高雄榮民總醫院

皮膚癌(SCC)診療原則

2016年03月21日第一版

皮膚癌醫療團隊擬定

注意事項:這個診療原則主要作為醫師和其他保健專家診療癌症病人參 考之用。假如你是一個癌症病人,直接引用這個診療原則並 不恰當,只有你的醫師才能決定給你最恰當的治療。

修訂指引

- 本共識依下列參考資料修改版本
- NCCN 2016版 診療指引

SCC診療指引審視修訂會議討論日期

- 上次會議: 2015/09/29
- 本共識經審視後與上一版之差異

上一版:

使用NCCN 2015版 診療指引

新版:

更新 NCCN 2016版 診療指引

診斷 初步評估

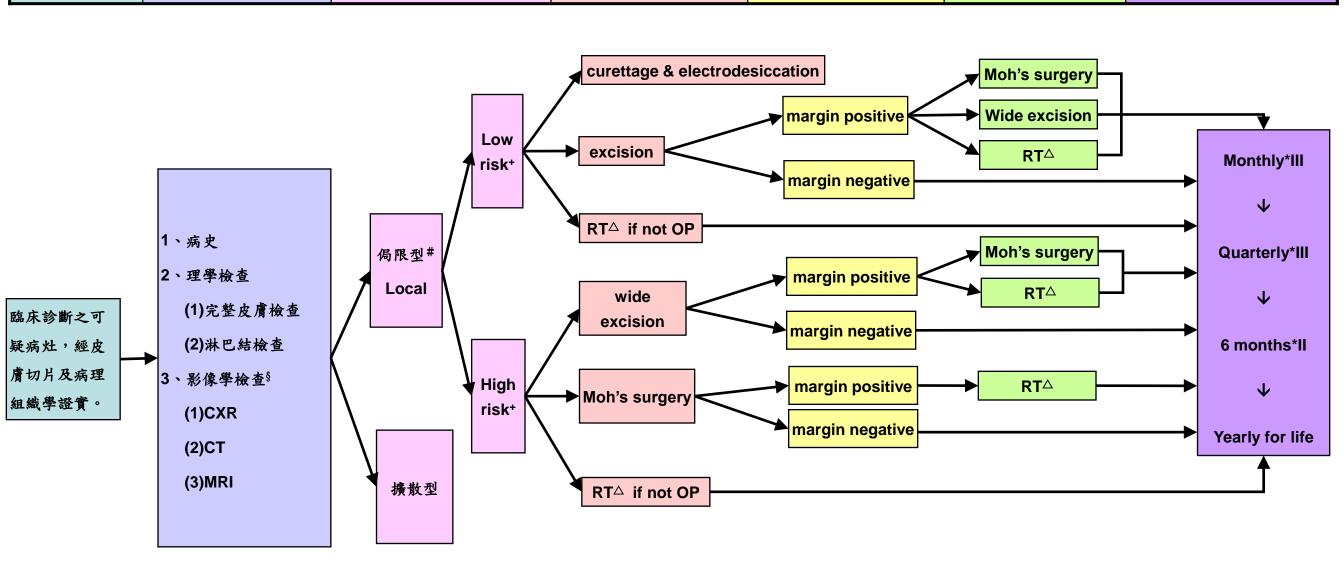
分期

初始治療

療效評估

輔助治療

追蹤



§ : Image studies is indicated for extensive disease (deep structural involvement such as bone, deep soft tissue, perineural disease)

#: Tany, N0, M0

△: RT主要針對手術不適用之情形, 附件二

+:附件一

鱗狀上皮細胞癌(SCC)

高雄榮民總醫院 臨床診療指引 2016第一版

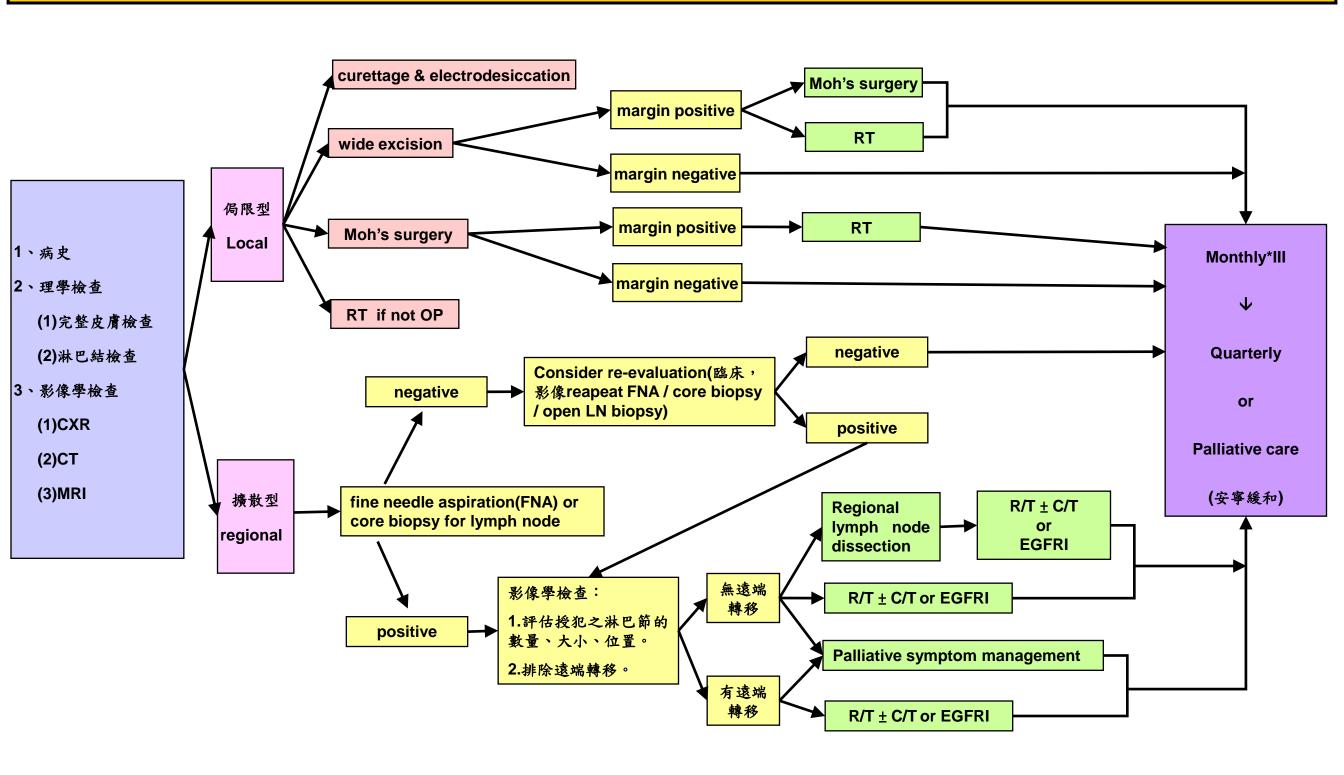
R/T△± C/T☆ or EGFRI

初步評估 診斷 分期 再評估(針對淋巴結) 初步治療 輔助治療 追蹤 1、病史 侷限型 negative 2、理學檢查 臨床診斷之 Consider re-evaluation(臨床, negative 影像reapeat FNA / core biopsy (1)完整皮膚檢查 可疑病灶, open LN biopsy) (2)淋巴結檢查 經皮膚切片 positive Monthly*III 3、影像學檢查§ 及病理組織 Quarterly*III (1)CXR 學證實。 擴散型# 6 months*II (2)CT fine needle aspiration(FNA) or core biopsy for lymph node regional (3)MRI Yearly for life or **Palliative care** Regional R/T△± C/T☆ (安寧緩和) lymph node **EGFRI** dissection 無遠端 影像學檢查: 轉移 R/T△± C/T☆ or EGFRI 1.評估授犯之淋巴節 positive 的數量、大小、位置 2.排除遠端轉移。 **Palliative symptom management** 有遠端 轉移

- § : Image studies is indicated for extensive disease (deep structural involvement such as bone, deep soft tissue, perineural disease) if perineural disease is suspected, MRI is preferred.
- #: Palpable regional lymph node(s) or abnormal lymph nodes identified by image studies. (擴散型的 "初始皮膚病灶" 治療同侷限型中high risk) T any, N1, M0 or M1 (附件三)
- ¥: Palliative symptom management, including salvage C/T
- △: RT主要針對手術不是用之情形, 附件二
- ☆: chemotherapy regimen & EGFRI, 附件四

鱗狀上皮細胞癌(SCC)

復發



鱗狀上皮細胞癌(SCC)

癌症藥物停藥準則

- ➤ 根據CTCAE (Common Terminology Criteria for Adverse Events, Version 4.0 Published: May 28, 2009 【v4.03: June 14, 2010】),出現Grade 3 ~ Grade 4 adverse event。
- ▶ 停藥至adverse event回復至Grade 1或Baseline時可再次用藥,但有些患者必須調整用藥劑量。
- ▶ 使用BRAF inhibitor時可能產生cutaneous SCC。此現象雖被CTCAE列為Grade 3 toxic effect, 但此現象不必停藥或調整劑量
- ▶特定藥物治療下疾病仍持續進展,根據追蹤及評估顯示疾病對此特定藥物治療無效 (考慮停止投藥並選擇其他治療方法)。
- > 病患要求 (Hospice care或其他因素)
- > 病患死亡

鱗狀上皮細胞癌(SCC)_ regional disease

高雄榮民總醫院 臨床診療指引 2016第一版

附件一:



NCCN Guidelines Version 1.2016 Squamous Cell Skin Cancer

NCCN Guidelines Index Squamous Cell TOC Discussion

RISK FACTORS FOR LOCAL RECURRENCE OR METASTASES

H&P	Low Risk	<u>High Risk</u>
Location/size ¹	Area L <20 mm	Area L ≥20 mm
	Area M <10 mm ⁴	Area M ≥10 mm
	Area H <6 mm ⁴	Area H ≥6 mm
Borders	Well-defined	Poorly defined
Primary vs. recurrent	Primary	Recurrent
Immunosuppression	(-)	(+)
Site of prior RT or chronic inflammatory process	(-)	(+)
Rapidly growing tumor	(-)	(+)
Neurologic symptoms	(-)	(+)
<u>Pathology</u>		
Degree of differentiation	Well or moderately differentiated	Poorly differentiated
Adenoid (acantholytic), adenosquamous (showing mucin production), desmoplastic or basosquamous (metatypical) subtypes	(-)	(+)
Depth ^{2,3} : Thickness or Clark level	<2 mm or I, II, III	≥2 mm or IV, V
Perineural, lymphatic, or vascular involvement	(-)	(+)

Must include peripheral rim of erythema.

Note: All recommendations are category 2A unless otherwise indicated. Clinical Trials: NCCN believes that the best management of any cancer patient is in a clinical trial. Participation in clinical trials is especially encouraged.

²If clinical evaluation of incisional biopsy suggests that microstaging is inadequate, consider narrow margin excisional biopsy.

³A modified Breslow measurement should exclude parakeratosis or scale crust, and should be made from base of ulcer if present.

⁴Location independent of size may constitute high risk.

Area H = "mask areas" of face (central face, eyelids, eyebrows, periorbital, nose, lips [cutaneous and vermilion], chin, mandible, preauricular and postauricular skin/sulci, temple, ear), genitalia, hands, and feet.

Area M = cheeks, forehead, scalp, neck, and pretibia.

Area L = trunk and extremities (excluding pretibia, hands, feet, nail units, and ankles).

鱗狀上皮細胞癌(SCC)_ regional disease 臨床診療指引 2016第一版

高雄榮民總醫院

附件二:



NCCN Guidelines Version 1.2016 Squamous Cell Skin Cancer

NCCN Guidelines Index Squamous Cell TOC Discussion

PRINCIPLES OF RADIATION THERAPY FOR SQUAMOUS CELL SKIN CANCER

Primary Tumor ¹	Dose Time Fractionation Schedule
<u>Tumor Diameter</u>	Examples of Dose Fractionation and Treatment Duration
<2 cm	64 Gy in 32 fractions over 6–6.4 weeks 55 Gy in 20 fractions over 4 weeks 50 Gy in 15 fractions over 3 weeks 35 Gy in 5 fractions over 5 days
≥2 cm	66 Gy in 33 fractions over 6–6.6 weeks 55 Gy in 20 fractions over 4 weeks
Postoperative adjuvant	50 Gy in 20 fractions over 4 weeks 60 Gy in 30 fractions over 6 weeks

Regional Disease: All doses at 2 Gy per fraction using shrinking field technique

After lymph node dissection

Head and neck; with ECE: 60-66 Gy over 6-6.6 weeks

56 Gy over 5.6 weeks Head and neck; without ECE: 60 Gy over 6 weeks Axilla, groin; with ECE: Axilla, groin; without ECE: 54 Gy over 5.4 weeks

No lymph node dissection

Clinically (-) but at risk for subclinical disease: 50 Gv over 5 weeks

Clinically evident adenopathy: head and neck: 66-70 Gy over 6.6-7 weeks

Clinically evident adenopathy: axilla, groin: 66 Gy over 6.6 weeks

ECE= Extracapsular extension

- Protracted fractionation is associated with improved cosmetic results.
- Radiation therapy is contraindicated in genetic conditions predisposing to skin cancer (eg, basal cell nevus syndrome, xeroderma pigmentosum) and connective tissue diseases (eg. scleroderma).

Note: All recommendations are category 2A unless otherwise indicated.

Clinical Trials: NCCN believes that the best management of any cancer patient is in a clinical trial. Participation in clinical trials is especially encouraged.

Field margins for <2 cm primary tumors should be 1-1.5 cm; for tumors >2 cm, field margins should be 1.5-2 cm. Tighter field margins can be used with electron beam adjacent to critical structures (eg., the orbit) if lead skin collimation is used. Bolus is necessary when using electron beam to achieve adequate surface dose. An electron beam energy should be chosen which achieves adequate surface dose and encompasses the deep margin of the tumor by at least the distal 90% line. Electron beam doses are specified at 90% of the maximal depth dose (Dmax). Orthovoltage x-ray doses are specified at Dmax (skin surface) to account for the relative biologic difference between the two modalities of radiation. If intensity modulated radiation therapy is used to treat primary tumors, appropriate focus must be directed at assuring that there is adequate surface dose. Appropriate medical physics support is essential.

鱗狀上皮細胞癌(SCC)_ regional disease

高雄榮民總醫院 臨床診療指引 2016第一版

附件三-1:



National Cancer

Network®

Comprehensive NCCN Guidelines Version 1.2016 **Squamous Cell Skin Cancer**

NCCN Guidelines Index Squamous Cell TOC Discussion

Differentiation

Staging			
Table 1 American Joint Co TNM Staging Clas Carcinoma (cSCC (7th ed., 2010)		Regio NX N0 N1	onal Lymph Nodes (N) Regional lymph nodes cannot be assessed No regional lymph node metastases Metastasis in a single ipsilateral lymph node, 3 cm or less in greatest dimension
T0 No evidence o	r cannot be assessed of primary tumor situ r less in greatest dimension with less than two	N2	Metastasis in a single ipsilateral lymph node, more than 3 cm but not more than 6 cm in greatest dimension; or in multiple ipsilateral lymph nodes, none more than 6 cm in greatest dimension; or in bilateral or contralateral lymph nodes, none more than 6 cm in greatest dimension
ľ	r than 2 cm in greatest dimension	N2a	Metastasis in a single ipsilateral lymph node, more than 3 cm but not more than 6 cm in greatest dimension
· ·	e with two or more high-risk feature vasion of maxilla, mandible, orbit, or temporal bone	N2b	Metastasis in multiple ipsilateral lymph nodes, none more than 6 cm in greatest dimension
	vasion of skeleton (axial or appendicular) or vasion of skull base the eyelid	N2c N3	Metastasis in bilateral or contralateral lymph nodes, none more than 6 cm in greatest dimension Metastasis in a lymph node,
** High-risk features Depth/invasion	for the primary tumor (T) staging > 2 mm thickness Clark level ≥ IV		more than 6 cm in greatest dimension int Metastasis (M)
Anatomic location	Perineural invasion Primary site ear Primary site non-hair-bearing lip	M0 M1	No distant metastases Distant metastases

Used with the permission of the American Joint Committee on Cancer (AJCC), Chicago, Illinois. The original and primary source for this information is the AJCC Cancer Staging Manual, Seventh Edition (2010) published by Springer Science+Business Media, LLC (SBM). (For complete information and data supporting the staging tables, visit www.springer.com.) Any citation or quotation of this material must be credited to the AJCC as its primary source. The inclusion of this information herein does not authorize any reuse or further distribution without the expressed, written permission of Springer SBM, on behalf of the AJCC.

Continue

Poorly differentiated or undifferentiated

鱗狀上皮細胞癌(SCC)_ regional disease

高雄榮民總醫院 臨床診療指引 2016第一版

附件三-2:



NCCN Guidelines Version 1.2016 Squamous Cell Skin Cancer

NCCN Guidelines Index
Squamous Cell TOC
Discussion

Table 1 Continued American Joint Committee on Cancer (AJCC) TNM Staging Classification for Cutaneous Squamous Cell Carcinoma (cSCC) (7th ed., 2010)			
Anatomic Stage/Prognostic Groups			
Stage 0	Tis	N0	MO
Stage I	T1	N0	MO
Stage II	T2	N0	MO
Stage III	T3	NO	MO
	T1	N1	MO
	T2	N1	MO
	T3	N1	MO
Stage IV	T1	N2	MO
	T2	N2	MO
	T3	N2	MO
	T Any	N3	MO
	T4	N Any	MO
	T Any	N Any	M1

Histologic Grade (G)

- GX Grade cannot be assessed
- G1 Well differentiated
- G2 Moderately differentiated
- G3 Poorly differentiated
- G4 Undifferentiated

Used with the permission of the American Joint Committee on Cancer (AJCC), Chicago, Illinois. The original and primary source for this information is the AJCC Cancer Staging Manual, Seventh Edition (2010) published by Springer Science+Business Media, LLC (SBM). (For complete information and data supporting the staging tables, visit www.springer.com.) Any citation or quotation of this material must be credited to the AJCC as its primary source. The inclusion of this information herein does not authorize any reuse or further distribution without the expressed, written permission of Springer SBM, on behalf of the AJCC.

附件四-1:chemotherapy regimen

chemotherapy regimen		
published C/T regimens	schedule	
Cisplatin, 100 mg/m2 IV D1	Q 21-28 days x 4 cycles	
5-FU, 1 g/m2 IV D1-3	Q 21-28 days x 4 cycles	
Bleomycin, bolus 16 mg IV D1 + 25 mg/m2 IV D1-3	Q 21-28 days x 4 cycles	

附件四-2:chemotherapy regimen & EGFRI

chemotherapy regimen & EGFRI		
published C/T regimens	schedule	
Cisplatin 100 mg/m2 IV D1	Q 21 days * 6 cycles	
5-FU 1 g/m2 IV D1-4	Q 21 days * 6 cycles	
* Cetuximab 400 mg/m2 ; 250 mg/m2 IV	400 mg/m2 * Week 1 ; then 250 mg/m2 * QW	

^{*} Cetuximab could be continued as long as the response or the stabilization persisted

附件四-2:chemotherapy regimen & EGFRI

chemotherapy regimen & EGFRI		
published C/T regimens	schedule	
Cisplatin 100 mg/m2 IV D1	Q 21 days * 6 cycles	
5-FU 1 g/m2 IV D1-4	Q 21 days * 6 cycles	
* Cetuximab, 400 mg/m2 IV Week 1, then 250 mg/m2 QW	Till IV or unacceptable toxicity	

^{*} Cetuximab could be continued as long as the response or the stabilization persisted

附件四-3:EGFRI

EGFRI		
published C/T regimens	schedule	
* Cetuximab, 400 mg/m2 IV Week 1, then 250 mg/m2 QW	Till IV or unacceptable toxicity	

^{*} Cetuximab could be continued as long as the response or the stabilization persisted

Reference

- 1. NCCN Clinical Practice Guideline in Oncology, Basal and Squamous Cell Skin Cancers, Version 2.2014
- 2. Gurudutt VV, Genden EM. Cutaneous squamous cell carcinoma of the head and neck. J Skin Cancer 2011;2011:502723.
- 3. Brantsch KD, Meisner C, Schonfisch B, et al. Analysis of risk factors determining prognosis of cutaneous squamous-cell carcinoma: a prospective study. Lancet Oncol 2008;9:713-720.
- 4. Lott DG, Manz R, Koch C, Lorenz RR. Aggressive behavior of nonmelanotic skin cancers in solid organ transplant recipients. Transplantation 2010;90:683-687.
- 5. Pack GT, Davis J. RADIATION CANCER OF THE SKIN. Radiology 1965;84:436-442.
- 6. Mendenhall WM, Ferlito A, Takes RP, et al. Cutaneous head and neck basal and squamous cell carcinomas with perineural invasion. Oral Oncol 2012;48:918-922.
- 7. Squamous cell carcinoma developing on burn scar. Ann Plast Surg 2006;56:406-408.
- 8. Classification of cutaneous squamous cell carcinoma. J Cutan Pathol 1991;18:225-226.
- 9. Neville JA, Welch E, Leffell DJ. Management of nonmelanoma skin cancer in 2007. Nat Clin Pract Oncol 2007;4:462-469.
- 10. Brodland DG, Zitelli JA. Surgical margins for excision of primary cutaneous squamous cell carcinoma. J Am Acad Dermatol 1992;27:241-248.
- 11. Ross AS, Schmults CD. Sentinel lymph node biopsy in cutaneous squamous cell carcinoma: a systematic review of the English literature. Dermatol Surg
 - 2006;32:1309-1321.
- 12. Veness MJ, Morgan GJ, Palme CE, Gebski V. Surgery and adjuvant radiotherapy in patients with cutaneous head and neck squamous cell carcinoma metastatic to lymph nodes: combined treatment should be considered best practice. Laryngoscope 2005;115:870-875.
- 13.Behshad R, Garcia-Zuazaga J, Bordeaux JS. Systemic treatment of locally advanced nonmetastatic cutaneous squamous cell carcinoma: a review of the literature.
- 14.Cranmer LD, Engelhardt C, Morgan SS. Treatment of unresectable and metastatic cutaneous squamous cell carcinoma. Oncologist 2010;15:1320-1328.
- 15.Phase II study of cetuximab as first-line single-drug therapy in patients with unresectable squamous cell carcinoma of the skin. J Clin Oncol. 2011 Sep 1;29(25):3419-26
- 16.Treatment of advanced squamous cell carcinoma of the skin with cisplatin, 5-fluorouracil, and bleomycin. Cancer 1990;66:1692–1696.
- 17. Platinum-Based Chemotherapy plus Cetuximab in Head and Neck Cancer. N Engl J Med 2008;359:1116-27.