< 0.001
No/ Unknown	1159441 (66.9)	341790 (86.0)	
Yes	574465 (33.1)	55842 (14.0)	
Chemotherapy ^b			< 0.001
No/ Unknown	1207743 (69.7)	350252 (88.1)	
Yes	526163 (30.3)	47380 (11.9)	

Characteristics	Included 1733906 (81.3)	Excluded 397632 (18.7)	P
Stage ^b			< 0.001
Early	1189957 (68.6)	193049 (48.5)	
Advanced	543949 (31.4)	48769 (12.3)	
Unknown	0 (0.0)	155814 (39.2)	
Insurance ^b			< 0.001
Insured	1475410 (85.1)	199736 (50.2)	
Medicaid	181047 (10.4)	27051 (6.8)	
Uninsured	39962 (2.3)	6337 (1.6)	
Unknown	37487 (2.2)	164508 (41.4)	
Povertya	32.40 (9.55)	32.47 (9.62)	< 0.001
Education ^a	14.87 (6.17)	14.76 (6.15)	< 0.001
Residence ^b			< 0.001
Metropolitan	1531155 (88.3)	350231 (88.1)	
Nonmetropolitan	201258 (11.6)	46226 (11.6)	
Unknown	1493 (0.1)	1175 (0.3)	

Unknown
 1493 (0.1)
 1175 (0.3)

 aMean (SD); bN (%) includes American Indian, Alaska Native, and Asian/Pacific Islander