



血液惡性疾病診療指引

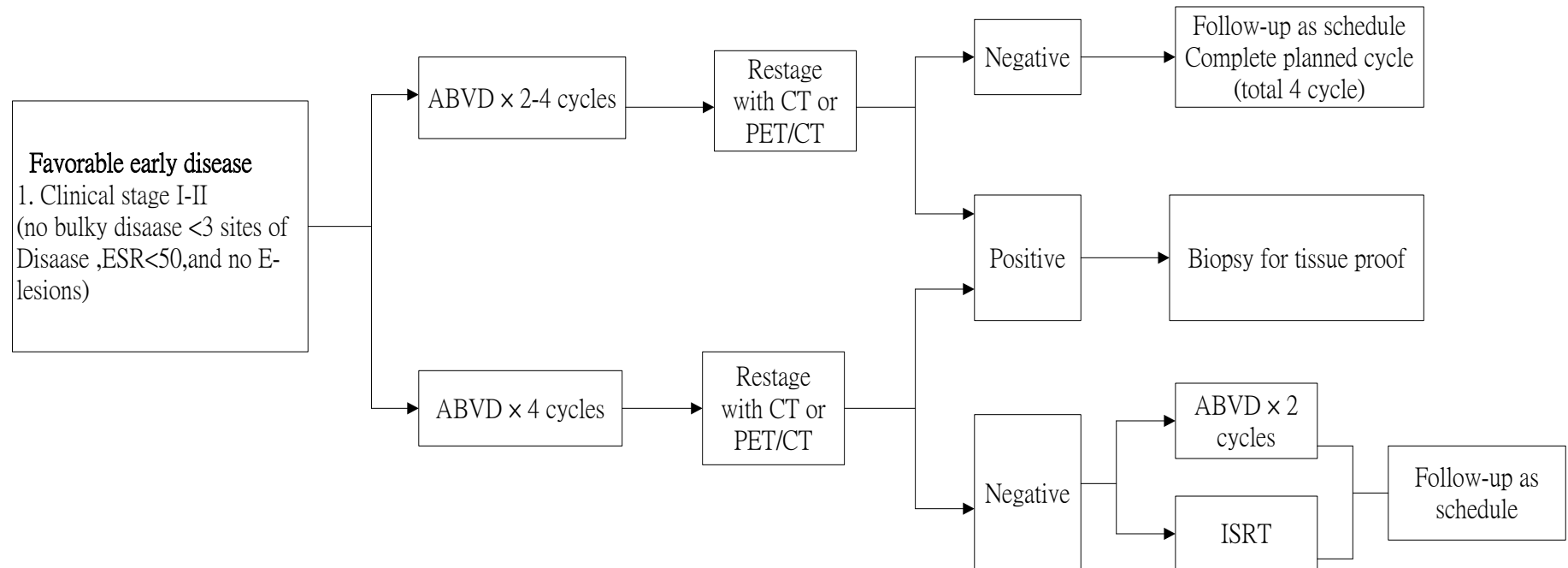
造血癌多專科團隊擬定

99.04 初制
113.03 修訂

Diagnosis	workup
1.Excisional biopsy (recommended) 2.Core needle biopsy may be adequate if diagnostic 3.Immunohistochemistry evaluation	1.Complete H&P B symptoms Performance status Examination of lymph nodes Liver, spleen 2.CBC/DC, ESR 3.Biochemistry: AST,ALT, uric acid, Cr, Ca, albumin, total protein, sugar, LDH,total bilirubin 4.Others: HBsAg, anti-HCV Ab 5.CXR (encouraged) 6.Diagnosis Whole body CT scan or PET scan 7.BM aspiration + biopsy (optional) 8.HIV (encouraged)

Staging		Risk factors																						
Lugano Classification for Hodgkin and Non-Hodgkin Lymphoma																								
Staging	Lugano Stage	※若為stage I or II, 有大於1個以上的risk factors, 為unfavorable risk unfavorable factors(localized presentations) (1)Bulky disease: ➤Mediastinal mass (chest x-ray): Maximum mass width 1 Maximum intrathoraci diameter 3 ➤Any mass > 10cm (CT) (2)Erythrocyte sedimentation rate ≥ 50, if asymptomatic (3)> 3 lymphoid regions (4)B symptoms ※ 若為stage III or IV,IPS大於4個以上的risk factors, 為unfavorable riskInternational prognostic score (IPS): International prognostic score (IPS)1 point per factor (advanced disease)2 • Albumin <4 g/dL • Hemoglobin <10.5 g/dL • Male • Age ≥ 45 years • Stage IV disease • Leukocytosis (white blood cell count ≥ 15,000/mm3) • Lymphocytopenia (lymphocyte count <8% of WBC count, and/or lymphocyte count <600/mm3) <table><tr><th>Score</th><th>5 years PFS (%)</th><th>5 years OS (%)</th></tr><tr><td>0</td><td>84</td><td>89</td></tr><tr><td>1</td><td>77</td><td>90</td></tr><tr><td>2</td><td>67</td><td>81</td></tr><tr><td>3</td><td>60</td><td>78</td></tr><tr><td>4</td><td>51</td><td>61</td></tr><tr><td>5~7</td><td>42</td><td>56</td></tr></table>		Score	5 years PFS (%)	5 years OS (%)	0	84	89	1	77	90	2	67	81	3	60	78	4	51	61	5~7	42	56
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1	77			90																				
2	67			81																				
3	60			78																				
4	51	61																						
5~7	42	56																						
I	Involvement of a single lymphatic site (i.e., nodal region, Waldeyer’s ring, thymus, or spleen)																							
IE	Single extralymphatic site in the absence of nodal involvement (rare in Hodgkin lymphoma)																							
II	Involvement of two or more lymph node regions on the same side of the diaphragm																							
IIE	Contiguous extralymphatic extension from a nodal site with or without involvement of other lymph node regions on the same side of the diaphragm																							
III	Involvement of lymph node regions on both sides of the diaphragm ; nodes above the diaphragm with spleen involvement																							
IV	Diffuse or disseminated involvement of one or more extralymphatic organs, with or without associated lymph node involvement ; or noncontiguous extralymphatic organ involvement in conjunction with nodal Stage II disease or any extralymphatic organ involvement in nodal Stage III disease Stage IV includes any involvement of the CSF, bone marrow, liver, or multiple lung lesions (other than by direct extension in Stage IIE disease)																							

Management of Classical Hodgkin lymphoma (CHL) CS IA-IIA Favorable Disease



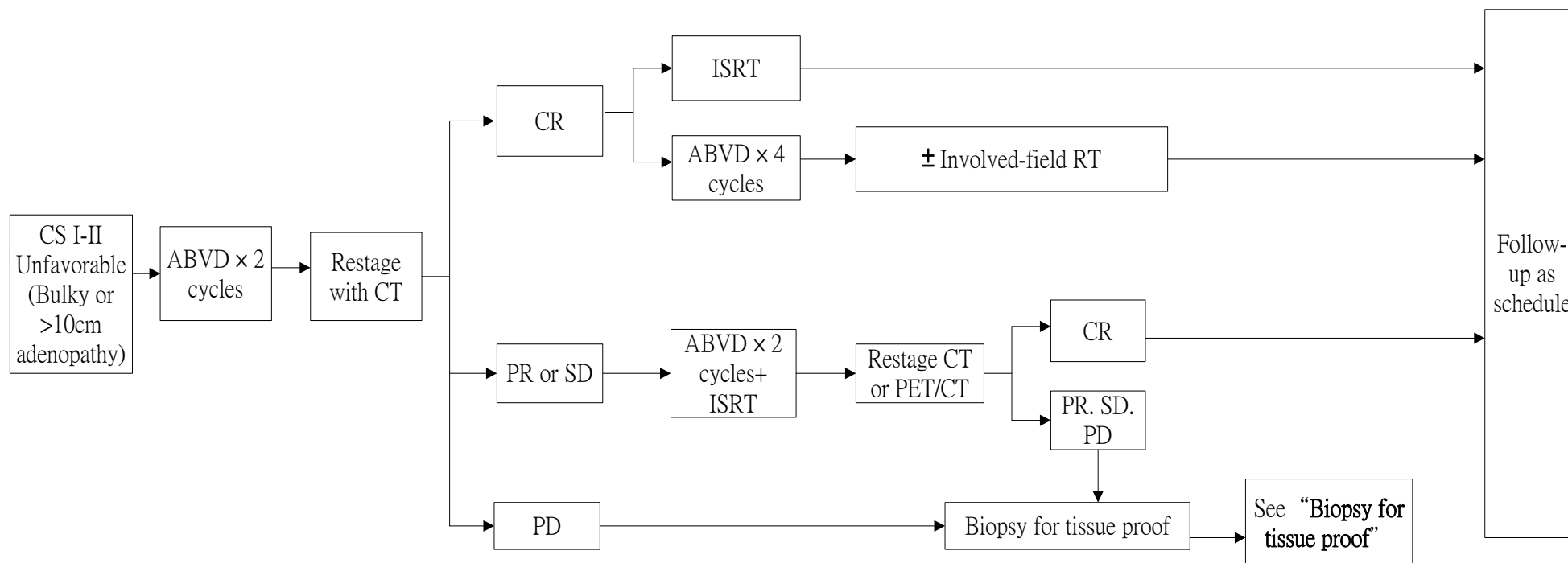
註：

ISRT: Involved site radiation therapy

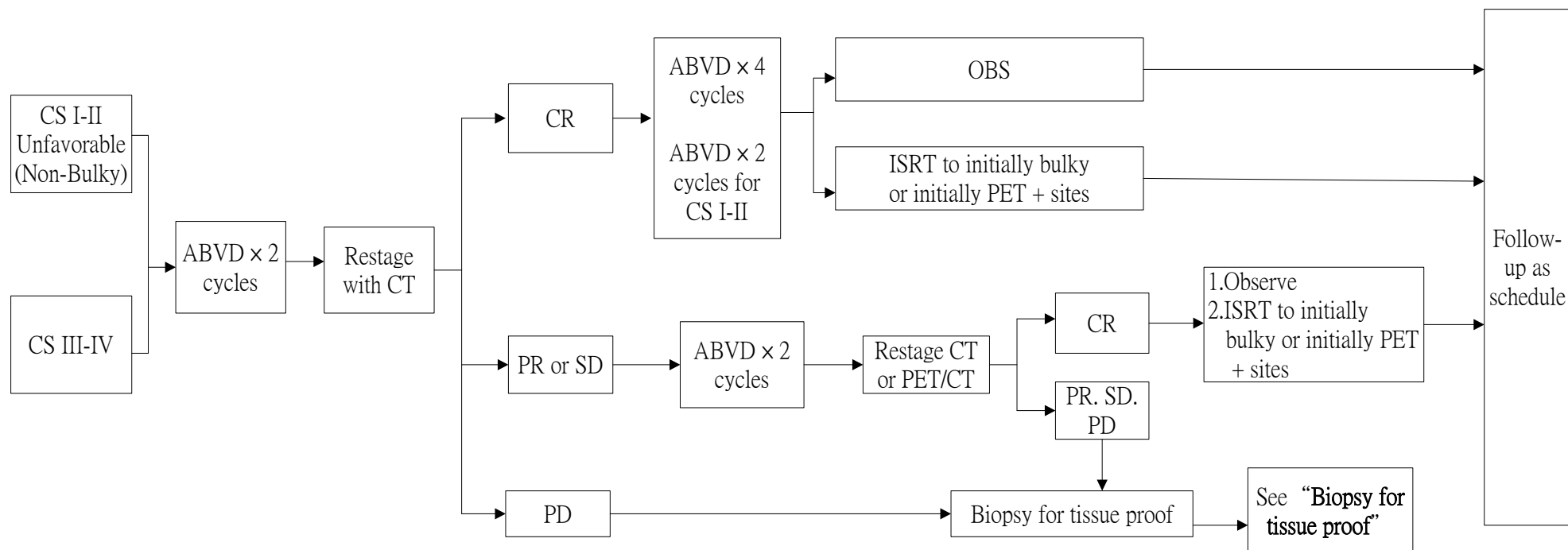
參考資料來源：

1. V2 2024. NCCN Hodgkin Lymphomas Guidelines

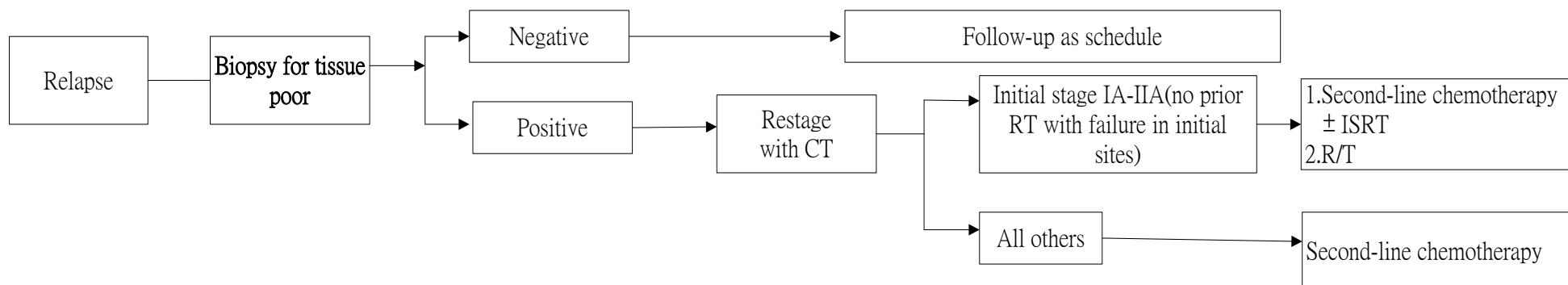
Management of CHL CS I-II Unfavorable Disease
Management of CHL CS III-IV Disease



Management of CHL CS I-II Unfavorable Disease
Management of CHL CS III-IV Disease



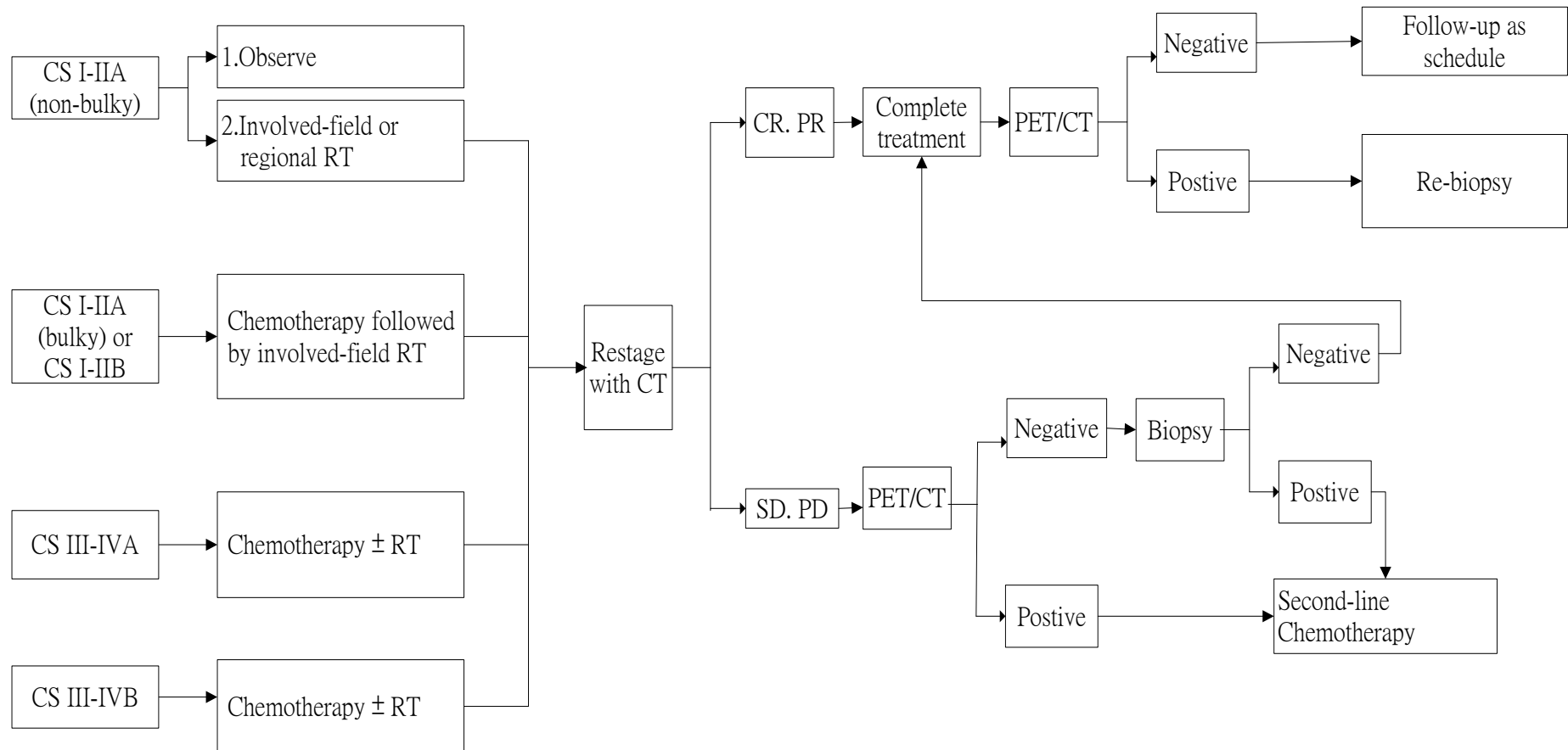
Management of Relapsed Hodgkin Lymphoma



Management of Nodular Lymphocyte-Predominant HD (NLPHL)

Stage

Primary Treatment



Follow-up schedule

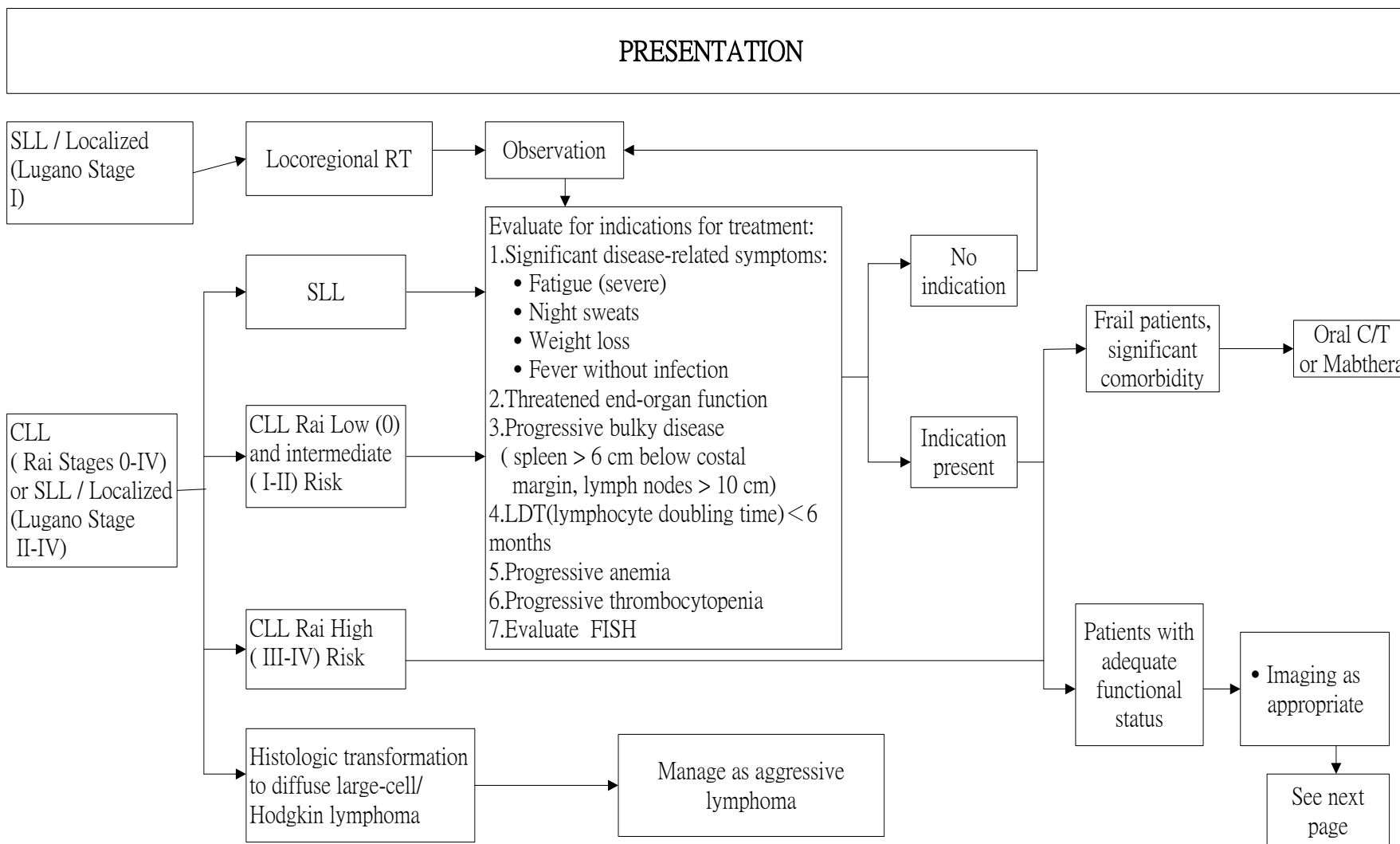
- Interim H&P:
Every 3-6 mo for 1-2 y, then every 6-12 mo for next 3-5 y
- Laboratory studies:
Every 3-6 mo for 1-2 y, then every 6-12 mo for next 3-5 y
TSH at least annually if RT to neck
- Imaging studies:
Every 6-12 mo during first 2-5 y
- After 5 years
Annually F/U

Diagnosis	Staging work-up
1.Surgical biopsy of the largest lymph nodes or mass lesion 2.Flow cytometry or cytogenetic studies(optional) 3.Immunohistochemistry evaluation	1.Complete history and physical examination including Waldeyer's rings, B symptoms, risk of HIV infection, infection, autoimmune diseases, immunosuppressive therapies 2.complete blood cell count with a differential, HBsAg, HCV Ab, HBc Ab testing (for all patients receiving anti-CD 20 antibody therapy) 3.Chemistry profiles: LDH, AST, ALT, uric acid, Cr, Ca, albumin, total protein, sugar 4.Whole body, Ga scan, Local CT,or PET 5.Bone marrow aspiration and biopsy(optional). 6.Lumbar puncture (optional) with cytology in selected patients. a. All patients with Burkitt lymphoma. b. Patients with NHL in certain sites e.g CNS, epidural space, testes, ethmoid sinus, breast and large cell lymphoma with bone marrow involvementc. c. HIV positive patients 7.Gastrointestinal studiesa. a. Esophagogastroduodenoscopy, upper gastrointestinal plus small bowel and lower gastrointestinal series for patients with gastrointestinal tract lymphoma b.Considered in patients with positive stool occult blood 8. Cytogenetic and molecular tests in selected patients (optional); cardiac ejection fractionfor age >60 if anthracycline will be used (optional). Anthracycline is contraindicated if ejection fraction is less than 50% 9.EB virus (NK/T-cell lymphoma) 10.Pregnancy testing in wonmen of child-bearing age (If chemotherapy planned) 11.Beta-2 microglobulin(optional)

參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024.NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

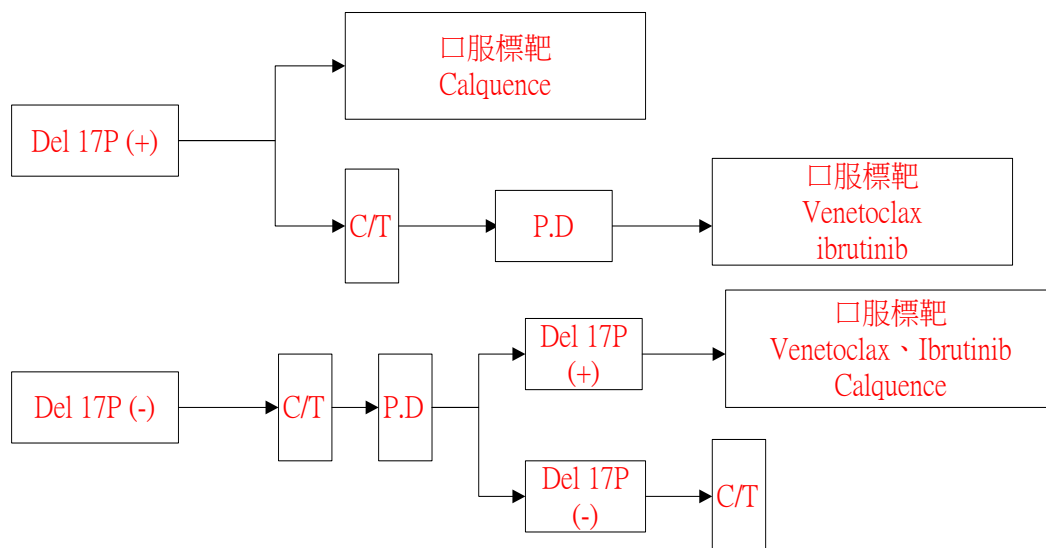
SLL/CLL Lymphoma



參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

SLL/CLL Lymphoma



(備註)

1. antibiotics for repetitive infections
2. if IgG < 500mg/dl → IVIG 0.3~0.5g/kg/month, to keep IgG > 500-700mg/dl
3. Acyclovir, for prevention or treatment of Herpes zoster
4. check Direct Coombs' test, reticulocyte count, haptoglobin for AIHA, especially for Fludarabine
5. PRCA, check parvo B19
6. 疫苗：
 - A. 每年流行性感冒疫苗，每5年肺炎雙球菌疫苗注射
 - B. 避免活菌/減毒疫苗

參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

SLL/CLL Lymphoma

CLL STAGING SYSTEMS

Rai System^a

Stage	Description	Risk Status
0	Lymphocytosis, lymphocytes in blood $>15,000/\text{mcL}$ and $>40\%$ lymphocytes in the bone marrow	Low
I	Stage 0 with enlarged node(s)	Intermediate
II	Stage 0–I with splenomegaly, hepatomegaly, or both	Intermediate
III ^c	Stage 0–II with hemoglobin $<11.0 \text{ g/dL}$ or hematocrit $<33\%$	High
IV ^c	Stage 0–III with platelets $<100,000/\text{mcL}$	High

Binet System^b

Stage	Description
A	Hemoglobin $\geq 10 \text{ g/dL}$ and Platelets $\geq 100,000/\text{mm}^3$ and <3 enlarged areas
B	Hemoglobin $\geq 10 \text{ g/dL}$ and Platelets $\geq 100,000/\text{mm}^3$ and ≥ 3 enlarged areas
C ^c	Hemoglobin $<10 \text{ g/dL}$ and/or Platelets $<100,000/\text{mm}^3$ and any number of enlarged areas

參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

SLL/CLL Lymphoma

SLL STAGING SYSTEM

Lugano Modification of Ann Arbor Staging System
(for primary nodal lymphomas)

<u>Stage^e</u>	<u>Involvement^g</u>	<u>Extranodal (E) status</u>
<i>Limited</i>		
Stage I	One node or a group of adjacent nodes	Single extranodal lesions without nodal involvement
Stage II	Two or more nodal groups on the same side of the diaphragm	Stage I or II by nodal extent with limited contiguous extranodal involvement
Stage II bulky ^f	II as above with “bulky” disease	Not applicable
<i>Advanced</i>		
Stage III	Nodes on both sides of the diaphragm	Not applicable
	Nodes above the diaphragm with spleen involvement	
Stage IV	Additional non-contiguous extralymphatic involvement	Not applicable

參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

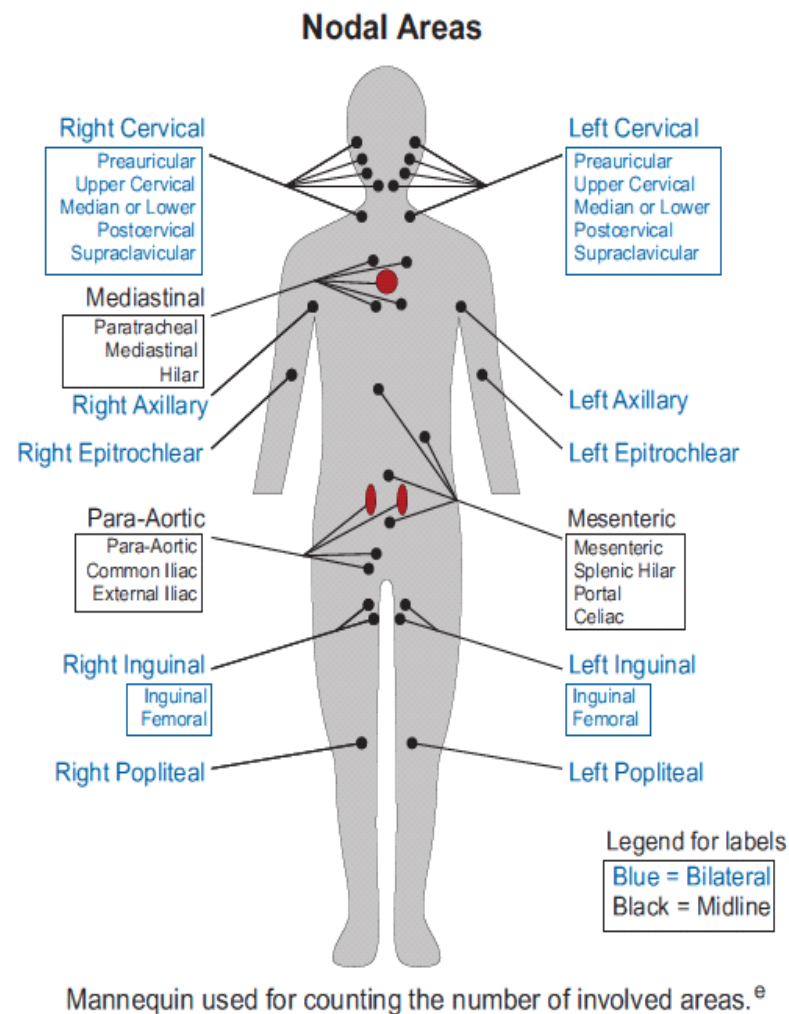
Treatment Guidelines for Follicular Lymphoma, Grades I and II (Grades III as DLBCL)

Follicular lymphoma international prognostic index (FLIPI)

1. Age ≥ 60
2. Ann Arbor stage III-IV
3. Hb level $< 12\text{g/dL}$.
4. LDH $> \text{normal}$
5. Involved nodal sites ≥ 5

score	Risk Group	5 years OS (%)	CR rate (%)
0 ~ 1	Low	91	71
2	Intermediate	78	51
3 ~ 5	High	52	36

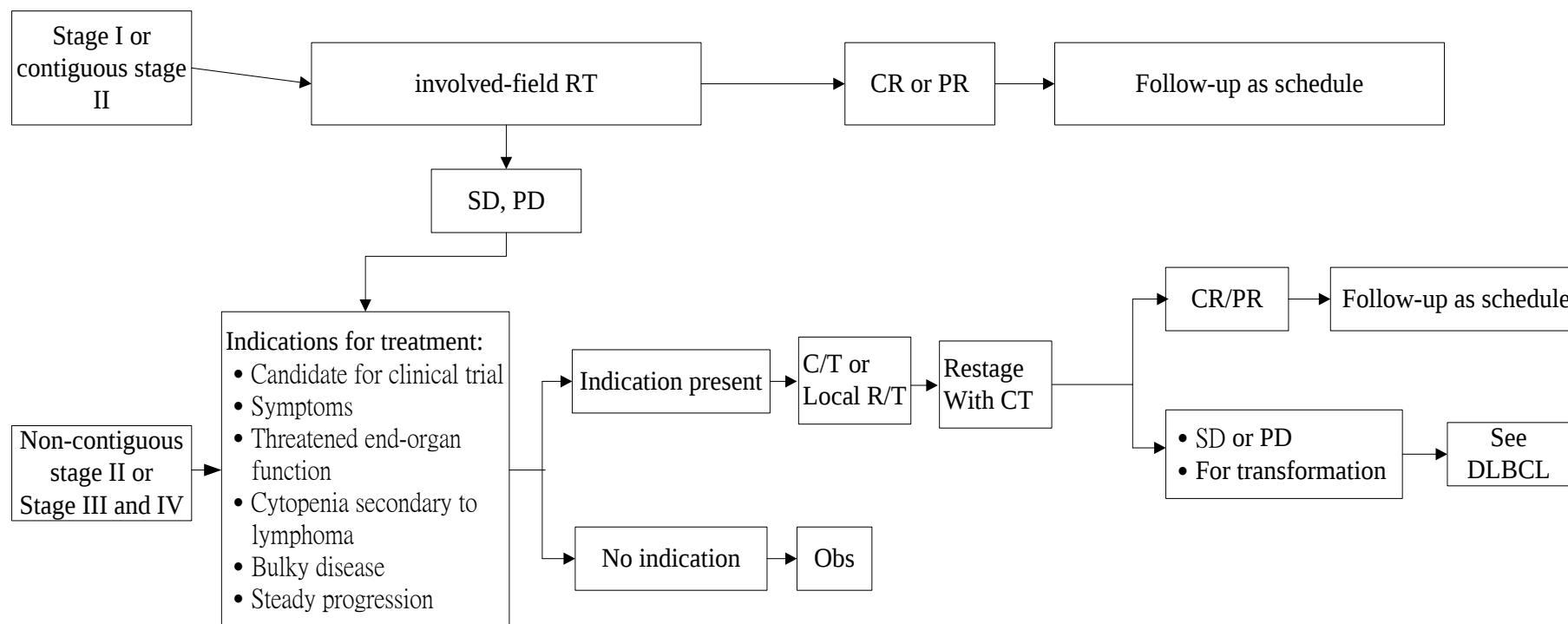
Blood 2004; 104: 1258



參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

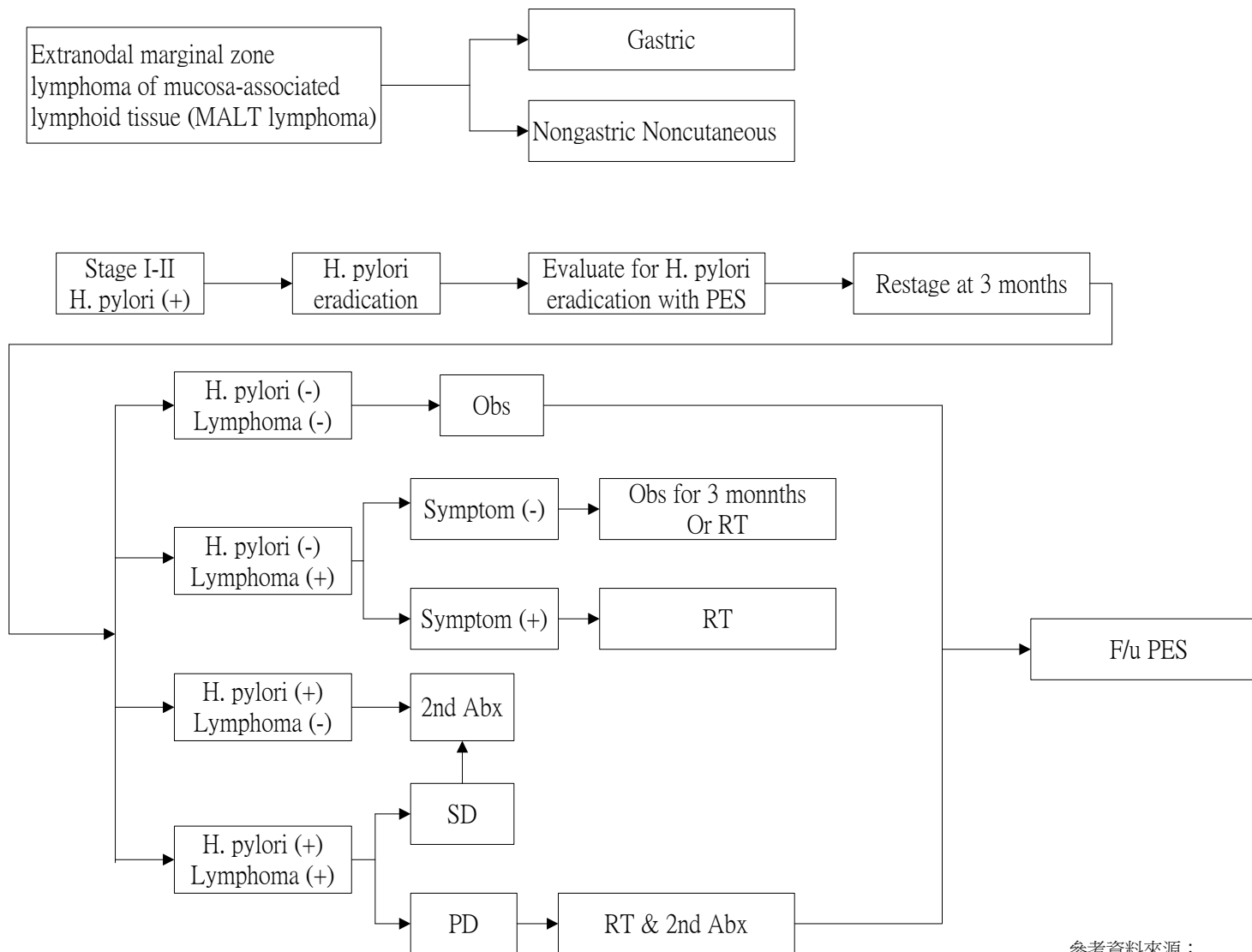
Treatment Guidelines for Follicular Lymphoma, Grades I and II (Grades III as DLBCL)



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2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

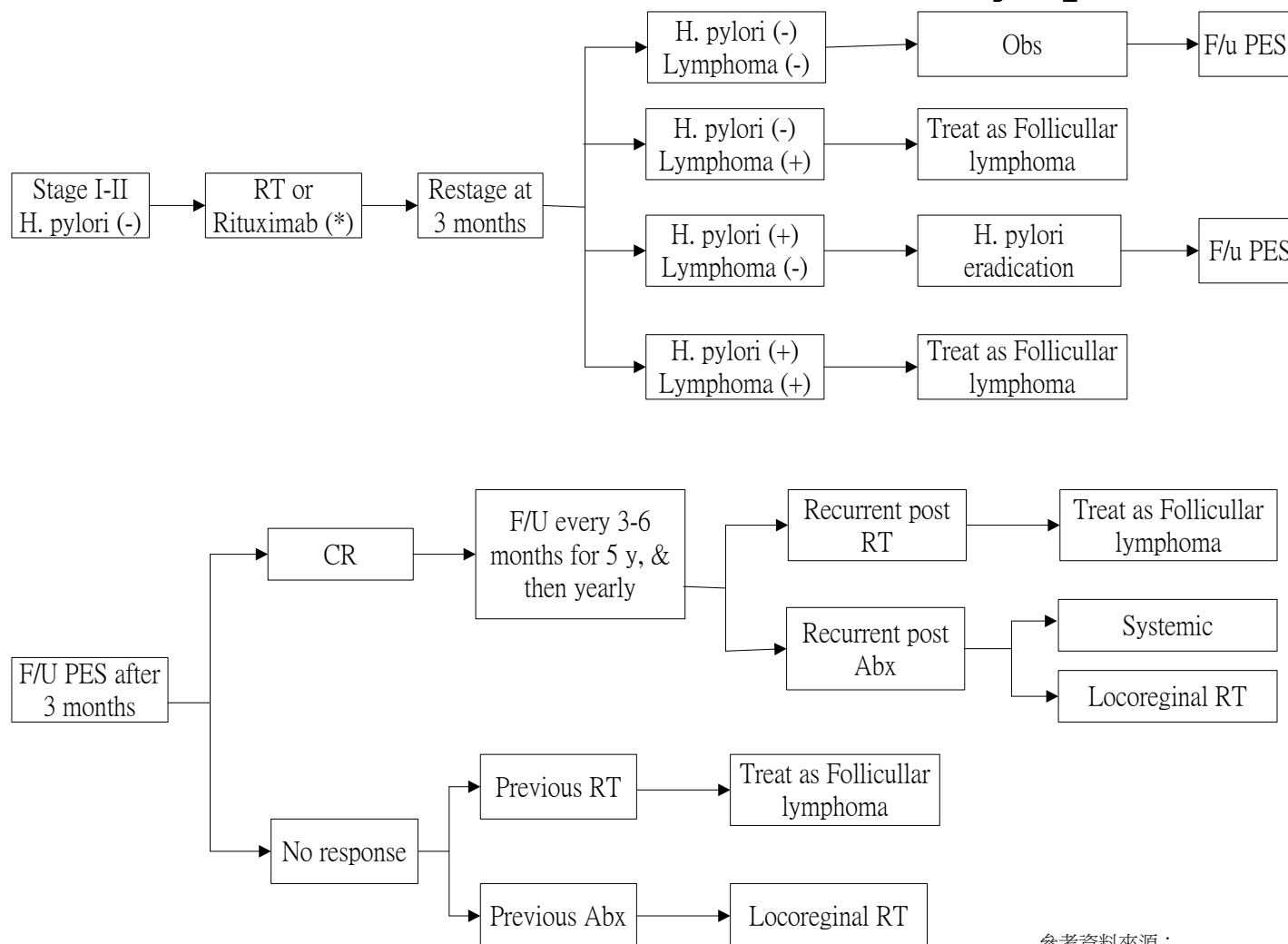
Gastric MALT Lymphoma



參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

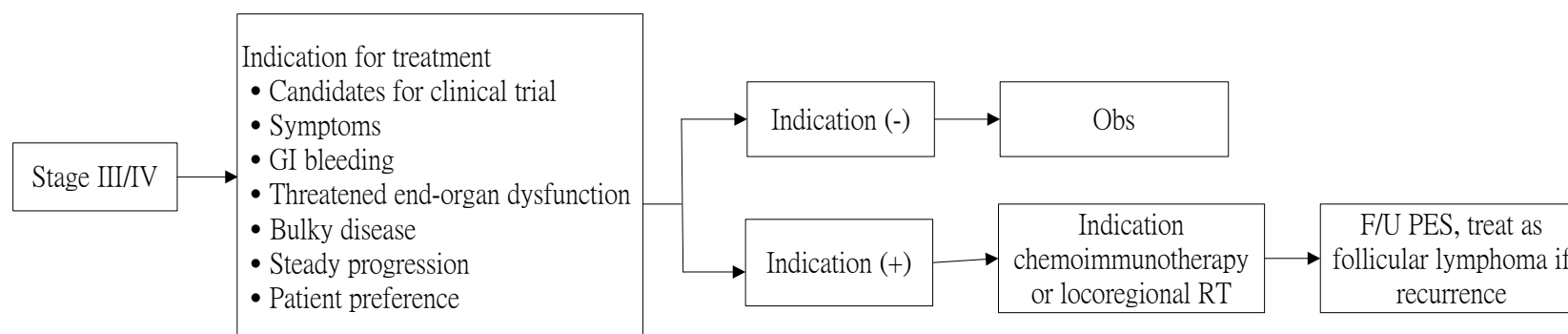
Gastric MALT Lymphoma



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2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

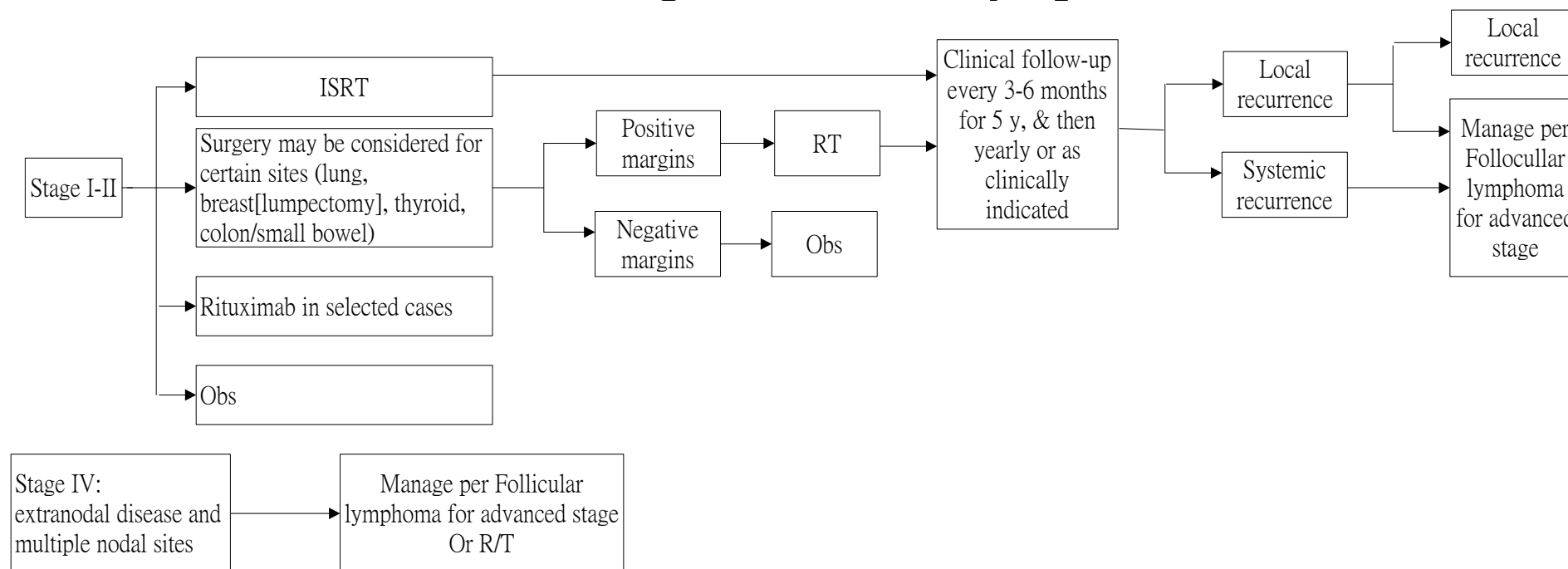
Gastric MALT Lymphoma



參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

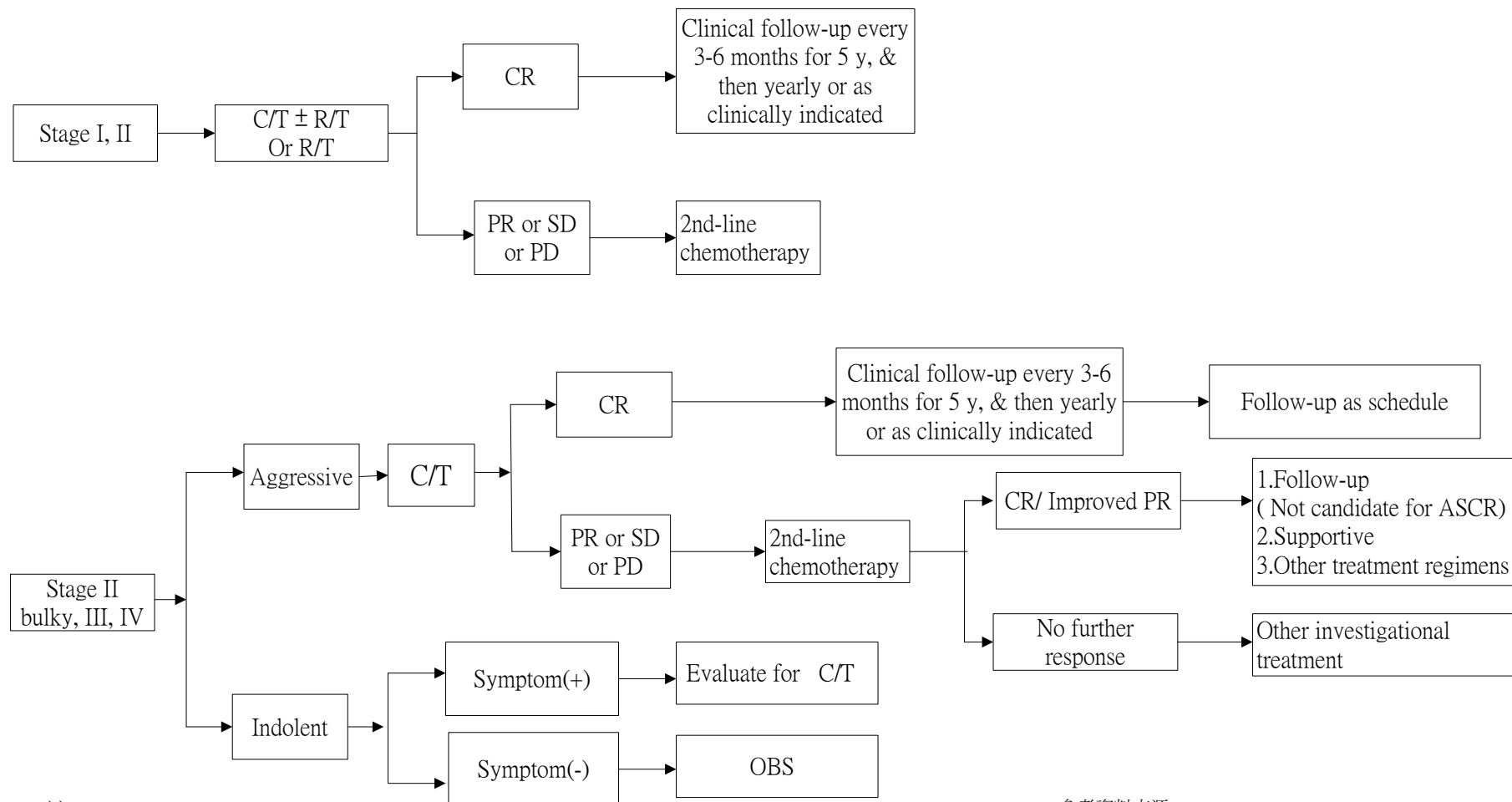
Nongastric MALT Lymphoma



參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

Treatment Guidelines for Mantle cell lymphoma



註：

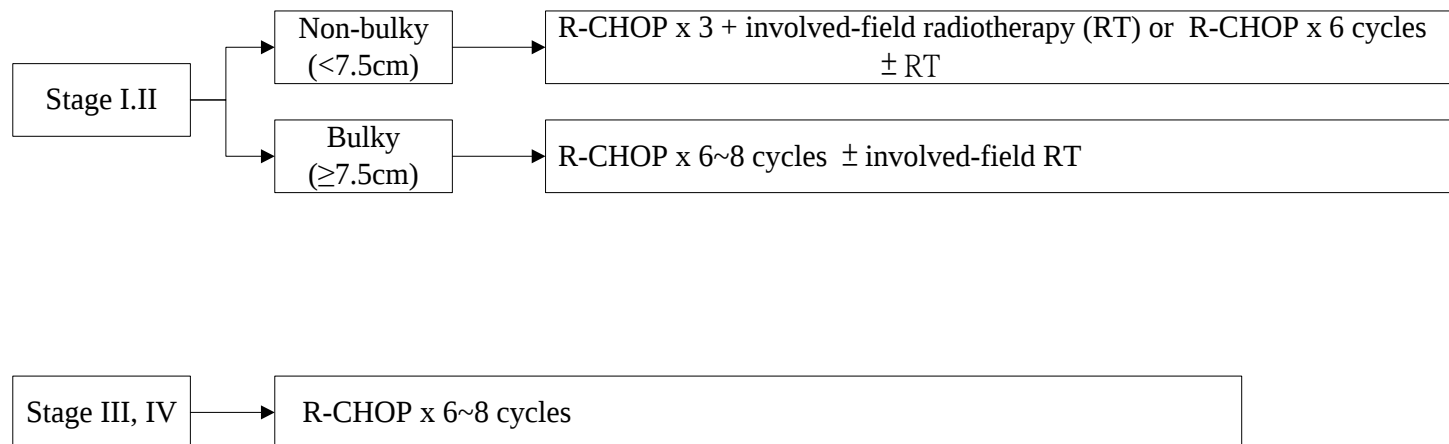
HDT: High dose therapy

ASCR: Autologous stem cell rescue

參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

Treatment Guidelines for Diffuse Large B-cell Lymphoma



註：

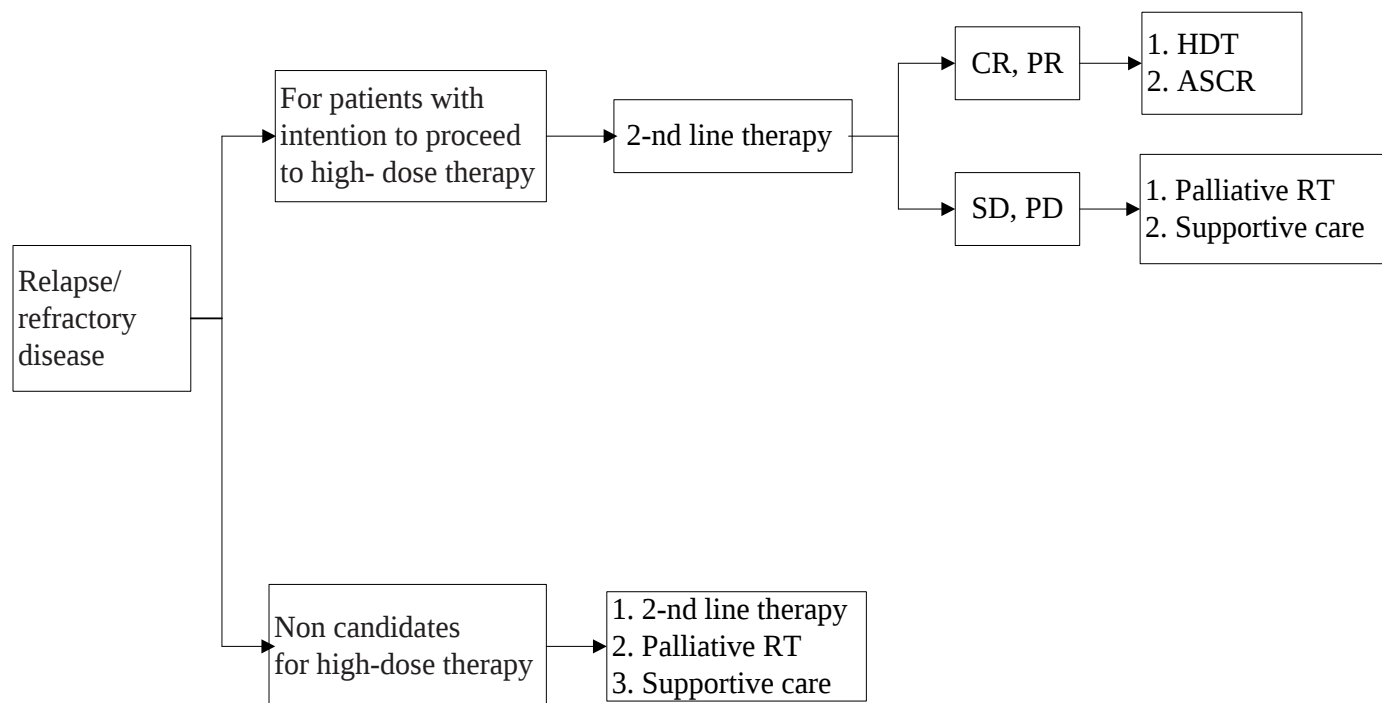
* R-CHOP or R-COP for old age

PET /CT if response is uncertain

參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

Treatment Guidelines for Diffuse Large B-cell Lymphoma



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1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

International prognostic index (IPI score)

International Prognostic Index (Aggressive NHL)

- Age >60 years
- Serum LDH >1x normal
- ECOG Performance status ≥ 2
- Ann Arbor Stage III or IV
- Extranodal involvement >1 sites

score	Risk Group	5 years OS (%)	CR rate (%)
0 or 1	Low	73	87
2	Low ~ intermediate	51	67
3	High ~ intermediate	43	55
4 or 5	High	26	44

NEJM 1993; 329: 987

Revised International Prognostic Index (for DLBCL)

score	Risk Group	4 years OS (%)	4 years PFS (%)
0	Very good	94	94
1~2	Good	79	80
3~5	Poor	55	53

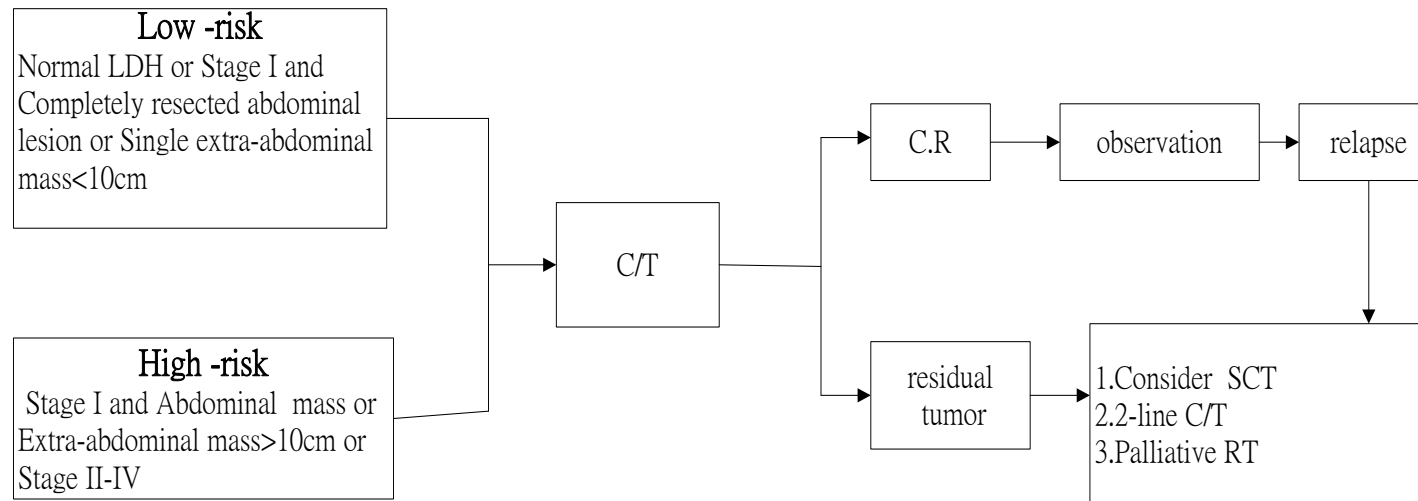
Blood 2007; 109: 1857

參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

Burkitt's Lymphoma

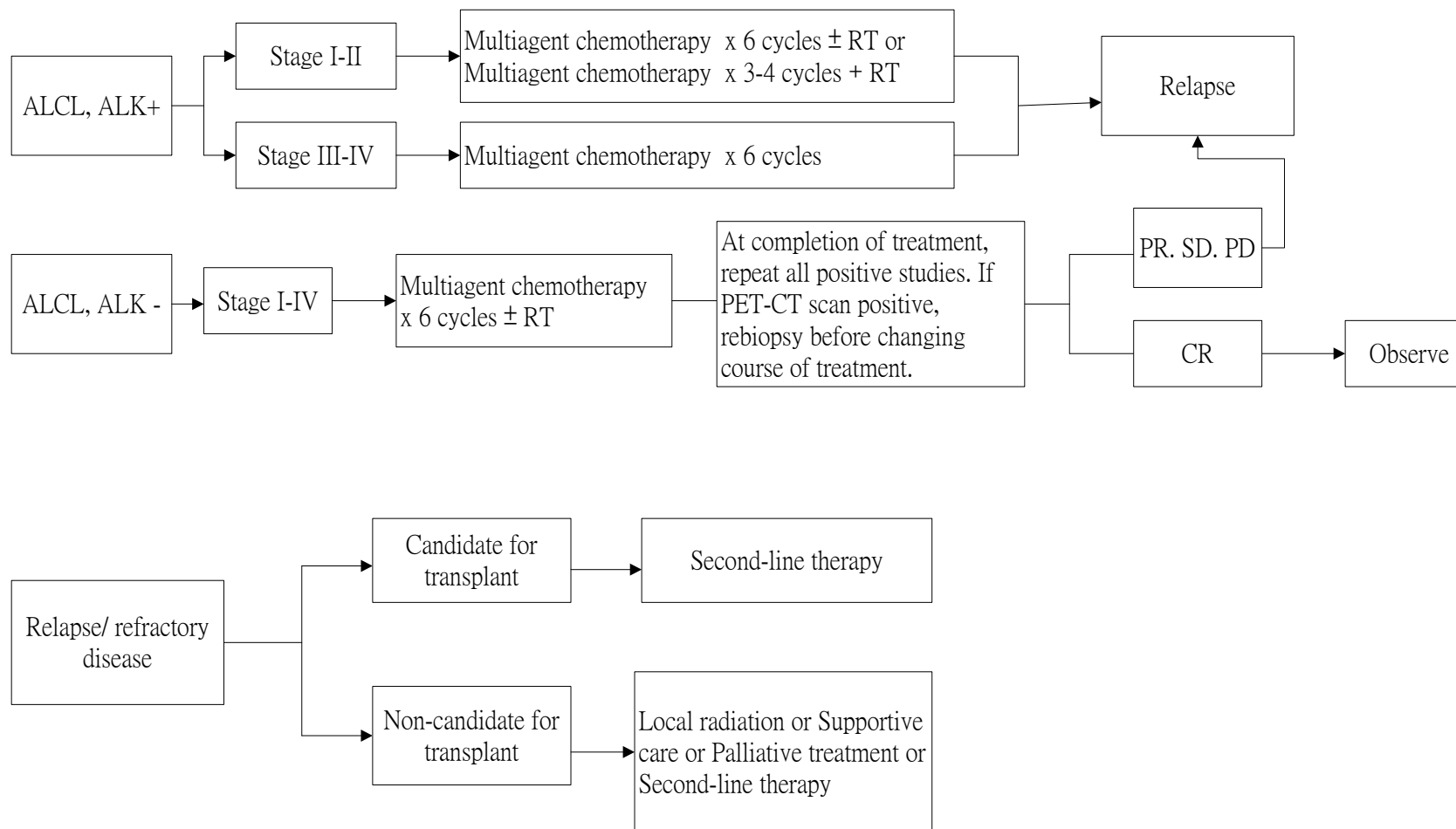
Risk assessment



參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

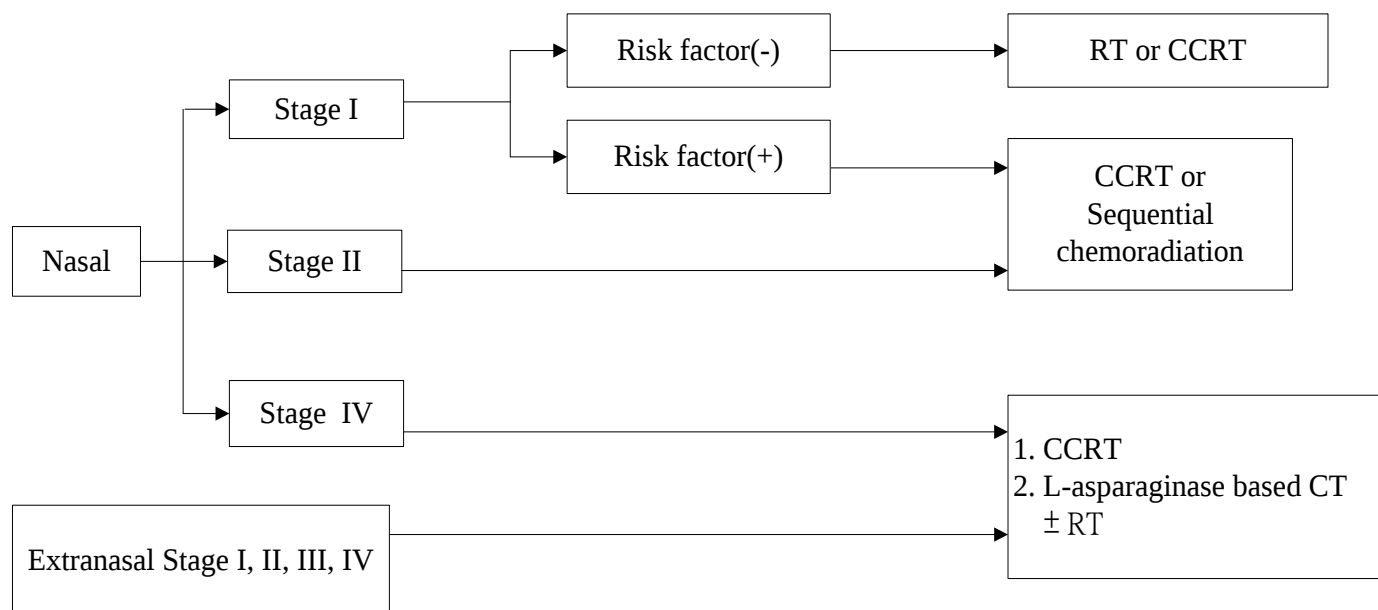
Treatment for Peripheral T-cell Lymphoma



參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

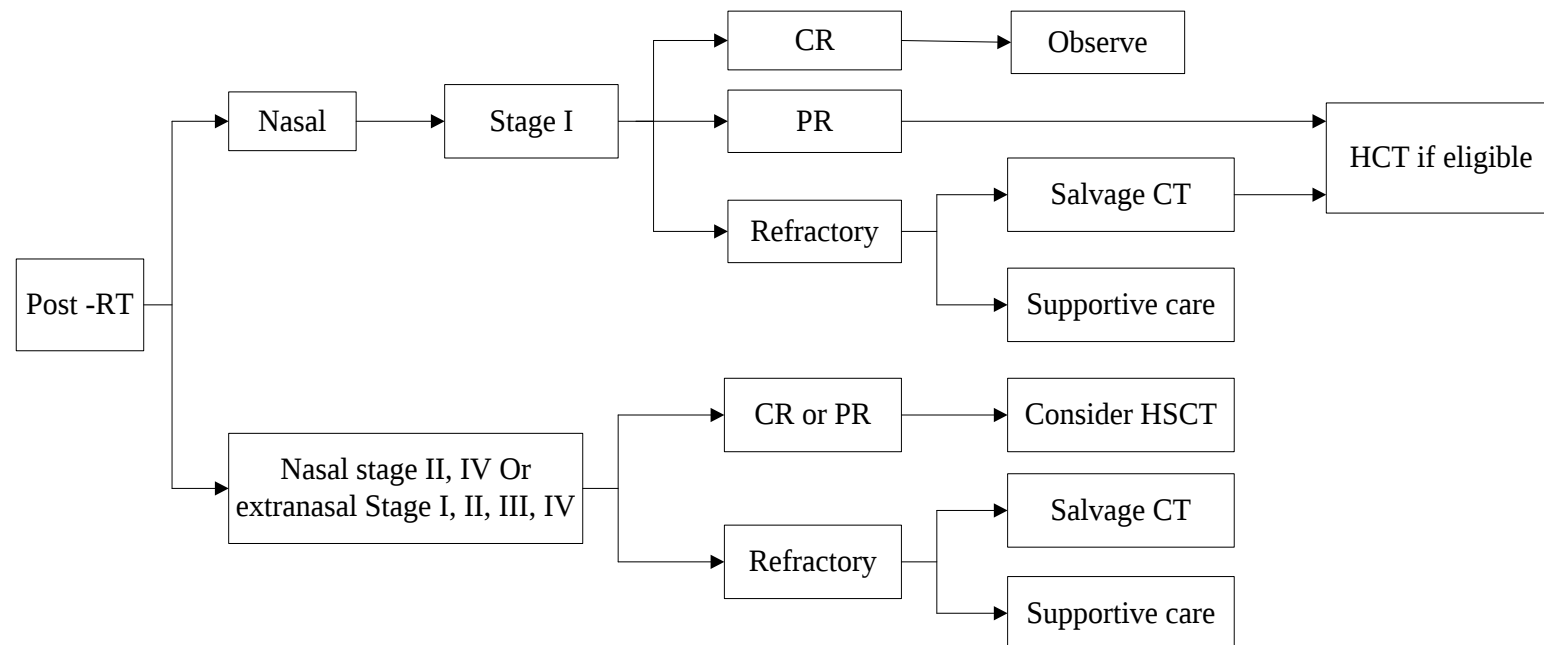
Treatment for Extranodal NK/T Cell lymphoma, nasal type



參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

Treatment for Extranodal NK/T Cell lymphoma, nasal type

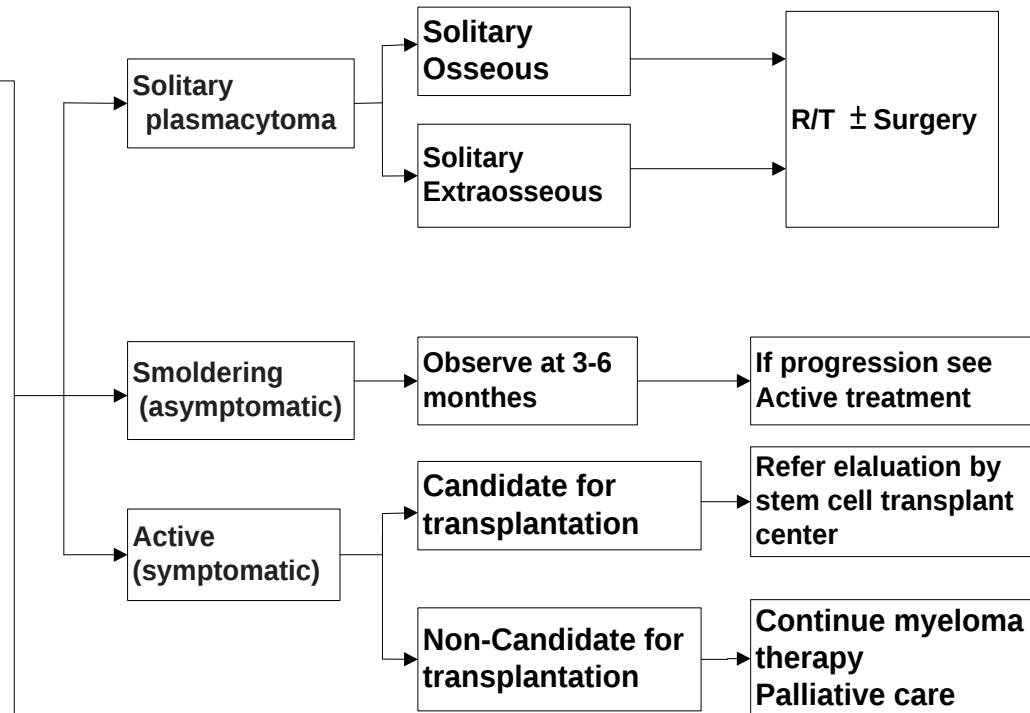


參考資料來源：

1. V1 2024. NCCN B-cell **Lymphomas** Guidelines
2. V1 2024. NCCN T-cell **Lymphomas** Guidelines
3. V1 2024. NCCN CLL/SLL Guidelines

WORKUP	Clinical presentation	Treatment
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- H&P
- CBC/DC
- Creatinine, electrolytes, BUN, urine acid
- LDH, Calcium/albumin
- Beta-2 microglobulin
- Serum free light chain assay
血清游離輕鏈球蛋白的分析
- Serum quantitative immunoglobulins
血清免疫球蛋白定量分析, serum protein electrophoresis 血清蛋白質電泳(SPEP)
- 24h urine for total protein, urine protein electrophoresis (UPEP), urine immunofixation electrophoresis (UIFE)
- Skeletal survey
- Bone marrow aspirate + biopsy, including bone marrow
- Cytogenetics
- Bence Jones Protein



STAGING SYSTEMS FOR MULTIPLE MYELOMA

Stage	ISS	RISS
I	Serum beta- 2 microglobulin < 3.5 mg/L, Serum albumin ≥ 3.5 g/dL	ISS stage I and atandard-risk Chromosomal abnormalities by FISH and Serum LDH ≤ the upper limit of nnormal
II	Neither stage I nor stage III	Not R-ISS stage I or III
III	Serum beta-2 microglobulin ≥ 5.5 mg/L	ISS stage III and either high-risk Chromosomal abnormalities by FISH and Serum LDH > the upper limit of nnormal

參考資料來源：

1. V2 2024.NCCN Multiple myeloma Guidelines