

# Chapter 7

## THE PLACE OF PARA-AORTIC LYMPH NODE DISSECTION (PAND) IN GASTRIC CANCER SURGERY

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### INTRODUCTION

Gastric cancer is responsible for about 7 millions of death every year worldwide ( Kamangar F, Dores GM & Anderson WF,2006). Lymph node metastasis is considered as one of the most significant prognostic factors in gastric cancer. D2 lymph node dissection is practiced as the standard surgical therapy, actually all around the world but it is more common in the east( Sasako M & et al,2006). Para-aortic lymph node (PAN, no:16) is the final station where lymph nodes drain into. (N3) It is known as distant metastasis and %18-40 PAN metastasis is observed in advanced stomach cancer(Japanese Gastric Cancer Association 2006, Isozaki H & et al,1999 ,Takashima S & Kosaka T,2005 ).

### DEFINITION

Para-aortic lymph node dissection represents the total removal of all lymph nodes following D2 lymph node dissection, namely, 16a2 (PALNs between the level of the celiac axis and left renal-vein) and 16b1 (PALNs between the left renal vein and inferior mesenteric artery). (If distal gastrectomy would be implemented, dissection of the left upper lateral nodes 16a2-lat should be sufficient(Japanese Gastric Cancer Association,2017, Sano T & et al,2004) .Para-aortic lymph node dissection is addressed in the literature with various names such as PAND, D3, R3 Para-aortic lymph node dissection.

### HISTORY

It was revealed 100 years ago by the biologists named Halsted, Moynihan and Snow that lymphatic system was effective in distal metastasis by serving as a filter holding cancer cells (Halsted WS. I,1907, BG M,1908, Neuhaus SJ, & et al,2004). Removal of perigastric lymph nodes in stomach cancer was adopted by Japanese surgeons and adhered as the main principle of cancer surgery.

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surgery with sole chemotherapy in patients with para-aortic lymph node involvement and is still continuing (started in 2016, planned to be completed in 2021.) Arm A: Surgery will be performed 4-6 weeks after preoperative chemotherapy, 4-8 cycles of FLOT will be implemented post-operatively, if necessary. Arm B: 4-8 cycles of FLOT will be implemented, surgical interventions might be permitted only for palliative purposes.( FLOT5,2015)

## REFERENCES

1. Amin MBES & et al,2017.AJCC cancer staging manual, 8th edn. Springer, New York. (13.01.2019 <https://www.springer.com/la/book/9783319406176>)
2. Baba M & et al,2000. Paraaortic lymphadenectomy in patients with advanced carcinoma of the upper-third of the stomach. *Hepato-gastroenterology*,47(33):893-896.
3. Bostanci EB & et al,2004. Comparison of complications after D2 and D3 dissection for gastric cancer. *Eur J Surg Oncol*, 30(1):20–25. doi: 10.1016/j.ejso.2003.10.008
4. Chang JS & et al, 2017.Locoregional relapse after gastrectomy with D2 lymphadenectomy for gastric cancer. *Br J Surg* ,104(7):877–884. <https://doi.org/10.1002/bjs.10502>
5. ChiCTR-TRC-11001675,2015. Prospective and open multicenter randomize control study for enlarged lymphadenectomy of gastric cancer. (13.01.2019 <http://www.chictr.org.cn/hvshowproject.aspx?id=1577>)
6. De Manzoni G & et al,2015. Impact of super-extended lymphadenectomy on relapse in advanced gastric cancer. *Eur JSurg Oncol* ,41: 534-540. <https://doi.org/10.1016/j.ejso.2015.01.023>
7. Douridas GN & Pierrakakis SK,2018. Is There Any Role for D3 Lymphadenectomy in Gastric Cancer? *Front. Surg*, 5;27. <https://doi.org/10.3389/fsurg.2018.00027>
8. FLOT5,2015.Chemotherapy Alone vs. Chemotherapy + Surgical Resection in Patients With Limited-metastatic Adenocarcinoma of the Stomach or Esophagogastric Junction ( 13.01.2019 <https://clinicaltrials.gov/ct2/show/record/NCT02578368>)
9. Fujiwara Y & et al,2015.A Multidisciplinary Approach for Advanced Gastric Cancer with Paraaortic Lymph Node Metastasis.*Anticancer Research* ,35;6739-6746
10. Halsted WS. I,1907. The Results of Radical Operations for the Cure of Carcinoma of the Breast. *Annals of surgery*,46(1):1-19.
11. Ito S & et al,2017.A phase II study of chemotherapy with docetaxel,cisplatin,and S-1 followed by gastrectomy with D2 plus paraaortic lymph node dissection for gastric cancer with extensive lymph node metastasis:JCOG1002.*Gastric Cancer* ,20:322-331. <https://doi.org/10.1007/s10120-016-0619-z>
12. Isozaki H & et al,1999. Effectiveness of paraaortic lymph node dissection for advanced gastric cancer. *Hepatogastroenterology*,46(25):549–554.
13. Japanese Gastric Cancer Association ,1998. Japanese Classification of Gastric Carcinoma - 2nd English Edition - *Gastric Cancer* ,1(1):10-24. <https://doi.org/10.1007/PL00011681>
14. Japanese Gastric Cancer Association,2017. Japanese gastric cancer treatment guidelines (ver.4) 2014.*Gastric Cancer*,20(1):1-19. <https://doi.org/10.1007/s10120-016-0622-4>

15. Kaito A & et al,2017. Prognostic Factors and Recurrence Pattern of Far-advanced Gastric Cancer with Pathologically positive Para-aortic Lymph Nodes. *Anticancer Research* ,37: 3685-3692. DOI: 10.21873/anticancer.11740
16. Kamangar F, Dores GM & Anderson WF,2006. Patterns of cancer incidence, mortality, and prevalence across five continents: defining priorities to reduce cancer disparities in different geographic regions of the world. *J Clin Oncol* ,24(14): 2137–2150. DOI:10.1200/JCO.2005.05.2308
17. Keighley MR & et al,1984. Incidence and prognosis of N4 node involvement in gastric cancer. *The British journal of surgery*,71(11):863-866. <https://doi.org/10.1002/bjs.1800711121>
18. Kodera Y & et al, 2015. Gastric adenocarcinoma with para-aortic lymph node metastasis: a borderline resectable cancer? *Surg Today* ,45:1082-1090. <https://doi.org/10.1007/s00595-014-1067-1>
19. Kumagai K & et al,2018. Survival benefit of “D2-plus” gastrectomy in gastric cancer patients with duodenal invasion. *Gastric Cancer*,21;296-302. <https://doi.org/10.1007/s10120-017-0733-6>
20. Lee WJ & et al ,1995. Lymph node metastases in gastric cancer: significance of positive number. *Oncology*,52(1):45-50. <https://doi.org/10.1159/000227426>
21. Liang H & Deng J,2016. Evaluation of rational extent lymphadenectomy for local advanced gastric cancer. *J Cancer Res*,28(4):397-403. doi: 10.21147/j.issn.1000-9604.2016.04.02
22. Matsumoto T & et al,2015. HER2 expression in locally advanced gastric cancer with extensive lymph node (bulky N2 or paraaortic) metastasis (JCOG1005-A trial.) *Gastric Cancer*,18:467-475. <https://doi.org/10.1007/s10120-014-0398-3>
23. Mengardo V & et al,2018. Bencivenga M, Weindelmayer J, Pavarana M, Giacomuzzi S, Manzoni G. Para-aortic lymphadenectomy in surgery for gastric cancer: current indications and future perspectives. *Updates Surgery*,70(2):207;211. <https://doi.org/10.1007/s13304-018-0549-x>
24. Morita S & et al,2016. The clinical significance of paraaortic nodal dissection for advanced gastric cancer, *European Journal of Surgical Oncology* ,42(9):1448-1454. <https://doi.org/10.1016/j.ejso.2016.01.002>
25. Moynihan BG,1908. The surgical treatment of cancer of the sigmoid flexure and rectum. *Surg Gynecol Obstet* 1908;6:463-6.
26. Neuhaus SJ & et al,2004. Dr. Herbert Lumley Snow, MD, MRCS (1847-1930): the original champion of elective lymph node dissection in melanoma. *Annals of surgical oncology*,11(9):875-8. <https://doi.org/10.1245/ASO.2004.02.031>
27. Oyama K & et al,2012. Efficacy of pre-operative chemotherapy with docetaxel, cisplatin, and S-1 (DCS therapy) and curative resection for gastric cancer with pathologically positive para-aortic lymph nodes. *J Surg Oncol* ,105(6): 535-541. <https://doi.org/10.1002/jso.22125>
28. Sano T & et al,2004. Gastric cancer surgery: Morbidity and mortality results from a prospective randomized controlled trial comparing D2 and extended para-aortic lymphadenectomy - Japan Clinical Oncology Group study 9501. *J Clin Oncol* ,22(14): 2767-2773. DOI: 10.1200/JCO.2004.10.184
29. Sasako M & et al,2006. Modern surgery for gastric cancer--Japanese perspective. *Scand J Surg* ,95:232-235. <https://doi.org/10.1177/145749690609500404>

30. Sasako M & et al,2008. D2 lymphadenectomy alone or with para-aortic nodal dissection for gastric cancer. *N Engl J Med*, 359(5): 453-462. DOI: 10.1056/NEJ-Moa0707035
31. Takashima S & Kosaka T,2005. Results and controversial issues regarding a para-aortic lymph node dissection for advanced gastric cancer. *Surg Today* ,35(6):425–431. <https://doi.org/10.1007/s00595-004-2976-1>
32. Tokunaga M & et al,2010. Can superextended lymph node dissection be justified for gastric cancer with pathologically positive paraaortic lymph nodes? *Ann Surg Oncol* ,17(8): 2031-2036. <https://doi.org/10.1245/s10434-010-0969-4>
33. Tsujinaka T & et al,2007. Influence of Overweight on Surgical Complications for Gastric Cancer:Results From a Randomized Control Trial Comparing D2 and Extended Para-aortic D3 Lymphadenectomy(JCOG9501).*Anal of Surgical Oncology*, 14(2):335-361. <https://doi.org/10.1245/s10434-006-9209-3>
34. Wang D & et al, 2016.Shen G,Shou C,Zhu K,Chen C,Yu J.Treatment of Gastric Cancer With Paraaortic Lymph Node Metastasis.*Int Surg* ,101:583-589. <https://doi.org/10.9738/INTSURG-D-16-00148.1>
35. Zhu Z & Chen J,2018. Explanation of the 7 Clinical Questions in Japanese Gastric Cancer Treatment Guidelines of Version 4.*Journal of Surgery*, 6(5): 112-115. doi: 10.11648/j.js.20180605.11