Editorial

Commentary on local therapy in men who present with a metastatic prostate cancer from special issue senior guest editor

Isaac Yi Kim

Section of Urologic Oncology, Rutgers Cancer Institute of New Jersey and Division of Urology, Rutgers Robert Wood Johnson Medical School, New Brunswick, NJ 08901, USA

Received April 23, 2019; Accepted April 23, 2019; Epub April 25, 2019; Published April 30, 2019

In contrast to localized disease, there is no effective cure for men who present with metastatic prostate cancer (mPCa) as the 5-year relative survival rate is only 28% [1]. More importantly, it has been reported that the outcome in men who present with mPCa has not improved over the last two decades [2]. Given such dismal prognosis of mPCa, new ideas and novel approaches must be explored. In this regard, recently emerging data suggest that local tumor control may enhance survival in men with mPCa [3-7]. In this special issue, we have attempted to review the latest science as well as the evidence for local therapy in men who present with a mPCa.

Prostate cancer (PCa) is the most common non-skin cancer diagnosed and the second leading cause of cancer deaths among men in the United States [8]. Although radiation and surgery are quite effective for localized disease, there is no effective cure for men with a metastatic disease. In these patients with mPCa, the NCCN Prostate Cancer Guideline states that ADT ± docetaxel is the standard of care [9]. The combination of ADT and docetaxel has also been shown to be effective especially in men with a high metastatic burden by multiple investigators [10, 11]. Most recently, a series of retrospective studies have reported that local tumor control may be beneficial in men with mPCa [12-15].

Combining cytoreduction with systemic therapies has been shown to improve survival in renal cell carcinoma, colon cancer, and ovarian cancer [16-18]. However, surgery or local therapy is not part of the standard treatment armamentarium in men with mPCa. Nevertheless, in 2006, Swanson et al have proposed that local therapy (radical prostatectomy) may be beneficial in men with mPCa due to tumor debulking, enhanced anti-tumor immunity, removing the primary site of tumor shedding, blocking paraneoplastic effects, and disrupting tumor production of hormones [19]. Since then, a body of evidence has accumulated supporting the hypothesis that treating the primary tumor improves outcome in PCa patients with clinical evidence of metastasis. First, analysis of the SEER database demonstrated that local tumor control, especially surgery, resulted in a significantly better overall survival and cancer-specific mortality in men with stage IV prostate cancer (M1a-c) [12, 14]. However, only 2.99% of eligible patients underwent surgery (245 out of 8185) [20]. Second, in patients found to have lymph node metastasis at the time of surgery. completing radical prostatectomy is associated with improved cancer-specific and overall survival [4, 5]. Third, primary PCa debulking has been shown to improve the effectiveness of ADT [6]. Since these studies are entirely retrospective, a well-designed and executed randomized clinical trials are necessary to determine whether local therapy should be incorporated into the armamentarium against mPCa. Herein, we have reviewed currently open clinical trials on this topic as well as potential systemic treatment that may be combined with local

Local therapy in metastatic prostate cancer

therapies to improve the outcome in men who have metastatic PCa on presentation.

Disclosure of conflict of interest

None.

Address correspondence to: Dr. Isaac Yi Kim, Section of Urologic Oncology, Rutgers Cancer Institute of New Jersey and Division of Urology, Rutgers Robert Wood Johnson Medical School, New Brunswick, NJ 08901, USA. E-mail: kimiy@cinj.rutgers.edu

References

- [1] Siegel R, Naishadham D and Jemal A. Cancer statistics, 2013. CA Cancer J Clin 2013; 63: 11-30.
- [2] Wu JN, Fish KM, Evans CP, Devere White RW, Dall'Era MA. No improvement noted in overall or cause-specific survival for men presenting with metastatic prostate cancer over a 20-year period. Cancer 2014; 120: 818-23.
- [3] Culp SH, Schellhammer PF and Williams MB. Might men diagnosed with metastatic prostate cancer benefit from definitive treatment of the primary tumor? A SEER-based study. Eur Urol 2014; 65: 1058-66.
- [4] Ghavamian R, Bergstralh EJ, Blute ML, Slezak J, Zincke H. Radical retropubic prostatectomy plus orchiectomy versus orchiectomy alone for pTxN+ prostate cancer: a matched comparison. J Urol 1999; 161: 1223-7.
- [5] Engel J, Bastian PJ, Baur H, Beer V, Chaussy C, Gschwend JE, Oberneder R, Rothenberger KH, Stief CG, Hölzel D. Survival benefit of radical prostatectomy in lymph node-positive patients with prostate cancer. Eur Urol 2010; 57: 754-61.
- [6] Qin XJ, Ma CG, Ye DW, Yao XD, Zhang SL, Dai B, Zhang HL, Shen YJ, Zhu Y, Zhu YP, Shi GH, Xiao WJ, Lin GW, Swanson GP. Tumor cytoreduction results in better response to androgen ablation—a preliminary report of palliative transurethral resection of the prostate in metastatic hormone sensitive prostate cancer. Urol Oncol 2012; 30: 145-9.
- [7] Shao YH, Kim S, Moore DF, Shih W, Lin Y, Stein M, Kim IY, Lu-Yao GL. Cancer-specific survival after metastasis following primary radical prostatectomy compared with radiation therapy in prostate cancer patients: results of a population-based, propensity score-matched analysis. Eur Urol 2014; 65: 693-700.
- [8] Siegel RL, Miller KD and Jemal A. Cancer statistics, 2016. CA Cancer J Clin 2016; 66: 7-30.
- [9] Mohler JL, Armstrong AJ, Bahnson RR, D'Amico AV, Davis BJ, Eastham JA, Enke CA, Farrington

- TA, Higano CS, Horwitz EM, Hurwitz M, Kane CJ, Kawachi MH, Kuettel M, Lee RJ, Meeks JJ, Penson DF, Plimack ER, Pow-Sang JM, Raben D, Richey S, Roach M 3rd, Rosenfeld S, Schaeffer E, Skolarus TA, Small EJ, Sonpavde G, Srinivas S, Strope SA, Tward J, Shead DA, Freedman-Cass DA. Prostate cancer, version 1.2016. J Natl Compr Canc Netw 2016; 14: 19-30.
- [10] James ND, Sydes MR, Clarke NW, Mason MD, Dearnaley DP, Spears MR, Ritchie AW, Parker CC, Russell JM, Attard G, de Bono J, Cross W, Jones RJ, Thalmann G, Amos C, Matheson D, Millman R, Alzouebi M, Beesley S, Birtle AJ, Brock S, Cathomas R, Chakraborti P, Chowdhury S, Cook A, Elliott T, Gale J, Gibbs S, Graham JD, Hetherington J, Hughes R, Laing R, McKinna F, McLaren DB, O'Sullivan JM, Parikh O, Peedell C, Protheroe A, Robinson AJ, Srihari N, Srinivasan R, Staffurth J, Sundar S, Tolan S, Tsang D, Wagstaff J, Parmar MK; STAMPEDE investigators. Addition of docetaxel, zoledronic acid, or both to first-line long-term hormone therapy in prostate cancer (STAMPEDE): survival results from an adaptive, multiarm, multistage, platform randomised controlled trial. Lancet 2016; 387: 1163-77.
- [11] Sweeney CJ, Chen YH, Carducci M, Liu G, Jarrard DF, Eisenberger M, Wong YN, Hahn N, Kohli M, Cooney MM, Dreicer R, Vogelzang NJ, Picus J, Shevrin D, Hussain M, Garcia JA, DiPaola RS. Chemohormonal therapy in metastatic hormone-sensitive prostate cancer. N Engl J Med 2015; 373: 737-46.
- [12] Culp SH, Schellhammer PF and Williams MB. Might men diagnosed with metastatic prostate cancer benefit from definitive treatment of the primary tumor? A SEER-based study. Eur Urol 2014; 65: 1058-66.
- [13] Heidenreich A, Pfister D and Porres D. Cytoreductive radical prostatectomy in patients with prostate cancer and low volume skeletal metastases: results of a feasibility and case-control study. J Urol 2015; 193: 832-8.
- [14] Antwi S and Everson TM. Prognostic impact of definitive local therapy of the primary tumor in men with metastatic prostate cancer at diagnosis: a population-based, propensity score analysis. Cancer Epidemiol 2014; 38: 435-41.
- [15] Fossati N, Trinh QD, Sammon J, Sood A, Larcher A, Sun M, Karakiewicz P, Guazzoni G, Montorsi F, Briganti A, Menon M, Abdollah F. Identifying optimal candidates for local treatment of the primary tumor among patients diagnosed with metastatic prostate cancer: a SEER-based study. Eur Urol 2015; 67: 3-6.
- [16] Glehen O, Mohamed F and Gilly FN. Peritoneal carcinomatosis from digestive tract cancer: new management by cytoreductive surgery

Local therapy in metastatic prostate cancer

- and intraperitoneal chemohyperthermia. Lancet Oncol 2004; 5: 219-28.
- [17] Bristow RE, Tomacruz RS, Armstrong DK, Trimble EL, Montz FJ. Survival effect of maximal cytoreductive surgery for advanced ovarian carcinoma during the platinum era: a metaanalysis. J Clin Oncol 2002; 20: 1248-59.
- [18] Flanigan RC, Salmon SE, Blumenstein BA, Bearman SI, Roy V, McGrath PC, Caton JR Jr, Munshi N, Crawford ED. Nephrectomy followed by interferon alfa-2b compared with interferon alfa-2b alone for metastatic renal-cell cancer. N Engl J Med 2001; 345: 1655-9.
- [19] Swanson G, Thompson I, Basler J, Crawford ED. Metastatic prostate cancer-does treatment of the primary tumor matter? J Urol 2006; 176: 1292-8.
- [20] Leyh-Bannurah SR, Gazdovich S, Budäus L, Zaffuto E, Briganti A, Abdollah F, Montorsi F, Schiffmann J, Menon M, Shariat SF, Fisch M, Chun F, Steuber T, Huland H, Graefen M, Karakiewicz Pl. Local therapy improves survival in metastatic prostate cancer. Eur Urol 2017; 72: 118-124.