be determined. 16.17 Although there is no agreement on the boundary of LND, previous studies demonstrated that 10% of single metastasis involved the common iliac nodes¹⁷ and nodal metastases in patients having multiple nodal metastases were found in common iliac and extrapelvic nodes such as paracavel, aortocaval and pataaortic nodes.¹⁸ Since preoperative assessment of nodal status is difficult even using modern imaging modalities, complete LND is necessary for accurate nodal staging of bladder cancer. In addition, the most effective therapy for metastatic nodes might be complete surgical dissection. it is mandatory to remove as many pelvic lymph nodes as possible for accurate staging, prognosis and radical treatment.

Conclusions

Removal of 13 or more lymph nodes has a significant impact on the disease-specific survival of node-positive patients. Adequate PLND provides accurate nodal staging and important pathological information for prognosis in node-positive patients.

References

- Bassi P, Ferrante GD, Piazza N, et al: Prognostic factors of outcome after radicalcystectomy for bladder cancer: Aretrospective study of a homogeneous patient cohort. J Urol 161:1494-1497, 1999.
- 2. Dalbagni G, Genega E, Hashibe M, *et al*: Cystectomy for bladder cancer: A contemporary series. J Urol **165**: 1111-1116, 2001.
- 3. Stein JP, Lieskovsky G, Cote R, *et al*: Radical cystectomy in the treatment of invasive bladder cancer: long-term results n 1,054 patients. J Clin Oncol **19**: 666-675, 2001.
- 4. Poulsen AL, Horn H and Steven K: Radical cystectomy: Extending the limits of pelvic lymph node dissection improves survival for patients with bladder cancer confined to the bladder wall. J Urol 160: 2015-2020, 1998.
- 5. Mills RD, Turner WH, Fleischmann A, *et al*: Pelvic lymph node metastases from bladder cancer: Outcome in 83 patients after radical cystectomy and pelvic

- lymphadenectomy. J Urol **166:** 19-23, 2001.
- 6. Whitmore WF Jr: Management of invasive bladder neoplasms. Sem Urol 1: 34-41, 1983.
- 7. Sobin LH and Wittekind Ch: *TNM Classification of Malignant tumors*, 5th ed. New York, Wiley-Liss Inc, 1997.
- 8. Mostofi FK, Davis CJ and Sesterhenn IA: Histological typing of urinary bladder tumours, 2nd ed. New York, Springer, 1999.
- 9. Lerner SP, Skinner DG, Lieskovsky G, *et al*: The rationale for en bloc pelvic lymph node dissection for bladder cancer patients with nodal metastasis: long-term results. J Urol **149**: 758-765, 1993.
- 10. Frank I, Cheville JC, Blute ML *et al*: Transitional cell carcinoma of the urinary bladder with regional lymph node involvement treated by cystectomy. Cancer **97**: 2425-2431, 2003.
- Stein JP. Cai J, Groshen S, Skinner DG: 11. Risk factors for patients with pelvic lymph node metastases following radical bloc pelvic cystectomy with en lymphadenectomy: the ssssconcept of lymph node density. J Urol 170: 35-41, 2003.
- 12. Herr HW, Bochner BH, Dalbagni G, et al: Impact of the number of lymph nodes retrieved on outcome in patients with muscle invasive bladder cancer. J Urol 167: 1295-1298, 2002.
- 13. Herr HW, Faulkner JR, Grossman B, *et al*: Surgical factors influence bladder cancer outcomes: A cooperative group report. J Clin Oncol **22**: 2781-2789, 2004.
- 14. Leissner J, Hohenfellner R, Thuroff JW, *et al*: Lymphadenectomy in patients with transitional cell carcinoma of the bladder; significance for staging and prognosis. BJU Int **85**: 817-823, 2000.
- 15. Konety BR, Joslyn SA and O'Donnell M: Extent of pelvic lymphadenectomy and its impact on outcome in patients diagnosed with bladder cancer: analysis of data from the surveillance, epidemiology and end results program data base. J Urol 169: 946-950, 2003.
- 16. Thalmann GN, Fleichmann A, Mills RD, *et al*: Lymphadenectomy in bladder cancer.

- EAU Update Series 1: 100-107, 2003.

 17. Leissner J, Ghoneim MA, Abol-Enein H, et al: Extended radical lymphadenectomy in patients with urothelial bladder cancer: results of a prospective multicenter study. J Urol 171: 139-144, 2004.
- 18. Abol-Enein H, El-Baz M, Abd El-Hameed MA, *et al*: Lymph node involvement in patients with bladder cancer treated with radical cystectomy: a patho-anatomical study--a single center experience. J Urol **172**: 1818-1821, 2004.

Table 1 Clinical and pathological stages and tumor histology

	No. of pts (%)
Clinical stage	
T1/Tis	5 (3.4)
T2	71 (48.6)
T3	43 (29.5)
T4	27 (18.5)
Pathological stage	
T0	19 (13.0)
T1/Tis	30 (20.6)
T2	41 (28.1)
T3	37 (25.3)
T4	19 (13.0)
Histology	
UC*	112 (76.7)
UC + other histological component	28 (19.2)
Other histological component	6 (4.1)

^{*} UC; urothelial carcinoma

 Table 2
 Multivariate analysis of parameters predicting disease-specific survival in node-positive patients

node-positive patients			
parameter	Multivariate p-value	Hazard ratio	95% CI
histology (pure UC* vs. other histological component ± UC)	0.6897	1.279	0.382 – 4.279
grade (G1/G2 vs. G3)	0.4450	1.936	0.355 - 10.550
pathological stage (= T2 vs. T3 =)	0.0132	8.205	1.553 – 43.343
No. of removed nodes (= 13 vs. < 13)	0.0008	9.363	2.526 – 34.704
No. of positive nodes (< 4 vs. 4 =)	0.0115	4.944	1.431 – 17.085
adjuvant chemotherapy (with vs. without)	0.6164	1.344	0.423 – 4.274

^{*} UC; urothelial carcinoma

Figure legends

Figure 1 Distribution of node-negative and node-positive patients stratified by the number of nodes removed and cumulative percentage of node-positive patients according to the number of nodes removed. White bars; the number of node-negative patients, black bars; the number of node-positive patients, and plots and line; the cumulative percentage of node-positive patients according to the number of lymph nodes removed.

Figure 2 Disease-specific survival of 146 patients stratified by nodal status. Node-positive (n=25) vs. node negative (n=121) patients, p < 0.0001 by log-rank test.

Figure 3 Disease-specific survival according to the number of nodes removed. A; Patients with pathologically organ confined tumor and negative nodes, the number of node removed of = 13 (n=46) vs. < 13 (n=44), p = 0.604. B; Patients with pathologically extravesical tumor and negative nodes, = 13 (n=28) vs. < 13 (n=28) nodes removed, p = 0.113. C; Node-positive patients, = 13 (n=14) vs. < 13 (n=11) nodes removed, p = 0.002. All p-values were determined by log-rank test.

Figure 4 Disease-specific survival of node-positive patients. A; According to the number of positive nodes; = 4 (n=7) vs. < 4 (n=18), p = 0.003. B; According to the combination of the numbers of nodes removed and positive nodes; = 13 nodes removed and < 4 positive nodes (n=12) vs. < <math>13 nodes removed and < 4 positive nodes (n=6), p = 0.016, and < 13 nodes removed and < 4 positive nodes (n=6) vs. = 4 positive nodes (n=7), p=0.499. C; According to lymph node density; = 20% (n=XX) vs. < 20% (n=XX), p = 0.0001. All p-values were determined by log-rank test.

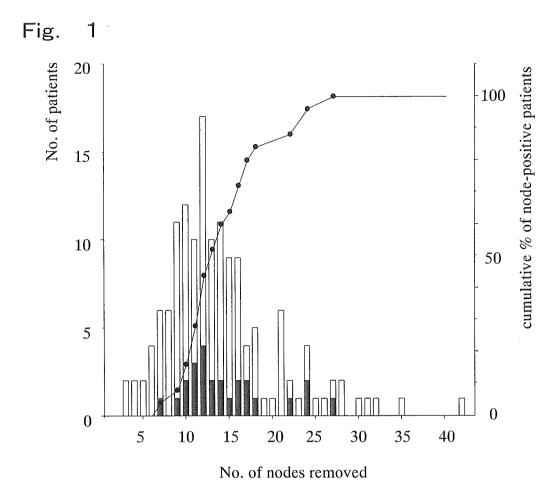


Fig. 2

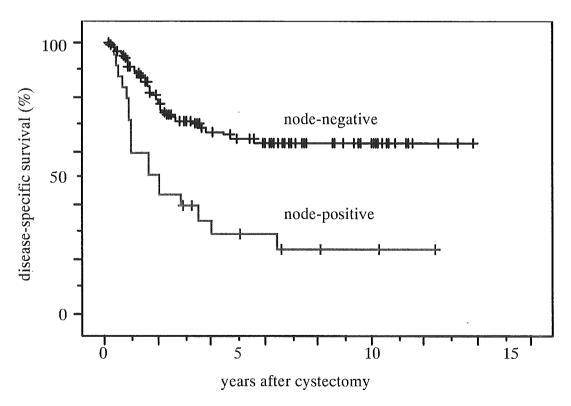


Fig. 3A

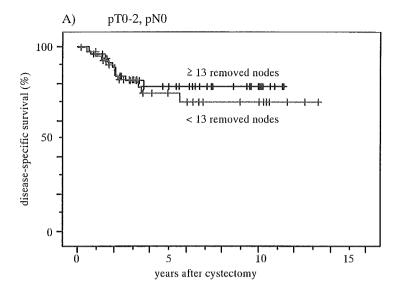


Fig. 3B

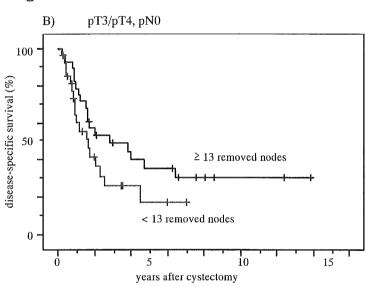


Fig. 3C

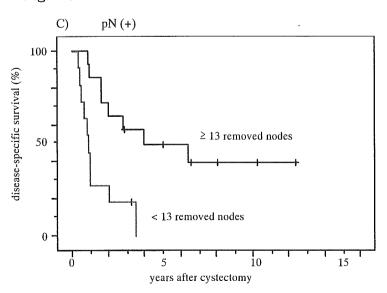


Fig. 4A

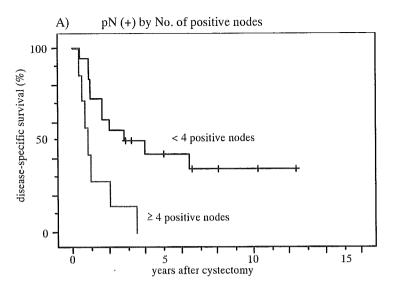


Fig. 4B

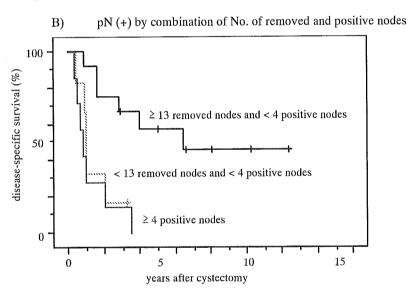


Fig. 4C

