

Supplemental Table 1. Risk of second primary malignancies (SPM) among pediatric lymphoma patients by specific lymphoma subtypes.

Lymphoma subtype	No. of patients	Observed no. of SPM	Expected no. of SPM	SIR	95%CI	Excess risk (per 10,000)	PY at Risk	Mean PY at Risk	Mean Age at Exposure	Mean Age at SPM
Total	6984	478	109.7	4.36#	3.98-4.77	32.9	112058	16.0	14.2	35.1
1 Hodgkin Lymphoma	3983	371	78.2	4.74#	4.27-5.25	42.3	69265	17.4	15.8	36.3
1(a) Classical Hodgkin lymphoma	3828	360	77.4	4.65#	4.19-5.16	41.8	67668	17.7	15.9	36.4
1(a)1 Lymphocyte-rich/mixed cell/lymphocyte depleted	599	57	14.0	4.08#	3.09-5.29	34.2	12575	21.0	14.6	39.1
1(a)1.1 Lymphocyte-rich	144	13	3.3	3.94#	2.10-6.74	30.9	3138	21.8	14.2	34.0
1(a)1.2 Mixed cellularity	429	39	9.9	3.93#	2.80-5.38	32.9	8838	20.6	14.7	39.2
1(a)1.3 Lymphocyte-depleted	26	5	0.8	6.69#	2.16-15.61	70.9	600	23.1	15.6	51.6
1(a)2 Nodular sclerosis	2826	270	57.5	4.69#	4.15-5.29	42.8	49674	17.6	16.1	36.2
1(a)3 Classical Hodgkin lymphoma, NOS	403	33	5.9	5.63#	3.87-7.91	50.1	5419	13.5	15.8	33.3
1(b) Nodular lymphocyte predominant Hodgkin lymphoma	155	11	0.9	13.00#	6.48-23.27	63.6	1598	10.3	14.1	35.0
2 Non-Hodgkin lymphoma	2995	107	31.5	3.48#	2.83-4.23	17.7	42781	14.1	12.1	30.7
2(a) Non-Hodgkin lymphoma, B-cell	1880	68	22.9	2.97#	2.31-3.77	15.7	28793	15.3	12.5	33.0
2(a)1 Precursor Non-Hodgkin lymphoma, B-cell	119	2	0.2	11.17#	1.25-40.32	25.3	720	6.1	8.5	9.2
2(a)2 Mature Non-Hodgkin lymphoma, B-cell	1461	50	17.6	2.84#	2.11-3.75	14.4	22447	15.4	12.9	33.2
2(a)2.1 Mantle cell lymphoma	24	4	0.6	6.75#	1.82-17.29	62.5	545	22.7	9.9	52.4
2(a)2.2 Diffuse large B-cell lymphoma (DLBCL)	711	29	10.7	2.72#	1.82-3.91	16.3	11252	15.8	14.7	30.3
2(a)2.3 Burkitt lymphoma/leukemia	568	12	4.6	2.63#	1.36-4.60	8.7	8571	15.1	10.2	31.0
2(a)2.4 Follicular lymphoma	105	5	1.5	3.36#	1.08-7.85	21.8	1611	15.4	14.5	40.4
2(a)3 Non-Hodgkin lymphoma, B-cell, NOS	300	16	5.1	3.14#	1.80-5.11	19.4	5626	18.8	11.9	35.1

2(b) Non-Hodgkin lymphoma, T-cell	623	15	2.3	6.44#	3.60-10.62	23.4	5414	8.7	12.0	23.8
2(b)1 Precursor Non-Hodgkin lymphoma, T-cell	199	3	0.3	9.10#	1.83-26.59	22.0	1213	6.1	10.4	16.0
2(b)2 Mature Non-Hodgkin lymphoma, T-cell	424	12	2.0	6.00#	3.10-10.48	23.8	4201	9.9	12.8	25.7
2(b)2.1 Mycosis fungoides	94	5	0.6	9.08#	2.93-21.19	48.4	919	9.8	13.5	28.5
2(b)2.2 Peripheral T-cell lymphoma	324	7	1.4	4.86#	1.95-10.00	17.1	3252	10.0	12.6	23.8
Peripheral T-cell lymphoma, NOS	66	3	0.4	6.91#	1.39-20.19	31.5	813	12.3	11.3	25.9
Anaplastic lar cell lymph, T-/Null-cell	185	3	0.8	3.79	0.76-11.09	11.8	1880	10.2	12.5	22.2
Cutaneous T-cell lymphoma, NOS	27	1	0.1	10.24	0.13-56.97	35.7	252	9.4	14.1	22.3
2(c) Non-Hodgkin lymphoma, unknown lineage	330	16	3.3	4.90#	2.80-7.96	22.6	5647	17.1	10.2	27.6
2(d) Non-Hodgkin lymphoma, NOS	162	8	3.0	2.64#	1.14-5.20	17.0	2927	18.1	12.6	30.0

SPM, second primary malignancy. SIR, standardized incidence ratio. PY, person-year. NOS, not otherwise specified.

#Significant difference was observed ($P < 0.05$).

All results in this table were calculated and analyzed by MP-SIR session of the software SEER*Stat with a 1-year latency of SPM after the first primary lymphoma.

Supplemental Table 2. Multivariate competing-risk analysis of second primary malignancies (SPM) among pediatric lymphoma patients for demographic characteristics.

Characteristic	HR	95%CI		P value
		Lower	Upper	
Sex				
Male	Reference	-	-	
Female	2.51	2.08	3.04	<0.001
Age, years				
<1	1.86	0.21	16.60	0.58
1-4	Reference	-	-	-
5-9	0.88	0.45	1.72	0.72
10-14	1.35	0.73	2.48	0.34
15-19	1.54	0.85	2.81	0.16
Race				
White	Reference	-	-	-
Black	1.21	0.89	1.65	0.23
Other†	1.25	0.81	1.92	0.32
Unspecified	NA	NA	NA	NA
Lymphoma subtype				
Hodgkin lymphoma	Reference	-	-	-
Non-Hodgkin lymphoma	0.66	0.51	0.86	0.002
Ann Arbor stage				
Stage I	Reference	-	-	-
Stage II	0.78	0.57	1.07	0.12
Stage III	0.92	0.62	1.36	0.68
Stage IV	0.94	0.63	1.38	0.74
Unstaged	NA	NA	NA	NA
Chemotherapy				
No	Reference	-	-	
Yes	0.94	0.77	1.16	0.58
Radiotherapy				
No	Reference	-	-	-
Yes	1.42	1.15	1.74	0.001
Unspecified	NA	NA	NA	NA

†Other in race correspond to American Indian/AK Native, Asian/Pacific Islander. The results in this table were calculated and analyzed by R software with package “cmprsk” and “survival”.

Supplemental Figure 1. Cumulative incidence of SPM by multivariate competing-risk analysis among pediatric lymphoma patients distributed by year of diagnosis. A. Follow up of 10 years; B. Follow up of 20 years; C. Follow up of 30 years

