

DOI:10.3724/SP.J.1008.2011.01016

• 研究快报 •

肾癌伴肉瘤样分化治疗及预后观察并文献复习

盛海波[△],王林辉[△],杨庆,刘冰,吴震杰,陈伟,徐遵礼,孙颖浩*

第二军医大学长海医院泌尿外科,上海 200433

[摘要] **目的** 收集、随访并分析 16 例肾癌伴肉瘤样分化患者临床资料并行文献复习,探讨肾癌伴肉瘤样分化患者临床治疗方法及预后。**方法** 2005 年至 2009 年我院泌尿外科行肾癌手术的患者共 547 例,其中伴肉瘤样分化的肾癌 16 例(2.9%)。男性 7 例,女性 9 例;年龄 52±10(36~70)岁;临床表现为肉眼血尿者 5 例,腰酸痛 5 例,乏力、消瘦 2 例,咳嗽、咯血 1 例,体检发现肾脏占位者 3 例;肿瘤均为单发,其中左侧 7 例、右侧 9 例;肿瘤最大径 7.8±4.1(2~15)cm。术前充分检查以明确肿瘤分期。术前检查提示腹膜后淋巴结肿大 2 例,右肺门淋巴结肿大 1 例,肝转移 1 例,肾静脉和(或)下腔静脉癌栓 2 例,骨转移 3 例,侵犯同侧肾上腺 1 例,侵犯输尿管上段 1 例。16 例患者均行手术治疗,病理标本常规行 H-E 染色及免疫组化检查。术后 5 例患者行 INF-α 治疗并进行随访。**结果** 所有患者行肾癌根治术,病理结果显示肾透明细胞癌伴肉瘤样分化 9 例,肾嫌色细胞癌伴肉瘤样分化 3 例,未分类肾细胞癌伴肉瘤样分化 4 例。术后分期与术前分期相同。4 例 pT1N0M0 患者随访至今无复发或转移,pT2~T4 患者已全部死亡,其无进展生存期为 5.6±4.5(2~16)个月,总生存期为 8.3±5.6(3~20)个月。**结论** 肾癌伴肉瘤样分化患者术前分期对患者预后影响较大,早期患者行手术治疗是最有效的治疗方法,建议早期患者术后定期复查以监视肿瘤复发、转移等情况,对于晚期患者靶向治疗可能有效。

[关键词] 肾肿瘤;肾细胞癌;肉瘤样分化;治疗

[中图分类号] R 737.11

[文献标志码] A

[文章编号] 0258-879X(2011)09-1016-03

Renal cell carcinoma complicated with sarcomatoid differentiation: treatment, prognosis and literature review

SHENG Hai-bo[△], WANG Lin-hui[△], YANG Qing, LIU Bing, WU Zhen-jie, CHEN Wei, XU Zun-li, SUN Ying-hao*

Department of Urology, Changhai Hospital Level, Second Military Medical University, Shanghai 200433, China

[Abstract] **Objective** To retrospectively analyze the clinical data of 16 patients with renal cell carcinoma complicated with sarcomatoid differentiation, so as to discuss the treatment and prognosis of the condition. **Methods** A total of 547 patients with renal cell carcinoma were admitted in our department during 2005-2009, and 16 (2.9%) of them also had sarcomatoid differentiation, including 7 males and 9 females, aged (52 ± 10) years old, ranging 36-70 years old. Gross hematuria was found in 5 patients, backache in 5, fatigue and weight loss in 2, cough and hemoptysis in 1; and 3 patients were accidentally found with renal occupation during physical examination. All tumors were solitary, with 7 in the left and 9 in right. The maximum tumor diameter was (7.8 ± 4.1) cm, ranging 2-15 cm. Full examination was done to confirm the tumor stage preoperatively. Preoperative examination showed that 2 patients had retroperitoneal lymph node enlargement, 1 had right pulmonary hilum lymphatic metastasis, 1 had liver metastasis, 2 had renal vein and (or) the inferior vena cava tumor thrombi, 3 had bone metastases, 1 had invasion of ipsilateral adrenal gland, and 1 had invasion of the upper ureter. All the 16 patients underwent surgical treatment, and their specimens were subjected to H-E staining and immunohistochemical examination. Five patients were treated with INF-α after operation and were followed up. **Results** All patients received radical nephrectomy. Pathological examination showed that 9 patients had clear cell renal cell carcinoma with sarcomatoid differentiation, 3 had chromophobe renal cell carcinoma with sarcomatoid differentiation, and the other 4 had unclassified renal cell carcinoma with sarcomatoid differentiation. No recurrence or metastasis was seen in the 4 pT1N0M0 patients. All the pT2-T4 patients died, with the progression-free survival time being (5.6 ± 4.5) months, ranging 2-16 months; the overall survival time was (8.3 ± 5.6) months, ranging 3-20 months. **Conclusion** Preoperation staging of patients with renal cell carcinoma complicated with

[收稿日期] 2011-07-02

[接受日期] 2011-08-04

[基金项目] 上海市市级医院新兴前沿技术联合攻关项目(SHDC12010115),上海市重点学科项目. Supported by the Municipal Hospital Level Project for Emerging and Frontier Technology of Shanghai (SHDC12010115) and Project for the Key Discipline of Shanghai.

[作者简介] 盛海波,第二军医大学临床医学八年制 2004 级学员. E-mail: urology_sean@163.com; 王林辉,博士,教授、主任医师,博士生导师. E-mail: wlhui@medmail.com.cn

△共同第一作者(Co-first authors).

*通信作者(Corresponding author). Tel: 021-81873409, E-mail: sunyh@medmail.com.cn

sarcomatoid differentiation has a great impact on patient survival. Operation is the most effective treatment for patients with early stage, and routine follow-up examination is recommended postoperatively. Targeted therapy may be an effective way for patients of advanced stage.

[Key words] kidney neoplasms; renal cell carcinoma; sarcomatoid differentiation; therapy

[Acad J Sec Mil Med Univ, 2011, 32(9): 1016-1018]

肾癌(renal cell carcinoma, RCC)约占成人恶性肿瘤的2%~3%,其发病率位居泌尿系统肿瘤第二位,但其病死率位居首位^[1]。2004年WHO肾脏肿瘤病理分类标准将肾癌分为肾透明细胞癌、肾嫌色细胞癌、肾乳头状腺癌、多房性囊性肾细胞癌、Xp11.2易位性肾癌、神经母细胞瘤伴发癌、黏液性管状及梭形细胞癌及未分类肾细胞癌等类型,而具有肉瘤样分化的肾癌在各亚型中约占1%~8%^[2-3],这部分肾癌肿瘤侵袭性及转移性很高,发现时大多已局部进展或远处转移,其中位生存期不超过1年^[3-8]。本研究收集、随访并分析16例肾癌伴肉瘤样分化患者临床资料,现报告如下。

1 资料和方法

2005年至2009年我院泌尿外科接收肾癌患者共547例,其中具有肉瘤样分化的肾癌16例,占同期收治肾癌患者的2.9%。男性7例,女性9例;年龄52±10(36~70)岁;临床表现为肉眼血尿者5例,腰部酸痛者5例,乏力、消瘦者2例,咳嗽、咯血者1例,体检发现肾脏占位者3例。肿瘤均为单发,其中左侧7例、右侧9例;肿瘤最大径7.8±4.1(2~15)cm。术前所有患者均行胸片、B超、上腹部CT

及KUB+IVP检查,必要时行胸部CT、骨ECT及上腹部MRI等检查以明确肿瘤分期。术前检查提示:腹膜后淋巴结肿大2例,右肺门淋巴结肿大1例,肝转移1例,肾静脉和(或)下腔静脉癌栓2例,骨转移3例,侵犯同侧肾上腺1例,侵犯输尿管上段1例。16例患者均行手术治疗,病理标本常规行HE染色(图1)及免疫组化检查。术后5例患者行INF-α治疗并定期随访。相关临床及随访资料见表1。

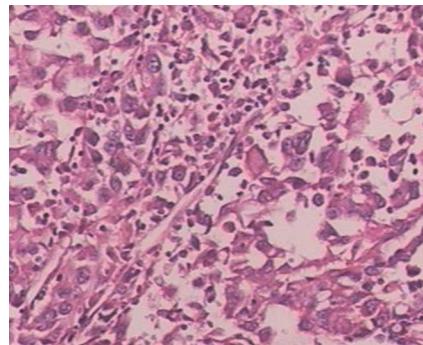


图1 肾嫌色细胞癌伴部分肉瘤样分化

Fig 1 H-E staining of chromophobe renal cell carcinoma with sarcomatoid differentiation

Original magnification: ×100

表1 患者临床相关及随访资料

Tab 1 Clinical data and follow-up data of patients

No.	Gender	Age (year)	Clinical manifestation	Tumor diameter d/cm	pTNM	Stage grouping	Follow-up t/month	PFS	Current status
1	F	51	Pain	4.5	T1bN0M1	IV	8	4	Death
2	F	44	Pain	9.0	T2N1M0	III	20	16	Death
3	M	53	Renal occupation	6.5	T1bN0M0	I	67	No	Wned
4	M	65	Cough, hemoptysis	6.0	T1bN0M1	IV	3	2	Death
5	M	43	Renal occupation	2.0	T1aN0M0	I	52	No	Wned
6	F	43	Pain	3.5	T1bN0M1	IV	8	4	Death
7	F	46	Hematuria	5.0	T4N0M0	IV	8	6	Death
8	F	58	Pain	15.0	T2N1M0	III	6	4	Death
9	F	60	Hematuria	7.0	T1bN2M0	IV	10	6	Death
10	M	36	Hematuria	9.0	T3bN0M0	III	4	2	Death
11	M	39	Weakness, weight loss	13.0	T4N0M0	IV	8	6	Death
12	F	45	Pain	12.0	T2N0M1	IV	3	2	Death
13	M	70	Renal occupation	5.0	T1bN0M0	I	36	No	Wned
14	F	62	Weakness, weight loss	15.0	T2N0M0	II	18	14	Death
15	F	52	Hematuria	4.5	T1bN0M0	I	20	No	Wned
16	M	61	Hematuria	7.0	T3bN0M0	III	3	2	Death

F: Female; M: Male; PFS: Progression-free survival; Wned: With no evidence of disease. The clinical data were collected until June 30, 2011

2 结 果

所有患者中局限性肾癌患者5例,局部进展期肾癌4例,转移性肾癌7例,皆行肾癌根治手术,1例下腔静脉癌栓患者术前行下腔静脉临时滤器置入术,根治术中成功取出癌栓及下腔静脉滤器。术前术后均无致命性并发症发生且后期死亡患者皆因癌症复发或转移致死。术后5例患者行皮下注射INF- α 治疗。术后病理诊断为肾透明细胞癌伴部分肉瘤样分化9例,肾嫌色细胞癌伴部分肉瘤样分化3例,未分类肾细胞癌伴部分肉瘤样分化4例。4例pT1N0M0患者随访至今无复发及转移,pT2~T4患者已全部死亡,其无进展生存期为5.6±4.5(2~16)个月,总生存期为8.3±5.6(3~20)个月。

3 讨 论

本研究中约81%(13/16)肾癌伴肉瘤样分化患者是因血尿、腰痛、乏力、消瘦等临床症状而被发现,只有3例患者为体检发现,且术前影像学检查和术中探查发现该类型肾癌肿瘤直径较大[(7.8±4.1)cm]且大多已发展至局部进展期(4例)或晚期(7例),这些数据与既往文献报道基本相符^[3]。肾嫌色细胞癌、透明细胞癌和集合管癌占大部分^[3],但本研究数据显示透明细胞癌伴肉瘤样分化占56.25%(9/16),与文献报道稍有出入。因本研究病例数较少,未能进行亚型间比较,但以往报道肾癌亚型伴肉瘤样变之间预后差别不大^[3,5]。既往文献报道肾癌伴肉瘤样分化pT1N0M0患者平均生存期为49.7个月,而pT2~T4的患者则为6.8个月^[9~10],截至2011年6月,本研究4例pT1N0M0期患者随访43.75(20~67)个月均无复发或远处转移,pT2~T4患者已全部死亡,其平均无进展生存期为5.6±4.5(2~16)个月,总生存期为8.3±5.6(3~20)个月,数据基本相符。

总结本组肾癌伴肉瘤样分化患者临床资料及对已有文献复习发现:(1)大部分该类型患者因血尿、腰痛、消瘦等临床表现就诊,且肿瘤直径往往较大,发现时大都处于肿瘤晚期(Ⅲ、Ⅳ患者各占25%、43.75%);(2)大部分肿瘤Furhman分级为Ⅲ、Ⅳ级,其肿瘤侵袭性及转移性很高,预后较普通肾癌差;(3)早期肾癌伴肉瘤样分化患者行肾癌根治手术能获得良好疗效,而局部晚期或远处转移患者手术治疗并不能明显延长其生存期限;(4)肾癌伴肉瘤样分化对化疗、放疗不敏感,近年来有文献报道靶向治

疗药物用于治疗肾癌伴肉瘤样分化患者取得较好的疗效^[11~12],但治疗报道例数相对较少,仍需后续大样本研究证实。

综上所述,肾癌伴肉瘤样分化患者术前分期对患者预后影响较大,早期患者行手术治疗是最有效的治疗方法,建议早期患者术后定期复查以监视肿瘤复发、转移等情况,对于局部进展或晚期肾癌伴肉瘤样分化患者靶向治疗可能有效,需要进一步研究证实。

[参 考 文 献]

- [1] Jemal A, Siegel R, Xu J, Ward E. Cancer statistics, 2010[J]. CA Cancer J Clin, 2010, 60: 277~300.
- [2] Tomera K M, Farrow G M, Lieber M M. Sarcomatoid renal carcinoma[J]. J Urol, 1983, 130: 657~659.
- [3] de Peralta-Venturina M, Moch H, Amin M, Tamboli P, Hailemariam S, Mihatsch M, et al. Sarcomatoid differentiation in renal cell carcinoma: a study of 101 cases[J]. Am J Surg Pathol, 2001, 25: 275~284.
- [4] Sella A, Logothetis C J, Ro J Y, Swanson D A, Samuels M L. Sarcomatoid renal cell carcinoma. A treatable entity[J]. Cancer, 1987, 60: 1313~1318.
- [5] Mian B M, Bhadkamkar N, Slaton J W, Pisters P W, Daliani D, Swanson D A, et al. Prognostic factors and survival of patients with sarcomatoid renal cell carcinoma[J]. J Urol, 2002, 167: 65~70.
- [6] Rodríguez-Covarrubias F, Castillejos-Molina R, Sotomayor M, Méndez-Probst C E, Gómez-Alvarado M O, Uribe-Uribe N, et al. Impact of lymph node invasion and sarcomatoid differentiation on the survival of patients with locally advanced renal cell carcinoma[J]. Urol Int, 2010, 85: 23~29.
- [7] Moch H, Gasser T, Amin M B, Torhorst J, Sauter G, Mihatsch M J. Prognostic utility of the recently recommended histologic classification and revised TNM staging system of renal cell carcinoma: a Swiss experience with 588 tumors[J]. Cancer, 2000, 89: 604~614.
- [8] Kwak C, Park Y H, Jeong C W, Jeong H, Lee S E, Moon K C, et al. Sarcomatoid differentiation as a prognostic factor for immunotherapy in metastatic renal cell carcinoma[J]. J Surg Oncol, 2007, 95: 317~323.
- [9] Selli C, Hinshaw W M, Woodard B H, Paulson D F. Stratification of risk factors in renal cell carcinoma[J]. Cancer, 1983, 52: 899~903.
- [10] Oda H, Machinami R. Sarcomatoid renal cell carcinoma. A study of its proliferative activity[J]. Cancer, 1993, 71: 2292~2298.
- [11] Lekili M, Muezzinoglu T, Nese N, Temeltas G. Sorafenib in metastatic renal cell carcinoma with sarcomatoid differentiation [J]. J Chin Med Assoc, 2010, 73: 262~264.
- [12] Arses M C, Herranz U A, Ferrán B B, Mateos L L, González J G, López R L. Temsirolimus in renal cell carcinoma with sarcomatoid differentiation: a report of three cases[J]. Med Oncol, 2011 May 11. [Epub ahead of print]