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Thyroid dysfunction in children after hematopoietic stem cell transplantation : short term follow-up for 12months

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Introduction

• Hematopoietic stem cell transplantation (HSCT) is a potentially curative therapeutic modality for patients with hematological malignancies.

• Endocrine dysfunctions are well-known complication after HSCT ; thyroid dysfunction and gonadal failure are most commonly observed adverse effect among the endocrinopathies.

Objective

To evaluate 12 months follow-up of thyroid function in patients who underwent HSCT during childhood and adolescence of hematological malignancy.
To evaluate the risk factors to predict thyroid dysfunction at 12 months.

Figure 2 shows that the percentage distribution of thyroid dysfunction after HSCT over 12 months. forty-four patients (53.0%) had thyroid dysfunction during follow up.
Significantly positive association between overall thyroid dysfunction incidence and time period after HSCT was observed (P for trend 0.035). among the thyroid dysfunction, subclinical hypothyroidism (SH) incidence increased significantly (P for trend 0.0015) and euthyroid sick syndrome (ESS) incidence decreased significantly (P for trend 0.031) (Figure 3).

• Figure 4 shows that the changes in thyroid hormone levels after HSCT during the 12 months follow-up.

• In univariate analysis, there was no significant risk factor of thyroid dysfunction at 12 months after HSCT (Table 3).

Methods

• This study is a retrospective chart review of 217 hematologic-malignancy patients who underwent HSCT between January 2006 and December 2011. Patients enrollement is illustrated in Figure 1.

• Thyroid function of the patients was evaluated before and 1, 3, 6, 9, and 12 months after HSCT.

Total **217** pediatric patients undergone HSCT at Seoul St. Mary's hospital From January 2006 to December 2011

Exclusion
 Solid tumor : 18 patients
 Non-malignant disease : 52 patients
 Known thyroid dysfunction : 23 patients
 Incomplete records : 41 patients

Eighty-three patients (46 males and 37 females) diagnosed as hematologic malignancy were included

Figure 1. Study scheme.

• There were 83 patients who met the inclusion criteria. The patient



Figure 2. Percentage distribution of thyroid dysfunction incidence after HSCT. (A) Overall distribution of thyroid dysfunction incidence after HSCT. (B) Distribution of thyroid dysfunction incidence in each patients.



characteristics and treatment variable are presented in Table 1 and thyroid dysfunction after HSCT for 1 years who were studied are showed in Table 2.

Table 1. Patient characteristics and treatment variables

	Number (n =83)	
Age at HSCT (years)	9.78 ± 4.42	
Sex Male/ female	46/37	
Disease and disease status		
ALL	25	
First CR	16	
Second CR	9	
AML	51	
First CR	49	
Second CR	2	
CML-First chronic phase	7	
Type of HSCT		
Sibling / Unrelated	21/62	
Cell source		
BM/PBSC/Cord	10 /72/ 1	
Acute GVHD		
0~I	53	
II~IV	30	
Chronic GVHD (Yes/No)		
None	34	
Clinical limited	11	
Clinical extensive	38	
Death	6	

	No. (%)
After 1 month	14/81 (7.3%)
normal	67
euthyroid sick syndrome	8
subclinical hyperthyroidism	2
hypothyroxemia	4
After 3 months	10/83 (12.0%)
normal	73
euthyroid sick syndrome	5
subclinical hypothyroidism	2
subclinical hyperthyroidism	2
overt hyperthyroidism	1
After 6 months	14/81 (17.3%)
normal	67
euthyroid sick syndrome	3
subclinical hypothyroidism	7
subclinical hyperthyroidism	2
hypothyroxemia	2
After 9 months	21/82 (25.6%)
normal	61
euthyroid sick syndrome	3
subclinical hypothyroidism	13
hypothyroxemia	4
overt hypothyroidism	1
After 12 months	20/83 (24.1%)
normal	63
euthyroid sick syndrome	2
subclinical hypothyroidism	13
hypothyroxemia	2
overt hypothyroidism	2
overt hyperthyroidism	1

months months

Figure 3. Frequency of thyroid dysfunction (A) and SH (B) and ESS (C) according to time.



Figure 4. Changes in T3 (A) and fT4 (B) and TSH (C) levels over 12 months after HSCT.

Table 3. Risk factors associated with thyroid dysfunction at 12 months after hematopoietic stem cell transplantation

		Univariate logistic regression analysis		
		OR (95% CI)	P - value	
Age at HSCT		0.913(0.813-1.025)	0.123	
Sex	Male (n=46)	1		
	Female (n=37)	1.023(0.372-2.812)	0.965	
Diagnosis	Lymphoid (n=25)	1		
	Myeloid (n=58)	1.008(0.336-3.019)	0.989	
Conditioning regimen	BU based (n=56)	1		
	TBI based (n=27)	0.857(0.288-2.550)	0.782	
aGVHD grade	0~I (n=53)	1		
	II~IV (n=30)	0.507(0.164-1.570)	0.239	
cGVHD grade	0~Limited (n=45)	1		
	Extensive (n=38)	0.556(0.196-1.578)	0.270	

Conclusion

After HSCT during childhood and adolescence, a significant number of patients experience thyroid dysfunction including ESS and SH. Short-term and continuous follow-up for thyroid function after HSCT is important to provide timely and appropriate treatment.