EDITORIAL



Prostate Cancer: a comparison of focal therapy and radical prostatectomy

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Focal therapy of prostate cancer (PCa) may have been born in the U.S. [1, 2], but it has grown to late adolescence (although not full maturity) in the UK. Early observational studies from University and Imperial Colleges in London have helped make focal therapy a reasonable consideration for many men with PCa [3, 4]. Now from the UK comes the present study by Shah, Reddy and associates, who compare results of focal therapy and radical prostatectomy. The work does not provide Level 1 evidence, but for now it provides a "next best thing", since attempts to perform a randomized trial (RCT) have failed to recruit.

What the Shah-Reddy study adds is the first meaningful comparison of focal therapy (N-246) vs a standard of care, radical prostatectomy (N = 246), in men with mostly intermediate-risk PCa. Methods of focal therapy were highintensity focused ultrasound (~80%) and cryoablation (~20%). Groups were propensity-matched; data were collected prospectively from centers in the UK (N = 16) and Europe (N=3) between 2005 and 2018; and results were compared using the primary outcome of "failure-free survival" (FFS). At eight years of follow-up, some 80% of each group were failure-free, i.e., had received no additional treatment or developed metastases or died of PCa. When judged by FFS, radical prostatectomy offered no advantage over focal therapy; thus, with this article, focal therapy has gained a measure of credibility in a rightfully skeptical world.

Propensity score matching is a method used by statisticians who, when dealing with observational data, attempt to equalize the effect of known covariates on responses to different treatments. The authors have done an admirable job of putting all the major covariates into their model, and their complicated statistical methods have withstood peerreview. The results are convincing for what they are, a comparison of the efficacy of the two treatments using a retrospective analysis. Because men have proven unwilling to accept a random assignment to surgery or focal therapy, these data may remain the best available for the indefinite future.

The achievement of the authors in developing this comparison is commendable. However, to cite two of the several limitations: more than one-third of men in each group had Gleason Score 3 + 3 = 6 lesions, which would substantially increase the FFS of each group; today they would likely enter active surveillance. And the success rate for focal therapy (FFS) would have almost certainly been lower than 80%, had success been based on biopsy. Follow-up biopsies were obtained in less than half of men (personal communication, Taimur Shah), and results are not reported in the article. The study remains in the hypothesis-generating category.

The real impetus behind focal therapy is the chance of a benefit—and an even better chance of avoiding complications—in situations where treatment is desired, but life is not in immediate danger. Such is the case for many men with intermediate-risk PCa, who are willing to trade cancerspecific survival for improved quality of llfe [5]. For men with intermediate-risk PCa, which is now the commonest cancer found on MR-guided biopsy [6], proving the exact efficacy of focal therapy may be less important than reconfirming its safety.

Compliance with ethical standards

Conflict of interest Dr. Marks is co-founder Avenda Health.

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