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## **Spine Patient Optimal Radiosurgery Treatment for Symptomatic Metastatic Neoplasms (SPORTSMEN): a randomized phase II study protocol**

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# Spine Patient Optimal Radiosurgery Treatment for Symptomatic Metastatic Neoplasms (SPORTSMEN): a randomized phase II study protocol

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## Abstract

**Background:** Approximately 40% of patients with metastatic cancer will have spinal metastatic disease. Historically treated with external beam radiation therapy (EBRT) with limited durability in pain control, the increased lifespan of this patient population has necessitated more durable treatment results via spine radiosurgery/stereotactic body radiation therapy (SBRT). The goal of this study is to assess three-month pain freedom rates via the Spine Patient Optimal Radiosurgery Treatment for Symptomatic Metastatic Neoplasms (SPORTSMEN) randomized trial.

**Materials and methods:** This study is a prospective randomized three-arm phase II trial which will recruit patients with symptomatic spine metastases. All patients will be

randomized to standard-of care SBRT (24 Gy in 2 fractions), single-fraction SBRT (19 Gy in 1 fraction), or EBRT (8 Gy in 1 fraction), with the primary endpoint of three-month pain freedom (using the Brief Pain Inventory). We expect that SPORTSMEN will help definitively answer the efficacy of spine SBRT versus EBRT for achieving pain freedom, while defining the safety and efficacy of 19 Gy single-fraction spine SBRT. Local control will be defined according to Spine Response Assessment in Neuro-Oncology (SPINO) criteria.

**Discussion:** This is the first phase II trial to objectively assess optimal spine SBRT dosing in the treatment of symptomatic spine metastatic disease, while assessing spine SBRT versus EBRT. Findings should allow for better determination of the efficacy of two-fraction spine SBRT versus EBRT in the United States, as well as for the novel single-fraction 19 Gy spine SBRT regimen in patients with symptomatic spine metastases.

**Trial Registration:** Clinicaltrials.gov identifier: NCT05617716 (registration date: November 14, 2022).

**Key words:** spine radiosurgery; stereotactic body radiation therapy; external beam radiation therapy; spine metastases; pain freedom

## **Introduction**

Metastatic spine cancer incidence is increasing; the primary treatment is radiation therapy. Approximately 40% of all patients with metastatic cancer have spine metastatic disease, with nearly 200,000 new cases of spine metastases occurring annually in the United States [1–3]. Metastatic spine disease has been historically treated with external beam radiation therapy (EBRT) with conventional fractionation over 1–2 weeks, with relatively limited durability in pain control [4]. The increased lifespan of patients with metastatic cancer has resulted in an increase in the incidence of spine metastases, which has led to a need for more durable treatment results. To this end, stereotactic radiosurgery (SRS)/stereotactic body radiation therapy (SBRT) of the spine has exponentially increased in popularity since first being reported in 1995 [5–7], with the theoretical

advantages of higher tumoricidal dose and more rapid fall off between tumor and surrounding normal tissue compared with EBRT.

### ***Clinical data to date***

The result of a single randomized trial has revealed that SBRT provides superior, more durable pain control of metastatic spine cancer than EBRT. The results of this recent Phase II/III randomized trial (CCTG SC.24/TROG 17.06) involving 229 patients over a five year period revealed that for patients with symptomatic metastatic spine disease involving no more than three contiguous levels, spine SBRT of 24 Gray (Gy) in 2 fractions compared to external beam radiation therapy (20 Gy in 5 fractions) provided superior freedom from pain (using the Brief Pain Inventory) at three-months post treatment (35% versus 14%;  $p < 0.001$ ) which remained durable at six-months post-treatment (32% versus 16%;  $p = 0.004$ ) [8, 9]. However, this trial did not involve any patients from the United States, where the question of the superiority of SBRT over EBRT remains to the point where insurance companies may deny coverage of SBRT over EBRT in this patient population.

### ***Rationale***

Disparities exist in utilization of spine SBRT. Despite the superiority of SBRT over EBRT in the treatment of metastatic spine cancer from the SC.24 trial, disparities in spine SBRT utilization exist and have unfortunately remained prevalent. Previous nationwide analyses conducted over a 10-year period indicate that race and insurance status are independently associated with reduced receipt of SBRT [4, 7].

Specifically, African-American race has been associated with 20% reduced likelihood of receiving spine SBRT and Medicaid insurance associated with 30% reduced likelihood of receiving spine SBRT [4]. These findings are unfortunately consistent with an overarching trend in oncology involving racial disparities in receipt of optimal radiation therapy regardless of disease site [10–15].

The optimal SBRT treatment regimen for metastatic spine cancer has yet to be established. While the 24 Gy in 2 fraction radiosurgery course has been well-

characterized in both the aforementioned randomized trial and in a single-center retrospective study [8, 16], it is unclear whether this regimen is in fact optimal for providing pain freedom in metastatic spinal disease. While there has been variable data for 1, 2, 3, and 5 fraction spine SBRT — each of which have been used for several years<sup>7</sup>, there have been no randomized controlled trials to date specifically comparing spine SBRT dosing regimens.

Consequently, we propose the Spine Patient Optimal Radiosurgery Treatment for Symptomatic MEtastatic Neoplasms (SPORTSMEN) multicenter randomized Phase II clinical trial. The SPORTSMEN trial proposes to determine the optimal spine radiosurgery/SBRT dose, comparing 24 Gy in 2 fractions to 19 Gy in 1 fraction in the SBRT arms. With comparable biologically effective doses and biologically equivalent doses (Tab. 1), this trial will be the first to provide objective data in assessing optimal spine SBRT dosing. A novel concept, the SPORTSMEN trial has the potential to reduce spine SBRT treatment to a single fraction which could markedly reduce financial toxicity and racial disparities in treatment completion in a manner similar to the potential for the recently adopted ultrahypofractionated FAST-Forward radiation treatment regimen for breast cancer [17, 18]. As the superiority of SBRT over EBRT has yet to be established in the United States, the third arm of this trial involves conventional EBRT (8 Gy in 1 fraction).

The 19 Gy in 1 fraction arm was arrived upon based on the following evidence:

A — shorter time of treatment for the patient compared with multiple fractions [19];

B — higher dose than that tested in a previous prospective (and to-date unpublished) RTOG 0631 trial comparing single-fraction SBRT versus EBRT where only 45% of patients received 18 Gy x 1 and 55% of patients received 16 Gy x 1 (this trial used a pain scale criteria which unlike the Brief Pain Inventory did not account for mechanical instability components of spinal pain) and retrospective work assessing failure-free survival [20];

C — a nationwide analysis revealed single-fraction to be the most popular spine SBRT fractionation scheme in the United States [7];

D — recent work examining post-SBRT vertebral compression fracture (VCF) revealed that minimizing VCF risk occurs when the percentage of the vertebral body receiving at least 20 Gy is less than 24% for single-fraction SBRT, which is consistent with earlier work demonstrating significantly increased risk of VCF for spine SBRT of 20 Gy/fraction or greater [21, 22] including an unacceptably high VCF rate of 39% previously demonstrated for 24 Gy single-fraction spine SBRT [23];

E — The 19 Gy x 1 will provide the highest biologically equivalent dose (BED) and equivalent total dose in 2-Gy fractions (EQD2) of any treatment arm (Tab. 1), yet remaining low enough to consistently meet the spinal cord maximum dose (Dmax) constraint of 14 Gy and cauda equina Dmax constraint of 16 Gy for single-fraction spine SBRT (Tab. 2) [24].

Given the frequency of spine involvement in metastatic cancer, the implications of SPORTSMEN are far-reaching as a prospectively determined optimal spine SBRT dose will impact a large number of medical subspecialties in addition to radiation oncology. The short and long-term results of the recent Phase II SABR-COMET randomized trial demonstrating improved durable overall survival for oligometastatic disease treated with SBRT versus palliative standard-of-care only further illustrate the increasing importance of SBRT for metastatic spine disease and, therefore, the necessity of determining the optimal spine SBRT treatment regimen [25, 26]. Treatment arms would be stratified by the following disease demographics: 1 — Intermediate versus radioresistant histology; 2 — presence/absence of epidural disease (for which the Bilsky scale for epidural spinal cord compression will be used [27]); 3 — Baseline opiate use; 4 — Baseline pain score; 5 — Extent of disease; 6 — Remaining lines of standard of care therapy available (0–1 versus 2+).

## **Materials and methods**

### ***Objective(s)***

Primary objective was to evaluate pain freedom from symptomatic metastatic spinal disease at 3 months following treatment (two-fraction standard-of-care spine SBRT, one-

fraction spine SBRT, or EBRT), defined using the Brief Pain Inventory for assessment [9].

Secondary objectives were:

- 6-month freedom from pain, defined as pain freedom at 6-months following treatment per the Brief Pain Inventory;
- 6-month local control, defined as an actuarial 6-month rate of any new, recurrent or progressing (as defined by SPINO criteria) tumor within the planning target volume on any post-treatment MRI by 6 months. Follow-up MRIs will be fused with the planning scan for this assessment
- 6-month vertebral compression fracture rate, assessed by post-treatment MRI by 6 months;
- 6-month overall survival;
- 12-month vertebral compression fracture rate, assessed by post-treatment MRI at 12 months;
- demographic disparities (by race, ethnicity, sex, insurance status) in spine SBRT access and outcomes at 3 months;
- demographic disparities (by race, ethnicity, sex, insurance status) in spine SBRT access and outcomes at 6 months.

### **Study design**

This is a prospective, real-world randomized trial to determine the optimal spine SBRT regimen (no SBRT, one fraction SBRT, or two-fraction SBRT) for achieving pain freedom at 3 months in subjects with a diagnosis of spine metastatic disease from MRI and tissue diagnosis of primary malignancy.

### ***Study design including dose escalation/cohorts***

Subjects will be randomized to standard-of-care SBRT (24 Gy in 2 fractions), one-fraction SBRT (19 Gy in 1 fraction) or conventional EBRT (8 Gy in 1 fraction). Subjects will be randomized in a 2:1 distribution between the SBRT and EBRT arms.

### ***Number of subjects***

Approximately 240 subjects will be enrolled in this trial.

***Replacement of subjects***

If a subject fails to complete SBRT/EBRT as prescribed, they will be replaced.

***Expected duration of treatment and subject participation***

Expected duration of treatment approximately one month including screening CT simulation followed by one (EBRT and one-fraction SBRT) or two (standard-of-care SBRT) days of treatment. Follow up for all patients is twelve months. Thus, the duration of subject participation is thirteen months.

**Eligibility criteria**

***Inclusion criteria***

1. Histologically or cytologically confirmed spinal metastatic disease
2. No previous radiation therapy encompassing the anatomic site to be treated with spine SBRT.
3. Age  $\geq 18$  years. This study requires informed consent by the subject; as children are not able to perform this without parental approval, subjects  $<$  age 18 are excluded from this study.
4. Life expectancy of  $\geq 3$  months, in the opinion of and as documented by the investigator.
5. Subject must have a worst pain score  $\geq 2$  of 10 according to the Brief Pain Inventory
6. Subject must have no intention of changing pain medications on the first day of SBRT
7. Subject must have a Spinal Instability score (SINS)  $\leq 12$
8. Subject must be a spine SBRT candidate per Radiation Oncology
9. Subject must have the ability to understand and the willingness to sign a written informed consent document.

***Inclusion of women and minorities***



Men, women and members of all races and ethnic groups are eligible for this trial. The goals for SPORTSMEN accrual of patients from groups underrepresented in medicine is for racial and ethnic demographics to mirror those of the US Census, which for African-Americans is 12.6% representation [29]. Furthermore, the principal investigator is also the creator of the Navigator-Assisted Hypofractionation (NAVAH) program utilizing patient navigation to increase underrepresented minority patient access to short-course radiation therapy to reduce radiation therapy access disparities; this program will assist in ensuring the SPORTSMEN trial actively recruits a diverse and representative patient population [30, 31].

### **Treatment plan**

The SBRT fractionation schema (comprising 2 of the 3 arms of the study) will be specified as one of the options noted in Table 1. Following randomization, all patients will be treated to their assigned fractionation schema to completion.

Food and Drug Administration (FDA)-approved stereotactic localization procedures for imaging and treatment delivery will be used for linear accelerator based stereotactic localization with an immobilization system, and similarly for proton-based stereotactic localization. The dose will be prescribed to the isodose surface, which encompasses the margin of the metastasis, as defined by the imaging studies. The 100% dose will be recorded for each patient.

For patients with multiple spine metastases, SBRT will be delivered to each lesion that has not previously undergone radiation therapy. The prescribed dose will be according to the SPORTSMEN schema as described in Table 1 above.

If the above constraints cannot be met utilizing the prescribed radiosurgery dose in Table 1, then the highest dose to the target volume will be used such that constraints can be met. This will be considered a minor deviation. The conventional EBRT arm will be treated to 8 Gy x 1.

### ***Radiation therapy***

### *General guidelines and timing*

Spine SBRT will be delivered on all patients utilizing linear accelerator or proton-based techniques as per SPORTSMEN dosing criteria

### *Equipment and techniques to be used*

For SBRT, patients will be treated with linear accelerator. For lesions superior to T4, a facemask will be used; those inferior to T4 will not require a facemask, but will require a Pro-Lok type immobilization system. The most recent MRI is to be fused with the CT simulation prior to contouring.

### *Target volumes*

The GTV is defined as the gross disease visible on pre-treatment MRI (when visible). The CTV is defined as the GTV + microscopic disease on CT simulation; of note, a GTV is not required for all treatment plans, a CTV covering the Radiation Oncologist's clinical judgment of disease extent is sufficient. The PTV is defined as setup error. For SBRT, the  $PTV = CTV + 0.2 \text{ cm}$ . For EBRT, the  $PTV = CTV + 0.7 \text{ cm}$ .

### *Dose to target and organs at risk constraints*

Constraints for organs-at-risk will be as per the CCTG/SC24 trial protocol<sup>8</sup> for patients receiving SBRT. Table 2 lists some of the common spine SBRT dose constraints.

### ***Follow-Up***

Subjects will be followed approximately every 3 months ( $\pm 30$  days) after SBRT/EBRT for 6 months. A detailed medical history, toxicity assessment and physical examination including vital signs will be performed at each visit. Each follow-up over this time period will also include a Spine MRI with and without contrast, which will be analyzed per Spine response assessment in Neuro-Oncology (SPINO) criteria (Section 7) for assessment of local control, and for development of vertebral compression fracture. Neurologic status will be assessed using the Neurologic Assessment in Neuro-Oncology (NANO) scale **(Nayak et al., 2017)**.

After the 6-month follow-up period, subjects will be followed according to their treating physician per standard of care every 3-6 months, with the final MRI obtained at 12 months of follow-up. MRI Spine obtained during this time period may be used for

assessment of primary and secondary endpoints; however, are not mandated to be obtained at particular time intervals.

### **Adverse events and potential risks**

Reported adverse events of spine SBRT include: spine fracture, spinal cord toxicity (myelopathy), esophageal fistula, bowel dysfunction, fatigue, and secondary cancer formation.

### **Study parameters and calendar**

#### ***Study parameters***

The following will be completed prior to spine SBRT:

1. Written informed consent and Health Insurance Portability and Accountability Act (HIPAA) authorization
2. Diagnostic MRI Spine, performed per institutional standard of care
3. Medical history and clinical examination performed by radiation oncology, neurosurgery, medical oncology and/or neuro-oncology
4. Baseline ds-GPA.

#### ***Calendar***

The SPORTSMEN calendar is depicted in Table 3.

#### ***Measurement of efficacy***

Local progression will be defined according to the SPIne response assessment in Neuro-Oncology (SPINO) group and will consist of at least one of the following [31–33]:

1. Gross unequivocal increase in volume or linear dimension.
2. New or progressive tumor in the epidural space.
3. Neurological deterioration attributable to pre-existing epidural disease dimensions specific to the target volume site.

## **Results**

### ***Statistical methods***

#### *General considerations*

The overarching aim of this study is to assess and compare the safety and efficacy of two regimens for treating symptomatic metastatic neoplasms: 24 Gy in 2 fractions and 19 Gy in 1 fraction in the SBRT arm, and 8 Gy in 1 fraction in the EBRT arm. The 24 Gy/2 fx regimen has been validated with Level I evidence [8]. The primary endpoint is 3-month pain-free rates post-treatment. Other safety and efficacy endpoints will be assessed as secondary. The statistical analysis methods are outlined below.

#### *Study design*

This is a phase 2, open-label, randomized control trial with 3 arms. Eligible patients will be randomized with equal probability to standard-of-care SBRT arm (24 Gy in 2 fractions), one-fraction SBRT arm (19 Gy in 1 fraction) versus the EBRT arm (8 Gy in 1 fraction).

#### *Analysis Datasets*

Enrolled population — the enrolled population comprises all subjects who meet the eligibility criteria and are registered onto the study.

Safety population — the safety population comprises all subjects who have received at least one dose of radiation. This set will be used for safety analysis.

Efficacy population — The efficacy population comprises all subjects who have completed SBRT or EBRT. This population will be used for efficacy analysis.

### ***Sample size***

The planned accrual is 240 evaluable patients. We assume the 3-month pain-free rates of the two SBRT arms will be the same. The power analysis is based on our primary hypothesis that the 3-month pain-free rate of the combined SBRT arm is superior to EBRT. Based on literature, we assume that the 3-month pain-free rates are 30% for SBRT and 15% for EBRT. A sample size of 218 patients (64 patients per arm) provides 80% power to detect such an effect size using a 0.05 level, one-sided two-sample proportion

test, while accounting for a formal interim analysis for efficacy. Based on our experience, we aim to enroll up to 60 patients each year at University Hospitals and contributing centers. The patient accrual is expected to be completed within the first two years.

### ***Interim analyses***

For this study, it is planned that the Data Safety and Monitoring Board (DSMB) will review safety data on a regular frequency, e.g., every six months. DSMB may request additional safety data review. Such safety analyses do not inflate the type I error for the primary hypothesis testing and thus require no multiplicity adjustments. No formal interim safety analyses are planned.

We will conduct an interim analysis at 3 months after 50% (or 120) patients complete RT. A one-sided group sequential design will be used at the interim analysis using a Hwang-Shih-DeCani spending function at 50% of the information. The critical p-values (boundaries) for rejecting the null (efficacy) or alternative hypothesis (futility) are 0.0009 and 0.9731 at interim analysis and are 0.0499 and 0.9094 at final analysis. If the test for primary endpoint does not cross the pre-specified boundaries at the interim analysis, a final efficacy and safety analysis will occur after all participants complete the study follow-up. In parallel, we will also compare the 3-month pain-free rates between the two SBRT arms, which requires no multiplicity adjustments.

If the test statistic of the interim analysis for the primary endpoint crosses the pre-specified efficacy boundary, the study will drop the EBRT arm, but will continue enrolling the two SBRT arms. Under the unlikely scenario when one SBRT arm is inferior, we will exclude that SBRT arm with excessive pain-free rate, and continue with the other two arms. Results of the interim analysis will be presented to the DSMB by the study team.

### ***Patient characteristics and significant protocol violations***

Baseline subject characteristics will be tabulated, such as demographics (age, race, gender), and disease characteristics [disease-specific Graded Prognostic Assessment scale (ds-GPA)].

### ***Disposition***

The reasons for patient treatment and study discontinuation will be summarized.

### ***Analysis of primary objectives***

The primary endpoint of 3-month pain-free rate post-SBRT/EBRT will be calculated as the proportion of patients who reported pain freedom at 3 months in the Brief Pain Inventory along with a 95% confidence interval. The calculation will take place in the combined SBRT arm and the EBRT arm. Two-sample proportion test will be used to compare the combined SBRT arm vs. EBRT arm. A two-tailed, multivariable logistic regression model will be used to assess the effect of treatment regimen on pain events, where the outcome is whether a patient is free from pain at 3 months (yes/no), and the independent variables include treatment arm (SBRT vs. EBRT) and baseline covariates for potential efficiency gain. The estimated effect will be reported as odds ratio (OR, SBRT vs. EBRT) with 95% CI.

To further compare the two SBRT regimens, we will calculate the 3-month pain-free rates separately for each SBRT regimen. Two-sample proportion test and two-tailed multivariable logistic regression will be used to compare the two regimens and quantify the effect.

### ***Analysis of secondary objectives***

The 6-month pain-free rate will be analyzed similarly as in Section 12.8. Other secondary endpoints, including local control, vertebral compression fracture, and overall survival will be estimated using the Kaplan-Meier methods, where the time to events will be calculated as the duration from randomization to the corresponding event of interest. Patients without the corresponding event of interest will be censored at their last clinical follow-up. Log-rank test will be used to compare the regimens. Multivariable Cox regression models will be used to quantify the effect of RT regimens on these time-to-event endpoints, while adjusting for baseline covariates as specified in Section 12.8. The aforementioned analyses will be performed to compare 1) combined SBRT vs. EBRT and 2) three-arm comparison. The proportional hazard assumption will be assessed by visualizing and testing Schoenfeld residuals.

To assess the disparities in outcomes, including pain-free rates and other secondary endpoints, descriptive statistics will be summarized stratified by groups defined using race, ethnicity, sex and insurance status. Two-sample tests (proportion tests for rates, and log-rank tests for time-to-event outcomes) or their multi-group extensions will be used for marginal comparisons between these groups. Regression models (logistic regression or Cox proportional hazard model) will be used to quantify the difference between these groups while adjusting for baseline covariates.

## **Discussion**

This is the first phase II to objectively assess optimal spine SBRT dosing in the treatment of symptomatic spine metastatic disease, while assessing the efficacy of spine SBRT versus EBRT. This will also be the first neuro-oncology trial to prospectively target recruitment of underrepresented minorities to ensure participation commensurate with US Census representation.

## **Conclusions**

Findings should allow for better determination of the efficacy of two-fraction spine SBRT versus EBRT in the United States, as well as for the novel single-fraction 19 Gy spine SBRT regimen in patients with symptomatic spine metastases. Furthermore, the commitment of this trial to diversity, equity and inclusion from its inception will allow for increased access to standard-of-care treatment of metastatic spine disease regardless of patient race, ethnicity, gender or socioeconomic status.

## ***Conflicts of interest***

Dr. McClelland serves as a consultant for Gilmartin Capital (a company that evaluates surgically targeted radiation therapies), receives travel funding from GT Medical Technologies Inc., and receives research funding from the University Hospitals Minority Faculty Career Development Award, the ASTRO Emerging Investigator Award, the

Robert Winn Diversity in Clinical Trials Career Development Award, and the National Cancer Institute Paul Calabresi K12 Clinical Oncology Research Career Development Program. Dr. Spratt receives research funding from the National Institutes of Health and the Prostate Cancer Foundation and personal fees from Astellas, AstraZeneca, Bayer, BlueEarth, Boston Scientific, Elekta, GT Medical Technologies Inc., Myovant, Pfizer, Janssen, Novartis, and Varian.

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### ***Author contributions***

Conception and design: S.McC., Y.S., D.E.S.; manuscript writing: S.McC.; study coordination: S.McC.

Participating centers: University Hospitals Seidman Cancer Center Case Western Reserve University School of Medicine, Cleveland, OH 44106

Final approval of manuscript: All authors

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**Table 1.** Dose equivalent measurements of the Spine Patient Optimal Radiosurgery Treatment for Symptomatic Metastatic Neoplasms (SPORTSMEN) spine stereotactic body radiation therapy (SBRT) treatment arms

<b>Spine SBRT regimen</b>	<b>24 Gy in 2 fractions</b>	<b>19 Gy in 1 fraction</b>
BED	52.8 Gy	55.1 Gy
EQD2	44.0 Gy	45.92 Gy
Alpha/Beta ratio	10 Gy	10 Gy

BED — biologically equivalent dose; EQD2 — equivalent total dose in 2-Gy fractions

**Table 2.** Critical structures

	<b>Maximum point dose constraint</b>	<b>D0.1cc constraint</b>
<b>Organ at risk</b>	<b>24 Gy in 2 fractions</b>	<b>19 Gy in 1 fraction</b>
Spinal cord (no PRV)		14 Gy
Spinal cord + 0.2 cm PRV	17 Gy	16 Gy

Cauda equina (no PRV)	17 Gy	16 Gy
Esophagus	20 Gy	16 Gy
Trachea	20 Gy	20 Gy
Pharynx	20 Gy	N/A
Stomach	20 Gy	14 Gy
Rectum	20 Gy	18 Gy
Duodenum		16 Gy
Small bowel	20 Gy	11 Gy
Large bowel (colon)	20 Gy	16 Gy
Kidneys	26 Gy	N/A
Liver	26 Gy	N/A
Sacral nerve roots (to be contoured for tumors involving S1–S5)	26 Gy	18 Gy
Heart		22 Gy
Great vessels		18 Gy
Brachial plexus		17.5 Gy
Brainstem		14 Gy

PRV — planning risk volume; N/A

**Table 3.** The Spine Patient Optimal Radiosurgery Treatment for Symptomatic Metastatic Neoplasms (SPORTSMEN) study calendar

	Baseline Screening	SBRT	3 Month Follow Up <sup>1</sup>	6 Month Follow Up <sup>1</sup>	9 Month Follow Up	12 Month Follow Up <sup>2</sup>
	Within 30 days of SBRT	Linear Accelerator (LINAC) or Proton	90 days post SBRT	180 days post SBRT	270 days post SBRT	360 days post SBRT
Radiation Oncology consult and consent	x					
Medical History (including Brief Pain Inventory score)	x		X	X	X	X
Physical Examination	x		X	X	X	X
Vitals	x	X	X	X	X	X
ds-GPA/ECOG performance status/KPS	x		X	X	X	X
Diagnostic MRI Spine	x					
MRI Spine Planning Scan <sup>4</sup>		X				
Toxicity assessment			X	X	X	X
MRI Spine with and without contrast <sup>3</sup>			X	X	X	X

Footnotes:

1. Variations of +/- 14 days from the scheduled visit are permitted
2. After 12-months post-SBRT, subjects will be followed at physician's discretion, approximately every 3-6 months per standard of care. Any MRI Spine, physical exam or vitals obtained at these appointments will be gathered. However, if these procedures are not performed per standard of care, this will not be a deviation.
3. MRI Spine performed at University Hospitals Cleveland Medical Center will have sequences including contrast, no contrast, FLAIR, DTI and PWI. If patient receives MRI Spine outside of University Hospitals Cleveland Medical Center, a minimum of contrast, no contrast and FLAIR will need to be obtained and all sequences mentioned above are encouraged.
4. Variations of -30 days from the scheduled visit are permitted for SBRT, and may include the baseline screening MRI at the treating radiation oncologist's discretion.