

# May 19, 2023 Meeting Materials Health Technology Clinical Committee

# Stereotactic body radiation therapy

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### **Personal Data**

Name: Simon S. Lo, M.B., Ch.B., FACR, FASTRO

Address:

**Telephone number:** 

Place of birth:

Citizenship:

Date of birth:

Email:

#### Education

M.B., Ch.B. (M.D. equivalent)
Faculty of Medicine
Chinese University of Hong Kong
Shatin, New Territories
Hong Kong
Studies completed: June 1991
Date conferred: December 1991
Medical degree fully accredited by General Medical Council, UK with no examination required
(US News & World Report Top Clinical Medicine Universities in the World 2020: Ranked no. 92)
https://www.usnews.com/education/best-global-universities/clinical-medicine
(QS World University Rankings® 2020: Ranked no. 43)
https://www.topuniversities.com/university-rankings/university-subject-rankings/2020/medicine

### **Postgraduate Training**

<b>Training Position</b>	Department	Institution	Date
Intern	Obstetrics and	Prince of Wales	July 1, 1991-June
	Gynaecology,	Hospital, Hong	30, 1992
	Surgery, Medicine	Kong SAR, China	
	and Paediatrics	_	
Resident	Orthopaedics	Kwong Wah	July 1, 1992-June
		Hospital, Hong	30, 1993
		Kong SAR, China	
Resident	Clinical Oncology	Queen Elizabeth	July 1, 1993-May
	(Royal College of	Hospital, Hong	31, 1997
	Radiologists, UK	Kong SAR, China	
	curriculum and also		
	eligible for FRCP		
	Canada Board		
	Examination)		
Resident	Radiation Oncology	University of	July 1, 1997-June
		Minnesota Medical	30, 2001
		Center,	
		Minneapolis, MN	

American College of Radiation Oncology Fellow in Gastrointestinal Radiation Oncology and IORT (funded by grant)- Away rotation during	Radiation Oncology	Mayo Clinic, Rochester, MN	July 1-August 31, 2000
residency Visiting Fellow (Observer)- Away rotation during residency	Radiation Oncology	Princess Margaret Cancer Centre, University of Toronto, Toronto, ON, Canada	January 1-31, 2001

### **Faculty Positions Held**

<b>Faculty Position</b>	Department	Institution	Date
Assistant Professor of	Radiation	Wayne State	July 1, 2001- June
Radiation Oncology	Oncology	University School of	30, 2002
		Medicine, Detroit, MI	
Assistant Professor of	Radiation	Loyola University	July 15, 2002- June
Radiation Oncology	Oncology	Chicago, Maywood,	30, 2004
		IL	
Assistant Professor of	Radiation	Indiana University	July 1, 2004- June
Radiation Oncology	Oncology	School of Medicine,	23, 2006
		Indianapolis, IN	
Associate Professor of	Radiation	The Ohio State	June 26, 2006-
Radiation Oncology	Oncology	University College of	February 28, 2011
		Medicine, Columbus,	
		OH	
Associate Professor of	Neurosurgery	The Ohio State	June 26, 2006-
Neurosurgery		University College of	February 28, 2011
(Adjunct appointment)		Medicine, Columbus,	
		OH	
Visiting Associate	Radiation	Case Western	March 1, 2011-
Professor of Radiation	Oncology	Reserve University	June 30, 2011
Oncology		School of Medicine,	
		Cleveland, OH	
Associate Professor of	Radiation	Case Western	July 1, 2011-June
Radiation Oncology	Oncology	Reserve University	30, 2015
		School of Medicine,	
		Cleveland, OH	
Professor of Radiation	Radiation	Case Western	July 1, 2015-July
Oncology	Oncology	Reserve University	22, 2016
		School of Medicine,	
		Cleveland, OH	
Professor and Vice-	Radiation	University of	August 1, 2016-
Chair for Strategic	Oncology	Washington School	Present
Planning		of Medicine	

Professor of	Neurological	University of	August 1, 2016-
Neurological Surgery	Surgery	Washington School	Present
		of Medicine	

# **Hospital Positions Held**

Faculty Position	Department	Institution	Date
Attending Radiation	Radiation	Barbara Ann	July 1, 2001- June
Oncologist	Oncology	Karmanos Cancer	30, 2002
		Institute, Detroit, MI	
Attending Radiation	Radiation	Loyola University	July 15, 2002- June
Oncologist	Oncology	Medical Center,	30, 2004
		Maywood, IL	
Director, Medical	Radiation	Loyola University	November 1, 2002-
Student Elective	Oncology	Medical Center,	June 30, 2004
Rotation		Loyola University	
		Chicago, Maywood,	
		IL	
Attending Radiation	Radiation	Indiana University	July 1, 2004- June
Oncologist	Oncology	Cancer Center,	23, 2006
		Indianapolis, IN	
Medical Director	Radiation	Richard L Roudebush	July 1, 2004-
	Oncology	VA Medical Center,	November 30, 2004
		Indianapolis, IN	
Attending Radiation	Radiation	Arthur G. James	June 26, 2006-
Oncologist	Oncology	Cancer Hospital,	February 28, 2011
		Columbus, OH	
Residency Program	Radiation	Arthur G. James	August 1, 2006-
Director	Oncology	Cancer Hospital, The	July 31, 2009
		Ohio State University	
		College of Medicine,	
		Columbus, OH	
Director of Neurologic	Radiation	Arthur G. James	June 26, 2006-
Radiation Oncology	Oncology	Cancer Hospital, The	March 31, 2009
and Stereotactic		Ohio State University	
Radiosurgery		College of Medicine,	
Dimentary	Dediction	Columbus, OH	Lune 26, 2006
Director of Starostastic Dady	Radiation	Arthur G. James	June 26, 2006-
Stereotactic Body	Oncology	Cancer Hospital, The	February 28, 2011
Radiotherapy		Ohio State University	
		College of Medicine,	
Director of Nourclosia	Radiation	Columbus, OH University Hospitals	March 1 2011
Director of Neurologic Radiation Oncology	Oncology	Seidman Cancer	March 1, 2011- July 22, 2016
Radiation Oncology and Gamma Knife	Oncorogy	Center, Cleveland,	July 22, 2010
Radiosurgery		OH	
Director of	Radiation	University Hospitals	January 1, 2012-
Radiosurgery Services	Oncology	Seidman Cancer	July 22, 2016
(Medical service	Oncorogy	Center, Cleveland,	July 22, 2010
		OH	
	1	UH	

agreement with cancer center)			
Attending Radiation	Radiation	University Hospitals	March 1, 2011-July
Oncologist	Oncology	Seidman Cancer	22, 2016
		Center, Cleveland, OH	
Attending Radiation	Radiation	University of	August 1, 2016-
Oncologist	Oncology	Washington Medical	Present
		Center/ Seattle	
		Cancer Care Alliance	
		(SCCA)	
Courtesy Attending	Neurological	Harborview Medical	August 1, 2016-
Physician	Surgery	Center, Seattle, WA	Present
Attending Radiation	Radiation	Seattle Proton	August 1, 2019-
Oncologist	Oncology	Therapy Center	Present
Director of SBRT	Radiation	University of	February 2018-
	Oncology	Washington	Present

## UW Medicine/ Fred Hutchinson Cancer Research Center Positions Held

Position	Committee	Institution	Date
Member	Credential Committee	UWMC	September 2016-Present
Key personnel	Leading Academic	Fred Hutchinson Cancer	March 1, 2019-February 29,
	Participating Site, NCTN	Research Center	2020
Member of Leadership Team	Leading Academic	Fred Hutchinson Cancer	March 1, 2020-Present
	Participating Site, NCTN	Research Center	

### **Department Positions Held**

Position	Committee	Institution	Date
Co-Chair	Appointment and Promotion Committee	UW SOM	2018-Present
Member	Clinical Trial Committee	UW Rad Onc	2020-Present

### Honors

Honor/ Award	Year
Berlex Oncology Foundation:	1998
Epidemiology and Clinical Trial Design	
(Training Workshop)	
Berlex Oncology Foundation: Clinical	1999
Pharmacology of Anticancer Agents	
(Training Workshop)	
American College of Radiation Oncology	2000
Fellowship Grant Award in	
Gastrointestinal Oncology	
Radiological Society of North America	2001
Roentgen Resident/ Fellow Research	
Award	

Teacher Of The Year Award, Department	2002
of Radiation Oncology, Wayne State	2002
University- Elected by the residents in the	
residency program	
Excellence in Teaching Award,	2008
	2008
Department of Radiation Medicine, The	
Ohio State University College of Medicine	2011
Listed in Marquis Who's Who in America	
Awardee for Fellowship of American	2014
College of Radiology (FACR)	2014
Outstanding Reviewer of the Year,	2014
International Journal of Radiation	
Oncology, Biology, and Physics	
Outstanding Reviewer of the Year,	2015
Radiotherapy and Oncology	
Association of Residents in Radiation	2014-2015
Oncology (ARRO) Educator of the Year	
Best Doctors, Inc	2015-2020
Cleveland Magazine, Top Doctor	2015, 2016
Top Doctor, Castle Connolly	2013-2022
Seattle Magazine, Top Doctor	2020, 2021
Faculty Mentor of the Year, Association of	2016
Residents and Fellows, University	
Hospitals Case Medical Center	
Awardee for Fellowship of American	2017
Society for Radiation Oncology (FASTRO)	
Nominee for Board of Trustees of	2017, 2018, 2019
American Board of Radiology	
Elected Member of Board of Directors,	2019-2025 (Re-elected in 2022)
Radiosurgery Society	
American College of Radiology (ACR)	2019, 2020
Chapter Recognition Award for	,
Membership to CARROS (Radiation	
Oncology Chapter of ACR)- Co-leader for	
the award application	
Society for Palliative Radiation Oncology	2019
Lifetime Achievement Award	
UW Radiation Oncology Residency	2019
Teaching Award	

### **Board Certification**

USMLE steps 1, 2 and 3 passed in 1994, 1995, and 1999 ECFMG certified in 1993

Eligible for Royal College of Radiologists, UK Fellowship Examination in Clinical Oncology Certified by American Board of Radiology in Radiation Oncology in June 2001 (Passed written and oral examinations in September 2000 and June 2001, respectively on first attempt) ABR (Written boards- September 2000): 96th percentile for Clinical Oncology (1<sup>st</sup> quartile for all 8 areas) 92th percentile for Radiation Physics 89th percentile for Radiobiology

Recertified May 2011 (Good through December 2021)

Basic Life Support re-certified August 2021 (valid till August 2023)

Advanced Cardiac Life Support certified in August 2016 (valid until August 2020)

Cleveland Clinic Foundation Gamma Knife ICON training October 25-27, 2017

### **Current License(s) to Practice:**

Full registration, General Medical Council, UK, August 1993-Present Unrestricted Minnesota medical license granted in November 1999 (active) Unrestricted Michigan medical license granted in March 2001 (active) Unrestricted Indiana medical license granted in December 2004 (active) Unrestricted Ohio medical license granted in April 2006 (active) Unrestricted Washington medical license granted in March 2016 (active)

### Diversity, Equity and Inclusion Activities (optional): None

### **Professional Organizations**

#### Service on professional societies:

1. American Society for Therapeutic Radiology and Oncology (ASTRO):

Member of the Publications Committee of the Education Council 2007-2011 Member of the Workforce Diversity Subcommittee of the Research Council 2008-2014 Member of Emerging Technology Committee 2013-2015 Member (Recognized expert) of the ASTRO brain metastasis guidelines taskforce 2010-2011 Member (Recognized expert) of the ASTRO bone metastasis guidelines taskforce (Responsible for drafting the guidelines for stereotactic body radiation therapy for spinal metastases together with Dr. Eric L. Chang from M.D. Anderson Cancer Center and Dr. Arjun Sahgal from University of Toronto, Canada) 2010-2011 Member of ASTRO bone metastasis guidelines work group (responsible for updating of current guidelines) 2014-2016 Member of ASTRO Guideline Subcommittee 2014-Present Panelist of ASTRO guidelines for stereotactic body radiation therapy for lung cancer (Invited) 2015-Present Member of ASTRO Payer Relations Subcommittee 2015-Present Member of ASTRO CNS Committee 2013-2016, 2019-Present

Abstract Reviewer for CNS tumors 2013-2017, 2019-Present Abstract Reviewer for Palliative Radiation Oncology 2019-Present SBRT Model Policy Work Group 2019-Present Member of ASTRO Science Education and Program Development Subcommittee 2019-Present Member of ASTRO Education Committee 2019-Present SRS Model Policy Work Group 2020-Present

2. Radiological Society of North America (RSNA):

Member of the Subcommittee in Radiation Oncology & Radiobiology, Committee for the RSNA Scientific Assembly April 2010-November 2013

Abstract Reviewer November 2009-November 2013

Refresher Course Radiation Oncology Track Chair November 2014-December 2019

Member of Oncologic Imaging and Therapies Task Force November 2014-December 2019

Member of AMPPC Subcommittee 2020-Present

3. International Consensus Conference:

Panel Member of International Consensus Conference Bone Metastasis Group 2009-2011 Panel Member of International Consensus Conference Brain Metastasis Group 2009-2011

4. American College of Radiology:

Vice-Chair and Panel Member, June 1, 2009-May 31, 2012 Panel Chair, Starting June 1, 2012-May 31, 2017 American College of Radiology Appropriateness Criteria® Expert Panel on Radiation Oncology-Bone Metastases

Panel Member, June 1, 2013-May 31, 2017 American College of Radiology Appropriateness Criteria® Expert Panel on Radiation Oncology-Brain Metastases

Radiation Oncology Representative, February 1, 2017-May 31, 2019 American College of Radiology Appropriateness Criteria® Neuro 2 Expert Panel ACR AC Management of Vertebral Compression Fractures document

Radiation Oncology Representative, June 1, 2019-Present American College of Radiology Appropriateness Criteria® Expert Panel on Urological Imaging ACR AC Post-Treatment Follow-up and Active Surveillance of Clinically Localized Renal Cell Cancer document

Chair, Program Committee Council of Affiliated Regional Radiation Oncology Societies (CARROS) January 2012-2014 Comments: Responsible for development of in informational sessions for the CARROS program before ASTRO every year

Secretary, Executive Committee Council of Affiliated Regional Radiation Oncology Societies (CARROS) January 2015-September 2016

President-Elect Council of Affiliated Regional Radiation Oncology Societies (CARROS) September 2016-2018

President Council of Affiliated Regional Radiation Oncology Societies (CARROS) September 2018-2020

Immediate Past President and Fellowship Committee Chair Council of Affiliated Regional Radiation Oncology Societies (CARROS) September 2020-October 2022

Member ACR Radiation Oncology Commission 2018-2020

Contributor of questions in pediatric and CNS radiation oncology and medical statistics in the American College of Radiology in-training examination 2006-2014

Member Population Health Management (PHM) Committee for the Patient- and Family-Centered Care (PFCC) Commission May 2018-Present

Member ACR Practice Parameters Committee (2022-Present)

Chair ACR Practice Parameters for Radiation Oncology (2022)

5. Radiation Therapy Oncology Group

Member, Symptom Management Committee March 2012-June 2014

6. NRG Oncology

Member, Cancer Prevention and Control Steering Committee June 2014-Present

Disease site liaison, Functional/Quantitative Imaging Working Group

July 2014-Present

Liaison to the NRG Oncology Lung Steering Committee September 2016-Present

7. Leksell Gamma Knife Society

Leksell Society Gamma Knife Meetings 2012-2017 Abstract Reviewer

8. Elekta International Oligometastasis Consortium

Core Member October 2013-March 2016

Lead for international survey/ consensus guidelines project

9. Radiosurgery Society (RSS)

Member, RSS Meeting Planning Committee 2016-Present

Volunteer, Webinar Development 2015-Present

National Medical Director for Distinction in Practice in SRT Accreditation Program 2017-Present

Board of Directors 4/1/2019-Present

10. American Radium Society

Chair, Bone Metastasis Panel, ACR-ARS Appropriate Use Committee May 2017-May 2020 Consultant, Bone Metastasis Panel, ACR-ARS Appropriate Use Committee May 2020-Present

Member, Brain Metastasis Panel, ACR-ARS Appropriate Use Committee May 2017-Present

11. ICON Gamma Knife Expert Group

Member September 2017- Present

12. American Society of Clinical Oncology

Member, Annual Meeting Education Committee - Central Nervous System Tumors June 2018-Present

13. American College of Radiation Oncology

Member, Development Committee 2018-Present

14. Congress of Neurological Surgeons

Member, AANS/ CNS Brain Metastases Guidelines Taskforce 2016-2018 Member, AANS/ CNS Low Grade Glioma Guidelines Taskforce 2020-Present Lead Writer, AANS/ CNS Low Grade Glioma Radiotherapy Guidelines 2020-Present

15. International Stereotactic Radiosurgery Society

Board Member, Non-CNS SBRT Guidelines 2021-Present

#### Memberships in professional societies:

American Society for Radiation Oncology, 2001- Present Radiological Society of North America, 2001- Present American College of Radiology, 2002- Present American Medical Association, 1997- Present Canadian Association of Radiation Oncology, 2013-Present Radiosurgery Society, 2015-Present American Radium Society, 2016-Present American Society of Clinical Oncology, 2001-Present

### **Teaching Responsibilities**

#### **Visiting Professorships:**

Visiting Professor, Department of Radiation Oncology University of Michigan, Ann Arbor, MI Lecture: "Does the extent of surgery have any impact on the survival of patients with low grade glioma who receive postoperative radiation therapy?" November 2001

Visiting Professor, Department of Radiation Oncology Yale University, New Haven, CT Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" January 2002

Visiting Professor, Division of Radiation Oncology Mayo Clinic, Rochester, MN Lecture: "Does the extent of surgery have any impact on the survival of patients with low grade glioma who receive postoperative radiation therapy?" January 2002

Visiting Professor, Department of Radiation Oncology

Loyola University Medical Center, Maywood, IL

Lecture: "Does the extent of surgery have any impact on the survival of patients with low grade glioma who receive postoperative radiation therapy?" January 2002

Visiting Professor, Department of Radiation Oncology Indiana University, Indianapolis, IN Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" September 12, 2003

Invited Faculty for Osler Institute Radiation Oncology Board Review Course (Pediatric Oncology and Lymphoma/ Leukemia), Louisville, KY, June 2004

Invited Faculty for Osler Institute Radiation Oncology Board Review Course (Pediatric Oncology, Neuro-oncology and Lymphoma/ Leukemia), Louisville, KY, June 2005

Visiting Professor, Department of Radiation Medicine Arthur James Cancer Hospital Ohio State University Comprehensive Cancer Center Lecture: "Stereotactic Radiosurgery With or Without Whole Brain Radiation Therapy for Brain Metastases" September 26, 2005

Visiting Professor, Brain Tumor Institute Cleveland Clinic Foundation Lecture: "The Role of Gamma Knife Radiosurgery (GKR) in the Management of Unresectable Gross Disease or Gross Residual Disease after Surgery in Ependymoma" October 31, 2005

Visiting Professor, Division of Radiation Oncology H. Lee Moffitt Comprehensive Cancer Center, Tampa, FL Lectures: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" and "Stereotactic Body Radiation Therapy- Clinical Applications" November 7, 2005

Visiting Professor, Department of Clinical Oncology Queen Elizabeth Hospital, Hong Kong SAR, China Lecture: "Stereotactic Body Radiation Therapy- Clinical Applications" January 10, 2006

Guest Faculty, Neurosurgical Symposium Sponsored by Asian Congress of Neurological Surgeons Hong Kong Sanatorium, Hong Kong SAR, China Lectures: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" and "The Role of Gamma Knife Radiosurgery (GKR) in the Management of Unresectable Gross Disease or Gross Residual Disease after Surgery in Ependymoma" January 12, 2006

Invited Faculty for Med Prep Radiation Oncology Board Review Course (Pediatric Oncology, Neuro-oncology and Lymphoma/ Leukemia), Louisville, KY, June 2006

Visiting Professor National University Cancer Institute of Singapore June 21-23, 2010

Comments: I was invited to National University Cancer Institute of Singapore to conduct a training course in stereotactic body radiotherapy (SBRT) for all organ sites and will help them set up an operational SBRT program.

Visiting Professor UT Southwestern Medical Center Dallas, TX Lecture: Stereotactic Body Radiation Therapy for Spinal Metastasis June 28, 2010

Visiting Professor University of Minnesota, Minneapolis VA Medical Center Minneapolis, MN Lecture: Stereotactic Body Radiation Therapy for Early Stage Non-Small Cell Lung Cancer July 12, 2010

Visiting Professor UH Case Medical Center Case Western Reserve University Cleveland, OH Lecture: Stereotactic Body Radiation Therapy for Spinal Metastasis July 21, 2010

Visiting Professor Houston Methodist Hospital, Cancer Center and Research Institute Weill Cornell Medical College Houston, TX Lectures: SBRT for spinal metastasis Toxicities associated with SBRT Tutorials given to radiation oncology residents from The Methodist Hospital and Baylor College of Medicine November 9, 2012

Visiting Professor Sunnybrook Health Science Centre Odette Cancer Centre University of Toronto Toronto, ON Canada Lecture: Toxicities associated with SBRT and strategies to mitigate the risk Cancer Research Rounds December 4, 2012 Comments: Odette Cancer Centre is one of the largest radiation oncology programs in the world, treating ~ 500 patients per day in one single department.

Visiting Professor Juravinski Cancer Centre at Hamilton Health Sciences McMaster University Hamilton, ON Canada Lectures: Emerging applications of SBRT (Regional Oncology Rounds) Toxicities associated with SBRT and strategies to mitigate the risk (Radiotherapy-Specific Rounds) Tutorial to residents April 25, 2013

Visiting Professor Cardinal Bernadine Cancer Center Loyola University Medical Center Chicago, IL Lecture: Emerging applications of SBRT December 2, 2013

Fifth Annual Dr. Roger Potish Memorial Visiting Professorship and Lecture University of Minnesota Minneapolis, MN USA Lectures: SBRT for spinal metastasis Tutorials to residents May 9, 2014

Visiting Professor Pamela Youde Nethersole Eastern Hospital (One of the teaching hospitals of University of Hong Kong and the Chinese University of Hong Kong with a residency training program in Clinical Oncology) Hong Kong SAR, China Lectures: Emerging applications of stereotactic body radiotherapy Toxicities associated with SBRT and strategies to mitigate the risk Stereotactic body radiotherapy for spinal metastasis Comments: Symposium and Training in SBRT May 17, 2014

Visiting Professor Tuen Mun Hospital (with a residency training program in Clinical Oncology) Hong Kong SAR, China Intracranial stereotactic radiosurgery May 19, 2014

Visiting Professor London Regional Cancer Program University of Western Ontario London, ON Canada Lecture: Toxicities associated with SBRT and strategies to mitigate the risk Tutorials to residents June 3, 2014

Visiting Professor National University Cancer Institute of Singapore Singapore Lecture: Oligometastasis in 2015 - Fact or Myth? May 15, 2015

Visiting Professor University of Washington Seattle, WA Lecture: Emerging applications of SBRT- Additions to armamentarium against cancer October 28, 2015

Visiting Professor Sunnybrook Odette Cancer Centre University of Toronto Toronto, ON Canada Cancer Research Rounds: Emerging applications of SBRT July 5, 2016

Visiting Professor Sunnybrook Odette Cancer Centre University of Toronto Toronto, ON Canada Rapid Response Radiotherapy Program Lecture: The medical systems in US and Hong Kong July 5, 2016

Visiting Professor/ Guest Lecturer Faculty of Medicine, The Chinese University of Hong Kong Hong Kong Guest lecture: The importance of basic sciences in radiation oncology October 31, 2016

Visiting Professor Division of Radiation Oncology Department of Surgery British Columbia Cancer Agency University of British Columbia Vancouver, BC Canada Lecture: Emerging applications of SBRT- Additions to armamentarium against cancer May 4, 2017

Visiting Professor Department of Radiation Oncology Cleveland Clinic Foundation Cleveland, OH Lecture: Toxicities associated with SBRT and strategies to mitigate the risk October 25, 2017

Visiting Professor Prince of Wales Hospital/ Chinese University of Hong Kong Hong Kong NTEC spine metastasis special multidisciplinary meeting Lecture: Contemporary SBRT for spine metastasis June 29, 2018

Visiting Professor Winship Cancer Institute / Emory University Atlanta, GA Cancer Center Grand Rounds: Serious toxicities from SBRT- How to navigate through the bumpy skies safely Meeting with the radiation oncology residents October 8-9, 2019

Visiting Professor (Virtual amid COVID-19) Department of Radiation Medicine Georgetown University Washington DC Lecture: Strategies to mitigate serious toxicities from SBRT Meeting with radiation oncology residents May 18, 2020

Virtual Visiting Professor Department of Radiation Oncology Allegheny Healthcare Network Lecture: The incorporaton of SBRT into multidisciplinary spine oncology care December 16, 2020

Visiting Professor (Virtual amid COVID-19) Department of Radiation Oncology USC Lecture: Toxicities of Stereotactic Body Radiotherapy- How to navigate through the bumpy skies March 18, 2021

Visiting Professor (Virtual amid COVID-19) Department of Radiation Oncology Indiana University Lecture: Serious toxicities of SBRT June 19, 2021

Visiting Professor (Virtual amid COVID-19) Spine Oncology Group Indiana University Lecture: How to build a multidisciplinary spine oncology program June 19, 2021

Visiting Professor (Virtual amid COVID-19) Queen's University, Kingston, ON, Canada Lecture: How to build a multidisciplinary spine oncology program May 19, 2022

#### Other invited lectures at major academic centers:

Division of Radiation Oncology Mayo Clinic, Rochester, MN Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" August 2000

Division of Radiation Oncology Johns Hopkins Cancer Center, Baltimore, MD Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" October 2000

Department of Human Oncology University of Wisconsin, Madison, WI Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" November 2000

Department of Radiation Oncology Barbara Ann Karmanos Cancer Institute, Detroit, MI Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" November 2000

Department of Radiation Oncology University of Cincinnati, Cincinnati, OH Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" November 2000

Department of Radiation Oncology University of Florida, Gainesville, FL Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" November 2000

Department of Radiation Oncology Thomas Jefferson University Bodine Cancer Center, Philadelphia, PA Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" December 2000

Department of Radiation Oncology Medical University of South Carolina, Charleston, SC Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" December 2000

Department of Radiation Oncology University of Alabama, Birmingham, AB Lecture: "Stereotactic Radiosurgery vs. Fractionated Stereotactic Radiotherapy for Meningiomas" December 2000

#### Grand Rounds and Seminars for Community Physicians:

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds Magruder Hospital Port Clinton, OH September 27, 2011

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds Elyria Mercy Hospital Elyria, OH November 2,, 2011 Stereotactic Radiosurgery for Intracranial Disease Grand Rounds St. John Medical Center Westlake, OH November 11, 2011

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds St. Vincent's Charity Hospital Cleveland, OH November 16, 2011

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds Firelands Medical Center Sandusky, OH January 4, 2012

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds Southwest General Hospital Middleburg Heights, OH January 27, 2012

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds Robinson Memorial Hospital Ravenna, OH February 15, 2012

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds UH Geauga Medical Center Chardon, OH May 8, 2012

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds Summa Barberton Hospital- Parkview (Cancer) Center Barberton, OH August 16, 2012

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds Memorial Hospital Fremont, OH September 27, 2012

Stereotactic Radiosurgery for Intracranial Disease Grand Rounds Christ Hospital Cincinnati, OH September 12, 2013

Modern Advances in Radiotherapy for Brain and Spinal Tumors Grand Rounds Northside Medical Center Youngstown, OH October 17, 2013

Stereotactic Radiosurgery and Stereotactic Body Radiotherapy Grand Rounds Blanchard Valley Medical Center Findlay, OH November 1, 2013

#### Grand Rounds and Seminars For Students, Residents and Fellows:

Monthly oral board preparation sessions for residents Gershenson Radiation Oncology Center Karmanos Cancer Institute Wayne State University, Detroit, MI July 2001 to June 2002

Brain Metastasis- From the Radiotherapeutic Perspective Neurology Grand Round on 3/1/02, Morse Auditorium, Wayne State University, Detroit, MI

Nasopharyngeal Carcinoma- the American, Canadian and Hong Kong perspectives Radiation Oncology Grand Round on 4/23/02 Gershenson Radiation Oncology Center Wayne State University, Detroit, MI

Mock oral board examiner in central nervous system tumors for residents at University of Minnesota on 4/27/02 Department of Therapeutic Radiology-Radiation Oncology University of Minnesota, Minneapolis, MN

Hodgkin's Disease Radiation Oncology Grand Round on 9/27/02 Loyola University Medical Center, Maywood, IL

Parameningeal Rhabdomyosarcoma Surgical Grand Round on 10/23/02 Loyola University Medical Center, Maywood, IL

Plasma Cell Tumors and Multiple Myeloma Radiation Oncology Grand Round on 1/3/03 Loyola University Medical Center, Maywood, IL

Esophageal Cancer Radiation Oncology Grand Round on 3/21/03 Loyola University Medical Center, Maywood, IL Extranodal Lymphoma Radiation Oncology Grand Round on 9/5/03 Loyola University Medical Center, Maywood, IL

Pediatric Sarcoma Radiation Oncology Grand Round on 6/18/04 Loyola University Medical Center, Maywood, IL

Wilm's Tumor Radiation Oncology Didactic Lecture on 10/20/04 Indiana University Medical Center

Rhabdomyosarcoma Radiation Oncology Didactic Lecture on 10/27/04 Indiana University Medical Center

Retinoblastoma Radiation Oncology Didactic Lecture on 11/3/04 Indiana University Medical Center

Hodgkin Lymphoma- Role of Radiation Therapy Cancer Center Grand Round on 3/4/05 Indiana University Medical Center

Role of Stereotactic Radiosurgery in Brain Metastases Radiation Oncology Didactic Lecture on 9/19/05 Indiana University Medical Center

Extranodal Lymphoma Radiation Oncology Didactic Lecture on 12/21/05 Indiana University Medical Center

Central Nervous System Tumors Mock Oral Boards for Residents on 3/1/06 Indiana University Medical Center

Stereotactic Radiosurgery for Pituitary Tumors Endocrinology Grand Rounds on 3/22/06 Indiana University Medical Center

Stereotactic Radiosurgery for Intracranial and Skull Base Tumors Lecture for Gamma Knife Radiosurgery Training Course on 5/17/06 Indiana University Medical Center

Radiation Therapy for Malignant Gliomas Neurosurgery and Neurology Grand Round on 8/3/06 Ohio State University Medical Center

Modern Radiation Techniques And Their Applications To Brain Tumors Neuro-oncology Symposium on 9/16/06 Ohio State University Medical Center Extranodal Lymphoma Didactic lecture for residents on 2/23/07 Ohio State University Medical Center

Plasma Cell Tumors Didactic lecture for residents on 3/2/07 Ohio State University Medical Center

Extracranial Stereotactic Radiotherapy. Lung Cancer Care for 21<sup>st</sup> Century on 3/3/07 Ohio State University Medical Center

Stereotactic Radiosurgery for Acoustic Neuroma Neurosurgery and Neurology Grand Round on 4/5/07 Ohio State University Medical Center

Adult and pediatric brain tumors 2 lectures for radiation therapy students October 2007

Pediatric cancer 2 lectures for radiation therapy students November 2007

Didactic lecture series on CNS tumors including radiosurgery for residents Weekly Starting January 2008

Stereotactic body radiation therapy for medically inoperable non-small cell lung cancer November 23, 2009

Imaging Solutions in Cancer Management: A Case-Based Conference UH Seidman Cancer Center Case Western Reserve University September 8, 2011

Local Aggressive Therapy for Oligometastases Cancer Center Grand Rounds UH Seidman Cancer Center Case Western Reserve University October 26 ,2011

Gamma Knife Radiosurgery Neuroscience Grand Rounds UH Case Medical Center Case Western Reserve University November 4, 2011

Serious Toxicities Associated with Stereotactic Body Radiation Therapy Radiation Oncology Grand Rounds UH Seidman Cancer Center Case Western Reserve University October 16, 2012

Mock oral boards for residents (CNS tumors) UH Seidman Cancer Center Case Western Reserve University May 31, 2013

Stereotactic Body Radiotherapy for Spinal Metastasis Lecture for Residents and Students UH Seidman Cancer Center Case Western Reserve University August 7, 2013

Stereotactic Body Radiotherapy for Oligometastases Lecture for Residents and Students UH Seidman Cancer Center Case Western Reserve University September 11, 2013

Emerging Applications of Stereotactic Ablative Radiotherapy Radiation Oncology Grand Rounds UH Seidman Cancer Center Case Western Reserve University December 17, 2013

Does conventional wisdom hold for SRS/ SBRT? Lecture for Residents and Students UH Seidman Cancer Center Case Western Reserve University May 7, 2014

Spinal SBRT- Practical Essentials Radiation Oncology Resident Teaching Conference UH Seidman Cancer Center Case Western Reserve University February 4, 2015

Stereotactic Radiosurgery (Part 1) Lecture for neurosurgery residents UH Seidman Cancer Center Case Western Reserve University March 4, 2015

Stereotactic Radiosurgery (Part 2) Lecture for neurosurgery residents UH Seidman Cancer Center Case Western Reserve University May 20, 2015

Strategies to guide safe and effective delivery of spine SBRT Teaching session for radiation oncology residents UH Seidman Cancer Center Case Western Reserve University February 18, 2016

Glioblastoma Teaching session for radiation oncology residents UH Seidman Cancer Center Case Western Reserve University February 25, 2016

Meningioma Teaching session for radiation oncology residents UH Seidman Cancer Center Case Western Reserve University March 3, 2016

AVM and Trigeminal Neuralgia Teaching session for radiation oncology residents UH Seidman Cancer Center Case Western Reserve University March 10, 2016

SBRT for spinal metastases Teaching session for radiation oncology residents University of Washington School of Medicine September 7, 2016

Radiotherapy for uncomplicated bone metastases Teaching session for radiation oncology residents University of Washington School of Medicine March 31, 2017

Stereotactic Body Radiotherapy- Additions to the Armamentarium against Spinal Metastases University of Washington Neurosurgery Grand Rounds University of Washington School of Medicine July 5, 2017

Stereotactic body radiotherapy for spinal metastasis – Target delineation, challenges in response assessment and SPINO University of Washington Radiation Oncology Grand Rounds University of Washington School of Medicine July 26, 2017

Brain metastasis Teaching session for radiation oncology residents University of Washington School of Medicine August 8, 2018

Spinal metastasis Teaching session for radiation oncology residents University of Washington School of Medicine August 24, 2018

Serious complications from SBRT- A word of caution

University of Washington Radiation Oncology Grand Rounds University of Washington School of Medicine September 26, 2018

Sacral SBRT CTV e-contouring and workshop for residents UW residents teaching session June 21, 2019

SBRT for spinal metastasis UW residents teaching session July 30, 2019

Primary brain glioma UW residents teaching session August 16, 2019

Glioma UW residents teaching session 8/14/2020

SBRT for spinal metastasis UW residents teaching session 8/18/2020

Econtouring for spinal metastasis UW residents teaching session 8/21/2020

Spinal metastasis and spinal cord compression UW residents teaching session August 2021

Spinal metastasis and spinal cord compression UW residents case conference August 2021

#### **Residents' Teaching Sessions:**

Regular attendance of teaching conferences for medical students and residents at Wayne State University, Loyola University Medical Center, Indiana University Medical Center, Arthur G. James Cancer Hospital (The Ohio State University), University Hospitals Seidman Cancer Center/ Case Western Reserve University and University of Washington Medical Center.

#### Clinical

Contact hours-40-50 hours per week for residents Variable for medical students and fellows

C Michael Wilkinson, M.D. Resident 7/01-6/02 Barbara Ann Karmanos Cancer Institute Detroit, MI Currently in private practice in Grand Rapids

Faheem Ahmad, M.D. Resident 7/01-6/02 Barbara Ann Karmanos Cancer Institute Detroit, MI Currently faculty at Medical University of Ohio, Toledo, OH

Tanya Powell, M.D. Resident 7/01-6/02 Barbara Ann Karmanos Cancer Institute Detroit, MI Currently in private practice in Chicago

Sameer Keole, M.D. Resident 7/01-6/02 Barbara Ann Karmanos Cancer Institute Detroit, MI Currently faculty at Mayo Clinic, Scottsdale, AZ

David Hsu, M.D. Resident 7/02-6/03 Loyola University Medical Center Maywood, IL Currently in private practice

Robert Prock, M.D. Resident 7/02-6/04 Loyola University Medical Center Maywood, IL Currently in private practice in Indiana

Richard Garza, M.D. Resident 7/02-6/04 Loyola University Medical Center Maywood, IL Currently faculty at Loyola University Medical Center

Paul Crossan, M.D. Resident 7/02-6/04 Loyola University Medical Center Maywood, IL Currently in private practice in Indiana Rajanish Singla, M.D. Resident 7/02-6/04 Loyola University Medical Center Maywood, IL Currently in private practice

Suneel Nagda, M.D. Resident 7/02-6/04 Loyola University Medical Center Maywood, IL Currently faculty at University of Pennsylvannia

Brent Tinnel, M.D. Resident 7/04-6/06 Indiana University Medical Center Indianapolis, IN Currently faculty ay Madigan Army Medical Ctr, Seattle, WA

Achilles Fakiris, M.D. Resident 7/04-6/06 Indiana University Medical Center Indianapolis, IN Currently faculty at University of North Carolina, Chapel Hill, NC

Ramzi Abdulrahman, M.D. Resident 7/04-6/05 Indiana University Medical Center Indianapolis, IN Currently faculty at UT Southwestern Med Ctr, Dallas, TX

David Hoopes, M.D. Resident 7/04-6/06 Indiana University Medical Center Indianapolis, IN Currently faculty ay Wright-Patterson Air Force Med Ctr, Dayton, OH

David Shaeffer, M.D. Resident 7/04-6/05 Indiana University Medical Center Indianapolis, IN Currently in private practice

Bedatri Sinha, M.D. Resident 7/04-6/06 Indiana University Medical Center Indianapolis, IN Currently in private practice

James W. Clarke, M.D. Resident 7/06-9/06 7/07-9/07 Arthur G. James Cancer Hospital Columbus, OH Currently in private practice in Utah

Moataz El-Ghamry, M.D. Resident 1/1/07-2/28/07 Arthur G. James Cancer Hospital Columbus, OH Currently faculty at University of Louisville, Louisville, KY

Granger Scruggs, M.D. Resident 10/1/06-12/31/06 Arthur G. James Cancer Hospital Columbus, OH Currently in private practice in Dallas

Andrew Figura, M.D. Resident 4/1/07-6/30/07 Arthur G. James Cancer Hospital Columbus, OH Currently in private practice in Erie, PA

Timothy Korytko, M.D. Resident 10/1/07-12/31/07 Arthur G. James Cancer Hospital Columbus, OH Currently in private practice in Wisconsin

Jeffrey Radawski, M.D. Resident 10/09-12/09 Arthur G. James Cancer Hospital Columbus, OH Currently in private practice in Kalamazoo, MI

Michael Guiou, M.D. Resident 1/10-3/10 Arthur G. James Cancer Hospital Columbus, OH Currently faculty at The Ohio State University, Columbus, OH

Allison Quick, M.D. Resident 4/08-6/08 Arthur G. James Cancer Hospital Columbus, OH Currently faculty at The Ohio State University, Columbus, OH

Mersiha Hadziahmetovic, MD Resident 5/09-6/09 Arthur G. James Cancer Hospital Columbus, OH First job- Faculty at Vanderbilt University (Lead head and neck radiation oncologist)

Nicholas Galanopoulos, M.D. Resident 4/11-6/11 UH Seidman Cancer Ctr, Case Western Reserve University Cleveland, OH Currently in private practice in Atlanta, GA

Charles Woods, M.D. Resident 7/11-9/11 UH Seidman Cancer Ctr, Case Western Reserve University Cleveland, OH Currently in private practice in Atlanta, GA

Anton Khouri, M.D. Resident 9/11-12/11, 9/13-10/13 Currently in private practice in Cincinnati, OH

Charlene Kan, M.D., Ph.D. Resident 7/12-9/12 10/15-12/15 UH Seidman Cancer Ctr, Case Western Reserve University Cleveland, OH Currently the Chief of Radiation Oncology at Cleveland VA

Christian Okoye, M.D. Resident 4/13-6/13 UH Seidman Cancer Ctr, Case Western Reserve University Cleveland, OH Currently in private practice

Ravi Patel, M.D. Resident 7/13-8/13 7/15-9/15 UH Seidman Cancer Ctr, Case Western Reserve University Cleveland, OH Currently Research Fellow at University of Wisconsin

Mazen Mislmani, M.D. Resident 3/12-4/12, 11/13-12/13 UH Seidman Cancer Ctr, Case Western Reserve University Cleveland, OH Currently in private practice in Kalamazoo, MI

Bryan Traughber, M.D. Resident 1/13-3/13 1/16-Present UH Seidman Cancer Ctr, Case Western Reserve University Cleveland, OH Currently on faculty at Cleveland VA

Mussadiq Awan, MD Resident 7/14-9/14 UH Seidman Cancer Ctr, Case Western Reserve University Cleveland, OH Currently faculty at Medical College of Wisconsin

Thomas Mullen, MD Resident 1/17-2/17 University of Washington Medical Center Seattle, WA Currently in private practice in Oregon

Matthew Spraker, MD, PhD Resident 7/17-9/17 University of Washington Medical Center Seattle, WA Currently faculty at Washington University

Stephanie Schaub, MD Resident 1/18-3/18 University of Washington Medical Center Seattle, WA

Samuel Day, MD, PhD Resident 7/18-9/18 University of Washington Medical Center Seattle, WA

Aileen Kim, MD Resident 10-12/18 University of Washington Medical Center Seattle, WA

Amber Post, MD Resident July 1-August 15, 2019 University of Washington Medical Center Seattle, WA

Matthew Greer, MD Resident January-March 2020 University of Washington Medical Center Seattle, WA

Khang Dinh, MD Resident 2020 University of Washington Medical Center Seattle, WA

Bory Eastman, MD Resident April 2021 University of Washington Medical Center Seattle, WA

August Anderson, MD Resident July-August 2021 University of Washington Medical Center Seattle, WA

Sasha Swensen, MD Resident September-October 2021 University of Washington Medical Center Seattle, WA

Macklin Nguyen, MD Resident December 2021-February 2022 University of Washington Medical Center Seattle, WA

Peter Goff, MD Resident April 2021 University of Washington Medical Center Seattle, WA

#### **Clinical Fellows:**

#### UH Seidman Cancer Center, Case Western Reserve University

Anton Khouri, MD SRS/ SBRT Fellow 11/14-6/15 UH Seidman Cancer Center, Case Western Reserve University Cleveland, OH

Tai-Chung Lam, MB, BS, FRCR Visiting Fellow 3/31/2014-4/4/2014 Dana-Farber Cancer Institute/ Brigham & Women Hospital Harvard Medical School Boston, MA

Kin-Chung Lee, MB, ChB, FRCR Visiting Fellow 9/18/2014-9/30/2014 Pamela Youde Nethersole Eastern Hospital Hong Kong

Jacky Li, MB, BS, FRCR Visiting Fellow 3/1/2015-3/31/2015 Queen Elizabeth Hospital Hong Kong

#### University of Washington

Dennis Leung, MB, BS, FRCR Visiting fellow 1/17-2/17 University of Hong Kong

Ka-Man Cheung, MB, ChB, FRCR Visiting fellow 5/17-6/17 Queen Elizabeth Hospital, Hong Kong

Leslie Fok, MB, ChB, FRCR Visiting fellow 9/18 Queen Elizabeth Hospital Hong Kong

Luke Lee, MB, BS, FRCR

Visiting fellow 10/18 Queen Elizabeth Hospital Hong Kong

Jane Cho, MD SBRT Fellow University of Washington Medical Center Seattle, WA

#### **Medical Student Mentorship:**

#### Wayne State University

Daniel Chang Wayne State University Medical Student One month in 2002 Currently Professor at Stanford University

#### Loyola University

Michelle Mierzwa University of Cincinnati Medical Student One month in 2002 Currently on faculty at University of Michigan

Amit Bhatt Southern Illinois University One month in 2003 Graduated from Northwestern University Radiation Oncology Program

Angela Babbo Rush University One month in 2003 Graduated from Northwestern University Radiation Oncology Program

#### **Ohio State University**

Katherine Tsao OSU Medical Student, Class 2007 One month in 2006 Currently attending at Mayo Clinic, Jacksonville

Mersiha Hadziahmetovic OSU Medical Student, Class 2007 One month in 2006 Resident 5/09-3/11 Arthur G. James Cancer Hospital Columbus, OH Currently faculty at Case Western Reserve University Eugene Hong OSU Medical Student, Class 2008 Columbus, OH Currently in private practice

Steven Register OSU Medical Student, Class 2008 One month in 2007 Currently in private practice

David Chang University of Louisville Medical Student, Class 2008 One month in 2007 Currently in private practice

Michael Crotty Medical University of Ohio Medical Student, Class 2008 One month in 2007 Currently in private practice

William A. Hall Loyola University School of Medicine Medical Student, Class 2009 Currently faculty at Medical College of Wisconsin

Michael K. Cheung Medical University of Ohio Medical Student, Class 2010 Currently faculty at Los Angeles VA

#### **Case Western Reserve University**

Kevin Shuie Medical student Case Western Reserve University Current radiation oncology resident at Indiana University

Brent Cameron, Ph.D. Medical student Case Western Reserve University Current radiation oncology resident at Vanderbilt University

Lindsay Hwang Medical student Case Western Reserve University Current radiation oncology resident at University of Southern California

Shaakir Hasaan Medical student Nova School of Osteopathic Medicine Current radiation oncology resident at Allegheny General Hospital

Sarah Zakem Medical student Case Western Reserve University Current radiation oncology resident at University of Colorado

Nicholas Damico Medical student Case Western Reserve University Current radiation oncology resident at University Hospitals Seidman Cancer Center

Scott Chen Medical student Case Western Reserve University Current radiation oncology resident at Johns Hopkins University

Prachi Jain, M.D. 9/11 Internal Medicine Resident UH Case Medical Center Case Western Reserve University Currently resident at Hofstra School of Medicine, Long Island, New York

Andrew Song Medical student Case Western Reserve University Current radiation oncology resident at Thomas Jefferson University

#### University of Washington

Samuel Kosydar University of Washington MS3

#### **Research Mentorship:**

RT vs. chemoRT for cervical cancer with periaortic nodal metastasis Ayman Saad, M.D., Barbara Ann Karmanos Cancer Institute Outcome: One full manuscripts published

Prostascint for postoperative prostate cancer patient receiving external beam radiotherapy Suneel Nagda, M.D., Loyola University Medical Center Outcome: One full manuscripts published

Invasion of inferior vena cava by neuroblastoma Suneel Nagda, M.D., Loyola University Medical Center Outcome: One full manuscripts published

Stereotactic radiosurgery for secretory pituitary adenoma Brent Tinnel, M.D., Indiana University Outcome: One full manuscripts published

Stereotactic radiosurgery for ependymoma Ramzi Abdulrahman, M.D., Indiana University Outcome: One full manuscripts published Stereotactic radiosurgery for uveal melanoma Achilles Fakiris, M.D., Indiana University Outcome: One full manuscripts published

Stereotactic radiosurgery for macular degeneration Mark Henderson, M.D., Indiana University Outcome: One full manuscripts published

Stereotactic radiosurgery for low grade glioma Mark Henderson, M.D., Indiana University Outcome: One full manuscripts published

P-32 for craniopharyngioma R Bryan Barriger, M.D., Indiana University Outcome: One full manuscripts published

PET and SBRT for non-small cell lung cancer David Hoopes, M.D., Indiana University Outcome: One full manuscripts published

Pathologic complete response of melanoma brain metastasis after SRS Steven Register, M.D. (OSU medical student) Outcome: One full manuscripts published

Stereotactic radiosurgery for radioresistant brain metastases James Clarke, M.D., Ohio State University Outcome: 3 full manuscripts published

Stereotactic radiosurgery for breast cancer brain metastasis Jeffrey Radawski, M.D., Ohio State University Outcome: One full manuscript published

Stereotactic body radiotherapy for adrenal metastasis Michael Guiou, M.D., Ohio State University Outcome: One full manuscript published

Application for Radiological Society of America Medical Student Grant Title: Gamma Knife Radiosurgery For Breast Cancer Brain Metastasis- Impact of HER-2 Status February 2012 Andrew Song, 1<sup>st</sup> year medical student Case Western Reserve University School of Medicine Outcome: Not funded

Ray Tracing vs. Monte Carlo algorithm for SBRT for thoracic spinal metastasis Mentee: Christian Okoye, M.D. (Resident at Case Western Reserve University) Outcome: ASTRO presentation in 2013 and manuscript has been published in a PUBMED indexed journal

CTSC Pilot Grant (Application in progress) RFA and SBRT for liver metastasis Mentee: Ravi Patel, M.D., Ph.D. (Resident at Case Western Reserve University) Consensus guidelines for stereotactic body radiotherapy for renal cell carcinoma Presented in ASTRO 2015 and published in Future Oncology Mentee: Shankar Siva, MB, BS, FRANZCR, Physician-Scientist, Peter MacCallum Cancer Centre, University of Melbourne, Australia

Consensus guidelines for stereotactic body radiotherapy for head and neck cancer Two abstracts accepted for poster presentation in ASTRO 2016 and published in Future Oncology Mentee: Irene Karam, M.D., FRCPC, Assistant Professor of Radiation Oncology, Sunnybrook Health Science Centre, University of Toronto, Canada

Consensus guidelines for stereotactic body radiotherapy for gynecologic cancer Mentee: Eric Leung, M.D., FRCPC, Assistant Professor of Radiation Oncology, Sunnybrook Health Science Centre, University of Toronto, Canada

Consensus guidelines for CTV delineation for stereotactic body radiotherapy for sacral metastasis Mentee: Emma Dunne, MB, BS, FRCR, British Columbia Cancer Agency

Meta-analysis of SRS and HSRT for uveal melanoma Mentee: Samuel Kosydar (MD candidate 2021, UWSOM) Independent Investigative Inquiry, UWSOM

#### **Undergraduate Mentorship**

Jacob Gardner Duke University graduate 2015-Current Comments: I have provided guidance and facilitated his plan to go to Cambodia to help build the nation's first ever cancer center. With the support from a grant, he has worked in Cambodia for over a year.

#### **Other Mentorship**

I have mentored Dr. Mark Henderson (Indiana University resident) in the preparation of a review paper in SRS and SRT for uveal melanoma.

I have mentored Dr. James Clarke (OSU resident) in the preparation of two book chapters and 2 review papers in CNS tumors.

I have mentored Dr. Allison Quick (OSU resident) in the preparation of one book chapter in meningioma in the new radiation oncology textbook "Decision Making in Radiation Oncology (Springer)" and one review paper on radiotherapy for liver tumors.

I have mentored Dr. Mersiha Hadziahmetovic (OSU resident) in the preparation of 3 review papers on medulloblastoma, CNS germ cell tumors and SBRT for lung cancer, respectively.

I have mentored Andrew Song and Kevin Shiue, both CWRU medical students on the writing of a review article, resulting in a two peer-reviewed papers:

Andrew Song, A.B., Kevin Shiue, B.Sc., Mitchell Machtay, M.D., Min Yao, M.D., Ph.D., Rodney J. Ellis, M.D., Zhibin Huang, Ph.D., Nina A. Mayr, M.D., Bin S. Teh, M.D., Simon S. Lo, M.D. Stereotactic body radiation therapy for metastasis in the lung: An undervalued treatment option with future prospects. Lung Cancer Management. Accepted for publication in the June 2012 issue.

Kevin Shiue, B.S., Andrew Song, A.B., Bin S. Teh, M.D., Rodney J. Ellis, M.D., Min Yao, M.D., Ph.D., Nina A. Mayr, M.D., Zhibin Huang, Ph.D., Jason Sohn, Ph.D., Mitchell Machtay, M.D., Simon S. Lo, M.D. Stereotactic body radiation therapy (SBRT) for metastasis to the adrenal gland. Expert Rev Anticancer Ther. In press.\* (Impact factor ~ 2.7)

I mentored Dr. Christian Okoye, M.D., a UHSCC/ CWRU resident in the preparation of a book chapter on pediatric stereotactic radiosurgery (SRS) in a comprehensive textbook "Intracranial SRS".

#### **Recent CME:**

Weekly neuro-oncology tumor board at UH March 2011-Present (with CME credits)

Monthly Departmental Grand Rounds at UH 2012-Present (with CME credits)

Annual Meeting of American Society for Radiation Oncology (ASTRO) 2007-2019 (with CME credits)

Annual Meeting of Radiological Society of North America (RSNA) 2009-2018 (with CME credits)

Radiosurgery Society Meeting 2014, 2019 (with CME credits)

Annual Meeting of American College of Radiology 2015, 2019 (with CME credits)

Annual Continued Risk Management Education at UH 4 hours per year 2011-2016

## **Editorial Responsibilities**

### Editorships

Editor, Discovery Medicine (Partnership with Johns Hopkins School of Medicine for CME)

Associate Editor-in-chief, Journal of Radiation Oncology, 2016-Present

Associate Senior Editor, Advances in Radiation Oncology, 2017-Present

Associate Editor, Neurosurgery, 2017-Present

Associate Editor-in-Chief, Annals of Palliative Medicine

Editor, Cancers

Editor, Frontiers in Oncology

## **Editorial boards**

Member of Editorial Board, Clinical Medicine: Case Reports

Member of Editorial Board, Rare Tumors (Italy)

Member of Editorial Board, Expert Review of Anticancer Therapy (UK)

Member of Editorial Board, Journal of Radiation Oncology (USA)

Member of Editorial Board, Cancers (Switzerland)

Member of Editorial Board, CNS Oncology (UK)

Member of Editorial Board, Cureus (USA)

Member of Editorial Board, Future Oncology (UK)

Member of Editorial Board, Annals of Palliative Medicine

Expert Ambassador, Oncology Central (UK)

Honorary Advisor, Hong Kong Journal of Radiology (Official journal of Hong Kong College of Radiologists)

Member of Editorial Board, Journal of Radiosurgery and SBRT (USA)

Member of Editorial Board, Cancers (Switzerland)

## **Special National Responsibilities:**

#### National examination board:

American Board of Radiology (Radiation Oncology)-

Contributor of questions in pediatric CNS for the written part of the American Board of Radiology examination (2009-2011)

Reviewer of SAM questions for recertification for the American Board of Radiology, starting 2009 Committee member of CNS/ pediatric section, American Board of Radiology, February 2015-Present (Responsible for creating written examination items for initial certification and maintenance of certification and oral board examination) Oral board examiner, CNS/ pediatric, 2016, 2018, 2019 and 2021 Examination creation panel for CNS/ pediatric, January 2018-Present

#### **External grant reviewer:**

Regular external grant reviewer, Associazione Italiana per la Ricerca sul Cancro (A.I.R.C.) (Italian Association for Cancer Research)- It is one of the largest funding bodies for cancer research and is providing 50% of research money in cancer in Italy February 2012-Present Website: www.airc.it

External grant reviewer, Juravinski Cancer Centre Foundation, McMaster University, Hamilton, Ontario, Canada May 2014

External grant reviewer, Cancer Research UK, United Kingdom August/ September 2014

Regular external grant reviewer, The Food and Health Bureau (FHB) of The Government of the HKSAR May 2015-Present

External grant reviewer, Cancer Research UK, United Kingdom July/ August 2018

External grant reviewer, Natural Sciences and Engineering Research Council of Canada (NSERC) November/ December 2018

Reviewer, NRG Oncology NCORP Pilot Grants May 2019

External grant reviewer 'Highly Specialised Care & Research' programme The Netherlands Organisation for Health, Research and Development (ZonMw) December 2019

External grant reviewer Palliantie Meer dan zorg The Netherlands Organisation for Health, Research and Development (ZonMw) March 2021

## **Special Local Responsibilities**

University of Minnesota Radiation Oncology Residency Program: Resident Representative (July 1998- June 1999) Chief Resident (July 1999- June 2000)) Chief Resident (October- December 2000 and April- June 2001) Committee Member of the Education Committee (July 1998- June 2001) Committee Member of the Resident Selection Committee (July 1998- June 2001)

Barbara Ann Karmanos Cancer Institute/ Wayne State University: Member, Protocol Review Committee (July 2001-June 2002)

The Ohio State University Department of Radiation Medicine Residency Program Director July 1, 2006-September 2009

Arthur G. James Cancer Hospital Director of SRS and SBRT June 2006-February 2011

Ohio State University Cancer Institutional Review Board Committee Member January 2007-February 2011

The Ohio State University Health System Physician Executive Council July 1, 2008-February 2011

University Hospitals Case Medical Center Radiation Safety Committee July 2011-Present

Case Comprehensive Cancer Center Data Safety and Toxicity Committee January 2012-September 2013

University Hospitals Seidman Cancer Center Radiation Oncology Residency Clinical competency committee September 2013-Present

University Hospitals Seidman Cancer Center Director of Neurologic Radiation Oncology and Radiosurgery Services January 2012- July 2016

University of Washington School of Medicine Department of Radiation Oncology Co-Chair, Appointment and Promotion Committee 2017-Present

UW Medicine Department of Radiation Oncology Member, Credentialing Committee 2016-Present

University of Washington School of Medicine Department of Radiation Oncology Co-Chair, Proton Faculty Search Committee 2018-Present

University of Washington School of Medicine Department of Radiation Oncology Co-Chair, Gynecologic Radiation Oncology Faculty Search Committee 2018-Present

University of Washington School of Medicine Department of Radiation Oncology Director of Stereotactic Body Radiotherapy February 2018-Present

Key Personnel Fred Hutchinson Cancer Research Center Hutchinson Center as Lead Academic Participating Site (UG1 grant) March 1, 2019-February 29, 2020

Member of Leadership Team Fred Hutchinson Cancer Research Center Hutchinson Center as Lead Academic Participating Site (UG1 grant) March 1, 2020-Present

## **Research Protocols**

#### Case Western Reserve University (March 1, 2011-July 22, 2016)

- SHSC 1312- STEREOTACTIC RADIOSURGERY (SRS) +/- WHOLE-BRAIN RADIOTHERAPY (WBRT) FOR THE TREATMENT OF BRAIN METASTASES: PATIENT PREFERENCE STUDY Institutional PI: Simon S. Lo, M.D.
   Study PI : Edward Chow, MB,BS, FRCPC, Sunnybrook Health Science Centre, University of Toronto Funding: Internal
- N107C- A Phase III Trial of Post-Surgical Stereotactic Radiosurgery (SRS) Compared with Whole Brain Radiotherapy (WBRT) for Resected Metastatic Brain Disease Institutional PI: Simon S. Lo, M.D. Study PI: Paul D. Brown, M.D., M.D. Anderson Cancer Center Funding: NRG Oncology Number of patients accrued: 2 (6 screened)
- RTOG 0933- A PHASE II TRIAL OF HIPPOCAMPAL AVOIDANCE DURING WHOLE BRAIN RADIOTHERAPY FOR BRAIN METASTASES Institutional PI: Simon S. Lo, M.D. Study PI : Vinai Gondi, M.D., Chicago Funding: Radiation Therapy Oncology Group (RTOG) Number of patients accrued: 4
- RTOG 1205- Randomized Phase II Trial of Concurrent Bevacizumab and Re-Irradiation Versus Bevacizumab Alone as Treatment for Recurrent Glioblastoma Institutional PI: Simon S. Lo, M.D.
   Study PI: Christina Tsien, M.D., FRCPC Funding: Radiation Therapy Oncology Group (RTOG) Number of patients accrued: 0
- RTOG 1119- Phase II Randomized Study of Whole Brain Radiotherapy in Combination With Concurrent Lapatinib in Patients With Brain Metastasis From HER2-Positive Breast Cancer: A Collaborative Study of RTOG and KROG Study PI: In Ah Kim, M.D., Seoul National University Funding: Radiation Therapy Oncology Group (RTOG) Number of patients accrued: 0
- CHRV0081- Database of Patients Undergoing Stereotactic Body Radiation Therapy Study PI: Simon S. Lo, M.D. Outcomes: One manuscript published and two submitted
- Case 8810- A Phase 1 Study of Carboplatin and Gemcitabine Chemotherapy and Stereotactic Body Radiosurgery for the Palliative Treatment of Persistent or Recurrent Gynecologic Cancer Co-investigator: Simon S. Lo, M.D. Study PI: Rodney J. Ellis, M.D. Funding: Internal
- 8. RTOG 0436- Paclitaxel, Cisplatin, and Radiation Therapy With or Without Cetuximab in Treating Patients With Locally Advanced Esophageal Cancer

Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Mohan Suntharalingam, M.D., University of Maryland Funding: RTOG

- RTOG 0539- Phase II Trial of Observation for Low-Risk Meningiomas and of Radiotherapy for Intermediate- and High-Risk Meningiomas Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Leland Rogers, M.D., Gamma West, Utah Funding: RTOG
- RTOG 0913- Phase I/II Trial of Concurrent RAD001 (Everolimus) With Temozolomide/Radiation Followed by Adjuvant RAD001/Temozolomide in Newly Diagnosed Glioblastoma Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Leland Rogers, M.D., Gamma West, Utah Funding: RTOG
- 11. RTOG 0938- A Randomized Phase II Trial Of Hypofractionated Radiotherapy For Favorable Risk Prostate Cancer-RTOG CCOP Study Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Himu Lukka, M.D., McMaster University, Hamilton, Canada Funding: RTOG
- 12. RTOG 1012- Phase II Randomized Trial of Prophylactic Manuka Honey for the Reduction of Chemoradiation Therapy Induced Esophagitis-Related Pain During the Treatment of Lung Cancer - RTOG CCOP Study Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Lawrence Berk, M.D., Tampa General Hospital Funding: RTOG
- ACRIN6684- Multicenter, Phase II Assessment of Tumor Hypoxia in Glioblastoma Using 18F-Fluoromisonidazole (FMISO) With PET and MRI Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Elizabeth R. Gerstner, M.D. Funding: ACRIN
- 14. RTOG 0538- CALGB 30610/Endorsed Study: Phase III Comparison of Thoracic Radiotherapy Regimens in Patients with Limited Small Cell Lung Cancer Also Receiving Cisplatin and Etoposide Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Jeffrey Bogart, M.D., SUNY Syracuse Funding: RTOG
- RTOG 0631-Phase II/III Study of Image-Guided Radiosurgery/SBRT for Localized Spine Metastasis---RTOG CCOP Study Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Samuel Ryu, M.D., Henry Ford Hospital, Detroit Funding: RTOG
- RTOG 0834- Phase III Trial on Concurrent and Adjuvant Temozolomide Chemotherapy in Non-1p/19q Deleted Anaplastic Glioma: The CATNON Intergroup Trial

Institutional Co-investigator: Simon S. Lo, M.D. N. American PI: Michael A. Vogelbaum, MD, PhD Funding: RTOG

17. RTOG 0924- Androgen Deprivation Therapy and High Dose Radiotherapy With or Without Whole-Pelvic Radiotherapy in Unfavorable Intermediate or Favorable High Risk Prostate Cancer: A Phase III Randomized Trial Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Mack Roach III, M.D., UCSF Funding: RTOG

18. RTOG 0937- Randomized Phase II Study Comparing Prophylactic Cranial Irradiation Alone To Prophylactic Cranial Irradiation And Consolidative Extra-Cranial Irradiation For Extensive Disease Small Cell Lung Cancer (ED-SCLC) Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Elizabeth Gore, M.D., Medical College of Wisconsin Funding: RTOG

 RTOG 1010- A Phase III Trial Evaluating the Addition of Trastuzumab to Trimodality Treatment of Her2-Overexpressing Esophageal Adenocarcinoma Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Howard Safran, M.D. Funding: RTOG

20. RTOG 1114- Phase II Randomized Study of Rituximab, Methotrexate, Procarbazine, Vincristine, and Cytarabine With and Without Low-Dose Whole-Brain Radiotherapy for Primary Central Nervous System Lymphoma Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Antonio Omuro, M.D. Funding: RTOG

21. E3F05- Radiation Therapy With or Without Temozolomide in Treating Patients With Low-Grade Glioma Institutional Co-investigator: Simon S. Lo, M.D. Study PI: David Schiff, M.D., University of Virginia Funding: ECOG

- 22. E6508- BLP25 Liposome Vaccine and Bevacizumab After Chemotherapy and Radiation Therapy in Treating Patients With Newly Diagnosed Stage IIIA or Stage IIIB Non-Small Cell Lung Cancer That Cannot Be Removed by Surgery Institutional Co-investigator: Simon S. Lo, M.D. Study PI: Jyoti Patel, M.D., Northwestern Memorial Hospital Funding: ECOG
- Case6307- O6-Benzylguanine and Temozolomide in Combination With Genetically Modified Peripheral Blood Stem Cells in Newly Diagnosed Glioblastoma Multiforme Co-investigator: Simon S. Lo, M.D Study Sponsor: Stanton Gerson, M.D. Study PI: Andrew E. Sloan, M.D.
- 23. NBIO1307- A Phase III Clinical Trial Evaluating DCVax-L, Autologous Dendritic Cells Pulsed with Tumor Lysate Antigen for the Treatment of

Glioblastoma Multiforme Co-investigator: Simon S. Lo, M.D. Institutional PIs: Andrew E. Sloan, M.D. (Case Western Reserve University) and David Peereboom, M.D. (CCF)

24. CLDX 1311- An International, Randomized, Double-Blind, Controlled Study of Rindopepimut/GM-CSF with Adjuvant Temozolomide in Patients with Newly Diagnosed, Surgically Resected, EGFRvIII-positive Glioblastoma (The "ACT IV" Study) Co-investigator: Simon S. Lo, M.D.

Institutional PIs: Andrew E. Sloan, M.D. (Case Western Reserve University) and Gene Barnett, M.D. (CCF)

- 25. CLDX 2311- A Phase II Study of Rindopepimut/GM-CSF in Patients with Relapsed EGFRvIII-Positive Glioblastoma (The "ReACT" Study) Co-investigator: Simon S. Lo, M.D. Institutional PIs: Andrew E. Sloan, M.D. (Case Western Reserve University) and David Peereboom, M.D. (CCF)
- 26. CHRV 0115- Tumor Volume as a Predictor of Overall Survival for Patients with Brain Metastases Treated with Stereotactic Radiosurgery Co-investigator: Simon S. Lo, M.D. Study PI: Mitchell Machtay, M.D.
- 27. Case7Z11- Establishment of a Cancer Image Segmentation Database for Training Resident Physicians and Testing Proficiency Prior to Submitting Clinical Cases to Cooperative Groups Co-investigator: Simon S. Lo, M.D. Study PI: Jason Sohn, Ph.D.
- 28. Case 1307- Ohio Brain Tumor Study Co-investigator: Simon S. Lo, M.D. Study PI: Jill Barnholtz-Sloan (CWRU) and Gene Barnett (CCF)
- 29. BTTC 1312- Randomized, Double-Blind, Placebo-Controlled Trial of Lacosamide for Seizure Prophylaxis in Patients with High-Grade Gliomas Co-investigator: Simon S. Lo, M.D. Institutional PIs: PI: Lisa Rogers, D.O. (CWRU) and Manmeet Ahluwalia, M.D. (CCF)
- 30. ABBT 1512- A Randomized, Double-Blind, Phase 2, Dose-Ranging Study to Evaluate the Safety and Efficacy of Veliparib and Whole Brain Radiation Therapy Versus Placebo and Whole Brain Radiation Therapy in Subjects with Brain Metastases from Non-Small Cell Lung Cancer Co-investigator: Simon S. Lo, M.D. Institutional PIs: PI: Mitchell Machtay, M.D. (CWRU) and John Suh, M.D. (CCF)
- 31. A071101- Phase II Randomized Trial Comparing the Efficacy of Heat Shock Protein-Peptide Complex-96 (HSPPC-96) (NSC#725085, Alliance IND #15380) Vaccine Given with Bevacizumab Versus Bevacizumab Alone in the Treatment of Surgically Resectable Recurrent Glioblastoma Multiforme (GBM) Co-investigator: Simon S. Lo, M.D.

Institutional PIs: Andrew E. Sloan, M.D.

32. NABTT0703- Iniparib and Temozolomide With or Without Radiation Therapy in Treating Patients With Newly Diagnosed Malignant Glioma Co-investigator: Simon S. Lo, M.D. Institutional PIs: Andrew E. Sloan, M.D

#### University of Washington Medical Center/ Fred Hutchinson Cancer Research Center

- 33. NRG BN001- Randomized Phase II Trial of Hypofractionated Dose-Escalated Photon IMRT or Proton Beam Therapy Versus Conventional Photon Irradiation With Concomitant and Adjuvant Temozolomide in Patients With Newly Diagnosed Glioblastoma Co-investigator at UW/ Fred Hutch: Simon S Lo, MD
- 34. NRG BN005- A Phase II Randomized Trial of Proton Vs. Photon Therapy (IMRT) for Cognitive Preservation in Patients With IDH Mutant, Low to Intermediate Grade Gliomas Co-investigator at UW/ Fred/ Hutch: Simon S. Lo, MD
- 35. Alliance N0577 (CODEL)- Phase III Intergroup Study of Radiotherapy With Concomitant and Adjuvant Temozolomide Versus Radiotherapy With Adjuvant PCV Chemotherapy in Patients With 1p/19q Co-deleted Anaplastic Glioma or Low Grade Glioma Co-investigator at UW/ Fred Hutch: Simon S Lo, MD
- 36. RTOG 3508/AbbVie M13-813- A Randomized, Placebo Controlled Phase 3 Study of ABT-414 with ConcurrentChemoradiation and Adjuvant Temozolomide in Subjects with Newly Diagnosed Glioblastoma (GBM) with Epidermal Growth Factor Receptor (*EGFR*) Amplification (Intellance1) Co-investigator at UW/ Fred Hutch: Simon S. Lo, MD
- 37. NRG BN003- Phase III Trial of Observation Versus Irradiation for a Gross Totally Resected Grade II Meningioma Co-investigator at UW/ Fred Hutch: Simon S. Lo, MD
- 38. ALLIANCE A211401: REDUCING SURGICAL COMPLICATIONS IN NEWLY DIAGNOSED LUNG CANCER PATIENTS WHO SMOKE CIGARETTES (PI: Ivana Croghan, PhD) Study champion (NRG): Simon S. Lo, MD

### Others

39. TROG 15.03 FASTRACK II clinical trial-Stereotactic Ablative Radiotherapy (SABR) as a New Precision Treatment Option in Kidney Cancer Associate Investigator: Simon S Lo, MD PI: Shankar Siva, MB,BS, PhD, FRANZCR

# **Research Funding/ Grants**

1. NIH R01

1R01CA196687 Title: Accurate MR-based PET Attenuation Correction for Quantitative Clinical Trials \$1.9M direct. \$2.9M total Sep. 2015 - Aug. 2019 PI: Raymond Muzic, Ph.D. (raymond.muzic@case.edu) Co-investigator (5% effort without salary support) Case Western Reserve University Funds stayed with Case Western Reserve University when I left the institution in July 2016

SHSC 1312- STEREOTACTIC RADIOSURGERY (SRS) +/- WHOLE-BRAIN RADIOTHERAPY (WBRT) FOR 2 THE TREATMENT OF BRAIN METASTASES: PATIENT PREFERENCE STUDY Institutional PI: Simon S. Lo, M.D. Study PI : Edward Chow, MB, BS, FRCPC, Sunnybrook Health Science Centre, University of Toronto Funding: Internal Amount: \$3,000 Comments: Study has been completed and published.

- 3. Elekta International Oligometastasis Consortium (CORE group) Institutional PI at UH/ Case Western Reserve University Amount: Canadian \$4,000 (Funds stayed with Case Western Reserve University when I left in July 2016)
- 4. North American Gamma Knife Consortium Registry Institutional PI at UH/ Case Western Reserve University
- 5. Elekta ICON Gamma Knife Expert Group Institutional-investigator at University of Washington/ HMC Amount: TBD
- 6. UG1 CA 233328 (Yu) 03/01/19-02/28/25 0.12 CM NIH/NCI \$19,830,301

Hutchinson Center as Lead Academic Participating Site (UG1)

The goal of the NCI National Clinical Trials Network (NCTN) is to develop and conduct state-of-the-art cancer treatment and advanced imaging clinical trials, especially large, definitive multi-institutional trials evaluating new cancer therapies and related clinical approaches. NCTN Lead Academic Participating Sites (LAPS) are one component of this network and will provide scientific leadership in the development and conduct of NCTN clinical trials in association with one or more adult Network Groups as well as substantial accrual to clinical trials conducted across the entire NTCN.

Role: LAPS Leadership

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## Manuscripts in refereed journals: Total 242

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6. Chang E, <u>Lo S</u>. Diagnosis and management of Central Nervous System Metastases from Breast Cancer. Oncologist. 2003;8(5):398-410. (ISI 2012 Impact factor: 6.7)

7. <u>Lo SS</u>, Hall WA, Cho KH, Orner J, Lee CK, Dusenbery KE. Radiation dose response for supratentorial low grade gliomas- Institutional experience and literature review. J Neurol Sci. 2003 Oct 15;214(1-2):43-8\*. (ISI 2012 Impact factor: 2.24)

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 First author: Hematology/ oncology mentee at Loyola University of Chicago

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### Manuscripts submitted or in preparation (Total: 12)

1.Shankar Siva, ..., <u>Simon Lo</u>. International Stereotactic Radiosurgery Society Guidelines on Stereotactic Body Radiotherapy for Primary Renal Cell Carcinoma.

2. ACR Appropriateness Criteria Expert Panels on Neurological Imaging, Interventional Radiology, and Musculoskeletal Imaging: Majid A. Khan, MBBS, BS<sup>2</sup>; Jack W. Jennings, MD, PhD, MPH<sup>b</sup>; Jonathan C. Baker, MD<sup>c</sup>; Amanda R. Smolock, MD<sup>d</sup>; Lubdha M. Shah, MD<sup>e</sup>; Jason W. Pinchot, MD<sup>f</sup>; Daniel E. Wessell, MD, PhD<sup>g</sup>; Charles Y. Kim, MD<sup>h</sup>; Leon Lenchik, MD<sup>i</sup>; Matthew S. Parsons, MD<sup>j</sup>; Gina Huhnke, MD<sup>k</sup>; <u>Simon Shek-Man Lo, MB, ChB<sup>l</sup></u>; Yi Lu, MD, PhD<sup>m</sup>; Christopher Potter , MD<sup>n</sup>; Charles Reitman, MD<sup>o</sup>; Scott S. Russo, MD<sup>p</sup>; Arjun Sahgal, MD<sup>q</sup>; Akash Sharma, MD, MBA<sup>r</sup>; Naga M. Yalla, MD<sup>s</sup>; Francesca D. Beaman, MD<sup>t</sup>; Baljendra S. Kapoor, MD<sup>u</sup>; Judah Burns, MD.<sup>v</sup> Management of Vertebral Compression Fractures. Will be submitted to JACR.

3. AANS/ CNS guideline updates on the role of RT in low grade gliomas, TBD.

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6. International survey of SBRT for head and neck cancer. In progress.

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- 22. Proton therapy and MR LINAC for HCC\*
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## **Guest Editorships:**

Focused issue "Advances in Stereotactic Radio Surgery". Chinese Clinical Oncology. Vol 6, Supplement 2 (September 2017)- Guest Editors: Kevin Chua, David Tan, Melvin Chua, Simon Lo (Indexed in PUBMED)

- 1. Kevin L.M. Chua, David B.H. Tan, Melvin. L.K. Chua, Simon S. Lo. The promise of stereotactic body radiotherapy—next phase of integration into oncological practice.
- 2. Hannah Tharmalingam, Peter J. Hoskin. The optimism surrounding stereotactic body radiation therapy and immunomodulation.
- 3. Connie Yip, Gary J. R. Cook, Kasia Owczarczyk, Vicky Goh. Challenges in imaging assessment following liver stereotactic body radiotherapy: pitfalls to avoid in clinical practice.
- 4. Lijun Ma, Lei Wang, Chia-Lin Tseng, Arjun Sahgal. Emerging technologies in stereotactic body radiotherapy
- 5. Stewart Gaede, Michael I. Lock. Advances in external beam stereotactic body radiotherapy: principle concerns in implementing a liver radiation program.
- 6. Philip Gilbo, Isabella Zhang, Jonathan Knisely. Stereotactic radiosurgery of the brain: a review of common indications.
- 7. Morten Høyer. Re-irradiation with stereotactic body radiation therapy (SBRT).
- 8. Thomas A. C. Kennedy, Mark T. Corkum, Alexander V. Louie. Stereotactic radiotherapy in oligometastatic cancer.
- 9. Gargi Kothari, Alexander V. Louie, David Pryor, Ian Vela, Simon S. Lo, Bin S. Teh, Shankar Siva. Stereotactic body radiotherapy for primary renal cell carcinoma and adrenal metastases.
- 10. Majed Alghamdi, Chia-Lin Tseng, Sten Myrehaug, Pejman Maralani, Chris Heyn, Hany Soliman, Young Lee, Mark Ruschin, Leodante da Costa, Victor Yang, Mikki Campbell, Arjun Sahgal. Postoperative stereotactic body radiotherapy for spinal metastases.
- 11. Janice S. H. Tan, Xiaotian Lin, Kevin L. M. Chua, Paula Y. Lam, Khee-Chee Soo, Melvin L. K. Chua. Exploiting molecular genomics in precision radiation oncology: a marriage of biological and physical precision.
- 12. Kevin L. M. Chua, Iris Sin, Kam W. Fong, Melvin L. K. Chua, Hiroshi Onishi. Stereotactic body radiotherapy for early stage lung cancer—historical developments and future strategies.

Focused issue "Radiation Therapy in Palliative Oncology Care". Annals of Palliative Medicine (Ed: Charles Simone). Guest Editors: Tracy Balboni, Yolanda Tseng, <u>Simon Lo</u>. Indexed in PUBMED. A total of 12 papers.

Focused issue "Oligometastases- Fallacy or Real Deal?". Annals of Palliative Medicine (Ed: Charles Simone). Guest Editors: <u>Simon Lo</u>, Michael Milano, Tithi Biswas and Charles Simone. Indexed in PUBMED. In progress. A total of 12 papers.

Balamurugan Vellayappan, <u>Simon S Lo</u>, Jonathan Knisely, Kevin Shiue. The modern approaches to the management of brain metastases. Chinese Clinical Oncology. In press.

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## Published books:

1.(Eds: <u>Simon S. Lo</u>, Bin S. Teh, Jiade J. Lu, Tracey E. Schefter). Medical Radiology: Stereotactic Body Radiation Therapy. Springer 2013. (<u>http://link.springer.com/book/10.1007%2F978-3-642-25605-9</u>)

(This is the only up-to-date comprehensive textbook in Stereotactic Body Radiation Therapy since 2004 and is available in medical libraries worldwide; this book has had  $\sim$ 49,400 chapter downloads from September 2012 through May 2019 apart from sales of hundreds of hardcopies)

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4.(Eds: Arjun Sahgal, Jason Sheehan, Lijun Ma, <u>Simon Lo</u>) Image-Guided Hypofractionated Stereotactic Radiotherapy (IG-HSRT). A Practical Approach to SRT For Brain And Spine Tumors. Taylor & Francis Books, Inc (UK) 2016

5.(Eds: Eric Chang, MD; Paul Brown, MD; Simon Lo, MD; Arjun Sahgal, MD, and John Suh, MD) Adult CNS Radiation Oncology. Springer Nature (2018) (This is the only comprehensive textbook in adult CNS radiation oncology and since it was released in August 2018, there

have been 46,718 chapter downloads in 12 months)

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#### Other publications e.g. in non-referred journals and letters to the editor:

#### Letters and Commentaries (Peer-reviewed):

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# National Practice Guidelines:

## American Society for Radiation Oncology (ASTRO)

1.Stephen Lutz, MD, Lawrence Berk, MD, PhD, Eric Chang, MD, Edward Chow, MBBS, Carol Hahn, MD, Peter Hoskin, MD, David Howell, MD, Andre Konski, MD, Lisa Kachnic, MD, <u>Simon Lo, MB, ChB</u>, Arjun Sahgal, MD, Larry Silverman, MD, Charles von Gunten, MD, PhD, FACP, Ehud Mendel, MD, FACS, Andrew Vassil, MD, Deborah Watkins Bruner, RN, PhD, and William Hartsell, MD. Palliative radiotherapy for bone metastases: An ASTRO evidence-based guideline (2011)

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# American Academy of Neurological Surgeons/ Congress of Neurological Surgeons

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## American College of Radiology (ACR)

1. Expert Panel On Radiation Oncology-Bone Metastases, Lutz ST, Lo SS, Chang EL, Galanopoulos N, Howell DD, Kim EY, Konski AA, Pandit-Taskar ND, Ryu S, Silverman LN, Van Poznak C, Weber KL. ACR Appropriateness Criteria: Non-Spine Bone Metastases (2012)

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6. Expert Panel on Radiation Oncology–Brain Metastases: Gregory M. M. Videtic, MD, Elizabeth M. Gore, MD, Jeffrey D. Bradley, MD, John M. Buatti, MD, Isabelle Germano, MD, A. Paiman Ghafoori, MD, Mark A. Henderson, MD, <u>Simon Shek-Man Lo, MB, ChB</u>, Stephen T. Lutz, MD, Gregory J. A. Murad, MD, Roy A. Patchell, MD, Samir H. Patel, MD, Jared R. Robbins, MD, H. Ian Robins, MD, PhD; Andrew D. Vassil, MD, Franz J. Wippold II, MD, Michael J. Yunes, MD. ACR Appropriateness Criteria<sup>®</sup> Multiple Brain Metastases (2014)

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8.Expert Panel on Musculoskeletal Imaging: Catherine C. Roberts, MD; Mark J. Kransdorf, MD; Francesca D. Beaman, MD; Ronald S. Adler, MD, PhD; Behrang Amini, MD, PhD; Marc Appel, MD; Stephanie A. Bernard, MD; Ian Blair Fries,

MD; Isabelle M. Germano, MD; Bennett S. Greenspan, MD, MS; Langston T. Holly, MD; Charlotte D. Kubicky, MD, PhD; <u>Simon Shek-Man Lo, MB, ChB</u>; Timothy J. Mosher, MD; Andrew E. Sloan, MD; Michael J. Tuite, MD; Eric A. Walker, MD; Robert J. Ward, MD; Daniel E. Wessell, MD; Barbara N. Weissman, MD. American College of Radiology. ACR Appropriateness Criteria<sup>®</sup> The Follow-up of Malignant or Aggressive Musculoskeletal Tumors (2015)

9.Expert Panel on Neurologic Imaging: Lubdha M. Shah, MD; Jack W. Jennings, MD, PhD; Claudia F. E. Kirsch, MD; Eric J. Hohenwalter, MD; Francesca D. Beaman, MD; R. Carter Cassidy, MD; Michele M. Johnson, MD; A. Tuba Kendi, MD; <u>Simon Shek-Man Lo, MB, ChB</u>; Charles Reitman, MD; Arjun Sahgal, MD; Matthew J. Scheidt, MD; Kristofer Schramm, MD; Daniel E. Wessell, MD, PhD; Mark J. Kransdorf, MD; Jonathan M. Lorenz, MD; Julie Bykowski, MD. American College of Radiology. ACR Appropriateness Criteria<sup>®</sup>Management of vertebral compression fractures.

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#### Abstracts:

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103. J.C. Grecula, S. Elias, K. Thelen, M. Knopp, G. Otterson, P. Ross, E. Kassis, M. Welliver, M. Villalona-Calero, K. Shila, <u>S. Lo</u>, G. Jia, W.C. Yu, B. Yuh, S. Ghosh, E. Bertino, N. Mayr. Dynamic-Contrast Enhanced MR and Volume Regression Rate as a Preoperative Predictive Assay in Patients With Non-Small Cell Lung Cancer. International Journal of Radiation Oncology • Biology • Physics, Vol. 90, Issue 1, S816. Published in issue: September 01 2014

104. J. Yuan, D. Albani, Y. Zheng, B.W. Wessels, <u>S.S. Lo</u>, M. Yao. Experimental Validation of Monte Carlo Simulations Based on a Virtual Source Model for Tomotherapy in a Rando Phantom. International Journal of Radiation Oncology • Biology • Physics, Vol. 93, Issue 3, E608. Published in issue: November 01 2015

105. Z. Huang, Y. Feng, <u>S.S. Lo</u>, N.A. Mayr, W. Yuh, T.G. Yu, X.Y. Feng, J.Z. Dai, H.J. Qian. Correlation Between Metabolic Information and Tumor Volume for Astrocytic Brain Tumors Before and During Radiation Therapy. International Journal of Radiation Oncology • Biology • Physics, Vol. 93, Issue 3, E67. Published in issue: November 01 2015

106. K.J. Redmond, S.P. Robertson, <u>S.S. Lo</u>, S.G. Soltys, S. Ryu, T.R. McNutt, S.T. Chao, I.J. Barani, Y. Yamada, A.J. Ghia, E.L. chang, J.P. Sheehan, A. Sahgal. International Consensus Contouring Guidelines for Postoperative Spine Stereotactic Body Radiation Therapy (SBRT). International Journal of Radiation Oncology • Biology • Physics, Vol. 93, Issue 3, S56.Published in issue: November 01 2015

107. S. Siva, R.J. Ellis III, L. Ponsky, B.S. Teh, A. Mahadevan, A. Muacevic, H. Onishi, P. Wersall, T. Nomiya, <u>S.S. Lo</u>. A Multinational Report on Technical Factors of Stereotactic Body Radiation Therapy for Primary Renal Cell Carcinoma. International Journal of Radiation Oncology • Biology • Physics, Vol. 93, Issue 3, E577–E578. Published in issue: November 01 2015

108. <u>S.S. Lo</u>, R.J. Ellis III, L. Ponsky, B.S. Teh, A. Mahadevan, A. Muacevic, H. Onishi, P. Wersall, T. Nomiya. A Multinational Report on Factors of Stereotactic Body Radiation Therapy for Primary Renal Cell Carcinoma: Patient Selection and Follow-up. International Journal of Radiation Oncology • Biology • Physics, Vol. 93, Issue 3, E208–E209. Published in issue: November 01 2015

# National invitational lectures and others

#### Invited and/or refereed international or national meetings (on behalf of co-authors and co-presenters, if applicable):

Single dose vs. fractionated stereotactic radiotherapy for meningioma. Poster discussion session on November 2, 1999 in American Society for Therapeutic Radiology and Oncology meeting in San Antonio, TX

Does the extent of surgery have an impact on survival for patients with supratentorial low grade gliomas who received postoperative radiation therapy? Poster presentation in American Society for Therapeutic Radiology and Oncology meeting in October 2000 in Boston, MA

Is stereotactic radiotherapy adequate treatment for atypical and malignant meningiomas? Poster discussion in American Society of Clinical Oncology meeting in June 2002 in Orlando, FL

Radiosurgery for Meningiomas and Acoustic Neuromas Guest Faculty at the VI Detroit Neurosurgery Symposium (Wayne State University): The Renaissance of Neurosurgery, September 13-14, 2002

Radiation therapy with or without chemotherapy for cervical cancer with periaortic lymph node metastasis. Oral presentation in American Society for Therapeutic Radiology and Oncology Annual Meeting in October 2002, New Orleans, LA.

Prognostic factors in patients who receive radiation therapy for supratentorial low grade glioma. Oral presentation in the Annual Meeting of the Congress of Neurological Surgeons in September 2002, Philadelphia, PA.

Radiation dose response for supratentorial low grade gliomas revisited. Oral presentation in Radiological Society of North America Annual Meeting in November 2002, Chicago, IL.

Post-Operative Radiotherapy for Endometrial Cancer with Pathologic Risk Factors: Patterns of Failure. Poster presentation in the American Society of Clinical Oncology Annual Meeting in June 2003, Chicago, IL.

Low Dose Gamma Knife-based Stereotactic Radiosurgery for Vestibular Schwannoma. Oral presentation in Radiological Society of North America in November 2007, Chicago, IL.

Stereotactic Radiosurgery with or without Whole Brain Radiotherapy for Patients with 2-4 Radioresistant Brain Metastases. Oral presentation in Radiological Society of North America in November 2008, Chicago, IL.

Stereotactic Radiosurgery Alone for Patients with 1-4 Radioresistant Brain Metastases. Poster presentation at American Society for Radiation Oncology in November 2009, Chicago, IL.

International Practice Patterns and Consensus on the Management of Brain Metastases. Nursing Program Faculty. American Society for Radiation Oncology on November 3, 2009, Chicago, IL. Invited.

Oncodiagnosis Panel: Detection and Treatment of Early Lung Cancer--Current Controversies and Future Directions: Oncodiagnosis (Plenary Session). Radiological Society of North America Annual Meeting, November 29, 2009, Chicago, IL. Invited.

Moderator/ Presiding Officer. Oral presentation in lung and sarcoma. Radiological Society of North America Annual Meeting, December 3, 2009, Chicago, IL. Invited.

History and Overview of Stereotactic Body Radiation Therapy Stereotactic Body Radiation Therapy Symposium National University Cancer Institute of Singapore Sponsored by Singapore Radiological Society June 21-23, 2010 Singapore Invited.

Stereotactic Body Radiation Therapy for Early Stage Non-Small Cell Lung Cancer Stereotactic Body Radiation Therapy Symposium National University Cancer Institute of Singapore Sponsored by Singapore Radiological Society June 21-23, 2010 Singapore Invited.

Stereotactic Body Radiation Therapy for Spinal Metastases Stereotactic Body Radiation Therapy Symposium National University Cancer Institute of Singapore Sponsored by Singapore Radiological Society June 21-23, 2010 Singapore Invited. Stereotactic Body Radiation Therapy for Non-Pulmonary Primary Tumors Stereotactic Body Radiation Therapy Symposium National University Cancer Institute of Singapore Sponsored by Singapore Radiological Society June 21-23, 2010 Singapore Invited.

Stereotactic Body Radiation Therapy for Oligometastases Stereotactic Body Radiation Therapy Symposium National University Cancer Institute of Singapore Sponsored by Singapore Radiological Society June 21-23, 2010 Singapore Invited.

Starting Stereotactic Body Radiation Therapy in a Comprehensive Cancer Center Stereotactic Body Radiation Therapy Symposium National University Cancer Institute of Singapore Sponsored by Singapore Radiological Society June 21-23, 2010 Singapore Invited.

Radiation and Cancer Biology of Stereotactic Body Radiation Therapy Stereotactic Body Radiation Therapy Symposium National University Cancer Institute of Singapore Sponsored by Singapore Radiological Society June 21-23, 2010 Singapore Invited.

Normal Tissue Constraints of Stereotactic Body Radiation Therapy Stereotactic Body Radiation Therapy Symposium National University Cancer Institute of Singapore Sponsored by Singapore Radiological Society June 21-23, 2010 Singapore Invited.

Stereotactic Body Radiation Therapy for Stage I Non-Small Cell Lung Carcinoma BIT's 3<sup>rd</sup> World Cancer Congress (WCC)-2010 Singapore June 23, 2010

Stereotactic body radiation therapy/ Image-guided radiotherapy. Refresher Course. Radiological Society of North America Annual Meeting, December 2, 2010, Chicago, IL. Invited.

Moderator/ Presiding Officer. Radiation Oncology and Radiobiology (Pediatrics and Central Nervous System). Radiological Society of North America Annual Meeting, November 30, 2010, Chicago, IL. Invited. Stereotactic Body Radiation Therapy for early stage non-small cell lung cancer. American Thoracic Society Annual Meeting, May 2011, Denver, CO. Invited.

Skull Base Tumors University of Miami Comprehensive Cancer Center Hypofractionation 2011- Scientific Concepts and Clinical Experiences Miami Beach, FL, September 30, 2011 Invited.

Stereotactic body radiation therapy/ Image-guided radiotherapy. Refresher Course. Radiological Society of North America Annual Meeting (RSNA), December 2, 2011, Chicago, IL. Invited.

Moderator/ Presiding Officer. Radiation Oncology and Radiobiology (Lung). November 30, 2011. RSNA 2011, Chicago, IL. Invited.

Central Nervous System BOOST. Refresher Course. December 1, 2011. RSNA 2011, Chicago, IL. Invited.

SBRT for spinal metastasis International SBRT Symposium (Supported by RTOG) National University Cancer Institute of Singapore March 22-24, 2012 Invited.

Complications from SBRT International SBRT Symposium (Supported by RTOG) National University Cancer Institute of Singapore March 22-24, 2012 Invited.

Local therapy for oligometastases International SBRT Symposium (Supported by RTOG) National University Cancer Institute of Singapore March 22-24, 2012 Invited.

Program Chair. Informational Session "Incorporation of advanced diagnostic imaging into the practice of radiation oncology", CARROS (ACR) program, Boston, MA. October 27, 2012. Invited.

Moderator and Panelist, Scientific Panel: Innovative Approaches To Management Of Spinal Metastasis. October 28, 2012. American Society for Radiation Oncology annual meeting. Invited.

Panelist, Scientific Panel: The Role of Innovation in Palliation – Is the Benefit Worth the Cost? October 29, 2012. Boston, MA. American Society for Radiation Oncology annual meeting 2012. Invited. (Presentations cancelled due to hurricane Sandy)

Moderator and Speaker, Controversy Session: Stereotactic Radiation for Oligo-Metastasis: New Paradigm or Wishful Thinking? Special Courses. November 26, 2012, Radiological Society of North America Annual Meeting (RSNA) 2012, Chicago, IL. Invited.

BOOST: Lung-Integrated Science and Practice (ISP) Session. November 26, 2012, RSNA 2012, Chicago, IL. Invited.

Moderator/ Presiding Officer.Radiation Oncology and Radiobiology (Lung). November 27, 2012, RSNA 2012, Chicago, IL. Invited.

Radiation Oncology and Radiobiology Afternoon CME Posters. November 27, 2012, RSNA 2012, Chicago, IL. Invited.

Stereotactic Ablative Radiotherapy for Oligometastasis. International Society of Radiosurgery Congress. Breakfast Seminar. June 19, 2013, Toronto, ON, Canada, Invited.

Session Co-Chair, Plenary Session. Spines. International Society of Radiosurgery Congress. June 19, 2013, Toronto, ON, Canada, Invited.

Program Chair. Informational Session "ACR Appropriateness Criteria- A Useful Practice Tool", CARROS (ACR) program, Atlanta, GA, September 21, 2013.

Panelist, Scientific Panel: Radiosurgery for Metastases – Improved Patient Care or Unnecessary Cost? September 25, 2013, American Society for Radiation Oncology (ASTRO) annual meeting 2013, Atlanta, GA.

Clinical Aspects of LINAC-based SBRT. October 12, 2013, AAPM Ohio Valley and Ohio-Penn Chapters Fall Meeting 2013, Beachwood, OH. Invited.

Moderator/ Presiding Officer.Radiation Oncology and Radiobiology (CNS). December 2, 2013, Radiological Society of North America Annual Meeting (RSNA) 2013, Chicago, IL. Invited.

BOOST: Head and Neck—Integrated Science and Practice (ISP) Session. December 2, 2013, RSNA 2013, Chicago, IL. Invited.

Session Chair and Panelist. Role of Stereotactic Ablative Radiotherapy (SABR) and Interventional Radiology in the Management of Oligometastases. Refresher Course. December 3, 2013. RSNA 2013, Chicago, IL. Invited.

Leksell Gamma Knife Meeting Moderator Breakfast Seminar Spinal Radiosurgery May 15, 2014 New York City, NY (Declined this invitation due to time conflict)

Showdown session: Tumor Radiobiology: Does Conventional Wisdom Hold for SRS/SBRT? Radiosurgery Society Biannual Meeting May 9, 2014 Minneapolis, MN

Invited Speaker (Together with Drs. Ben Slotman, Arjun Sahgal, Bin Teh) Strategies to mitigate complications from SBRT Educational session. American Society for Radiation Oncology (ASTRO) Annual Meeting, September 17, 2014, San Francisco, CA

Co-moderator Oral scientific presentation: Brain and spinal metastases American Society for Radiation Oncology (ASTRO) Annual Meeting, September 17, 2014, San Francisco, CA A Multi-National Report on Technical Factors of Stereotactic Body Radiotherapy for Oligometastases. Poster presentation at American Society for Radiation Oncology Annual Meeting, September 15, 2014, San Francisco, CA

Program Chair. Informational Session "Cost-effectiveness of cancer therapy with special focus on radiotherapeutic procedures for breast and prostate cancer", pre-ASTRO CARROS (ACR) program, September 13, 2014, San Francisco, CA

Course Director and Moderator BOOST: Lung Cancer Case-based Review December 1, 2014 RSNA 2014, Chicago, IL

Moderator BOOST: Lung Cancer Integrated Science and Practice December 1, 2014 RSNA 2014, Chicago, IL

Course Director and Moderator BOOST: Lung Cancer Anatomy and Contouring December 1, 2014 RSNA 2014, Chicago, IL

Session Chair and Panelist. Role of Stereotactic Ablative Radiotherapy (SABR) and Interventional Radiology in the Management of Oligometastases. Refresher Course. December 2, 2014. RSNA 2014, Chicago, IL. Invited.

Invited presenter

"A Phase 2 Randomized, Double Blind, Placebo-Controlled Clinical Trial of D-methionine For the Prevention of Cisplatin-Induced Ototoxicity and Other Side Effects in Patients with Gynecological (Ovarian, Endometrial, Cervical) and Head and Neck Cancer" NRG Oncology Meeting Cancer Prevention and Control Committee Meeting February 6, 2015 San Diego, CA

Invited presenter

"A Phase 2 Randomized, Double Blind, Placebo-Controlled Clinical Trial of D-methionine For the Prevention of Cisplatin-Induced Ototoxicity and Other Side Effects in Patients with Gynecological (Ovarian, Endometrial, Cervical) and Head and Neck Cancer" NRG Oncology Meeting Cervical Cancer Work Group Meeting February 6, 2015 San Diego, CA

Invited Speaker IAEA Stereotactic Body Radiotherapy Symposium Sydney, Australia July 2015 Declined due to time conflict Invited Speaker Spinal SRS American Radium Society Hawaii April 2015 Declined due to time conflict

Invited Speaker SBRT for gastrointeestinal malignancies Yashoda International Cancer Conference: Recent Advances and Newer Horizons Hyderabad, India January 24- 25, 2015 Declined due to time conflict

Invited Speaker Radiobiology of SRS/ SBRT and Strategies to mitigate risk of complication of SBRT Pre-European Society for Radiotherapy and Oncology (ESTRO) Stereotactic Body Radiotherapy Symposium Barcelona, Spain April 24, 2015

Invited Speaker SBRT for bone metastasis Pre-European Society for Radiotherapy and Oncology (ESTRO) Stereotactic Body Radiotherapy Symposium Barcelona, Spain April 24, 2015

Invited Speaker The State of Brain and CNS Research Continuing Education Session Radiotherapy Research: Building on the Past and Looking to the Future Amercian College of Radiology Washington, DC May 17, 2015

Distinguished Speaker Management Of Spine Mets In The Era Of Spine SRS Session Chair, Oligometastasis in 2015 - Fact or Myth? Singapore Radiological Society Annual General Meeting Singapore May 14-15, 2015

Distinguished Speaker SRS Should Be First Line For Oligo Brain Metastasis Session Chair, Oligometastasis in 2015 - Fact or Myth? Singapore Radiological Society Annual General Meeting Singapore May 14-15, 2015

Distinguished Speaker Integration Of Systemic Agents With SRS In Oligometastatic Disease Session Chair, Oligometastasis in 2015 - Fact or Myth? Singapore Radiological Society Annual General Meeting Singapore May 14-15, 2015

Invited Speaker CT and MRI Guided Radiosurgery Procedures North American Gamma Knife Consortium Meeting Cleveland, OH June 27, 2015

Invited presenter "A Phase 2 Randomized, Double Blind, Placebo-Controlled Clinical Trial of D-methionine For the Prevention of Cisplatin-Induced Ototoxicity and Other Side Effects in Patients with Gynecological (Ovarian, Endometrial, Cervical) and Head and Neck Cancer" NRG Oncology Meeting Head and Neck Core Committee Meeting July 17, 2015 Denver, CO

Invited Speaker (Together with Drs. Ben Slotman, Arjun Sahgal, Bin Teh) Strategies to mitigate complications from SBRT Educational session. American Society for Radiation Oncology (ASTRO) Annual Meeting 2015 San Antonio, TX October 21, 2015

Invited Speaker (On behalf of Dr. Arjun Sahgal) Strategies to mitigate complications from SBRT: Spine Educational session. American Society for Radiation Oncology (ASTRO) Annual Meeting 2015 San Antonio, TX October 21, 2015

Co-moderator Oral scientific presentation: Brain SRS American Society for Radiation Oncology (ASTRO) Annual Meeting 2015 San Antonio, TX October 19, 2015

Session Chair and Panelist Image-guided stereotactic body radiotherapy (SBRT) for spinal metastases- Spinal imaging, target delineation and post-SBRT response evaluation Refresher Course RSNA 2015 Chicago, IL December 1, 2015

Moderator Fundamentals of Imaging for the Radiation Oncologist Refresher course RSNA 2015 Chicago, IL November 29, 2015

Keynote Speaker BOOST: CNS Tumors Integrated Science and Practice RSNA 2015 Chicago, IL December 1, 2015

Invited speaker Toxicities associated with SBRT and strategies to avoid them Langendorff Congress 2015 University of Freiberg, Germany October 10, 2015

Invited speaker Development of SBRT in the past decade- Global perspective University of North Carolina/ Accuray User Meeting November 13, 2015

Invited speaker Prospective clinical trials of SBRT for gynecologic malignancies- University Hospitals Seidman Cancer Center/ Case Comprehensive Cancer Center experience University of North Carolina/ Accuray User Meeting November 13, 2015

Invited speaker Response assessment after stereotactic body radiotherapy for spinal metastasis: a report from the SPIne response assessment in Neuro-Oncology (SPINO) group NRG Oncology Semi-Annual Meeting Functional Imaging Work Group Atlanta, Georgia January 22, 2016

Invited Speaker Safe and effective delivery of spine SBRT: Webinar (Invited) Radiosurgery Society (RSS) February 4, 2016

Invited Speaker Safe and effective delivery of spine SBRT: Webinar (Invited) American Association of Medical Dosimetrists (AAMD) April 13, 2016

Invited speaker Precision Radiotherapy Technologies in Management of Brain Metastases American Society of Clinical Oncology Palliative Care in Oncology Symposium San Francisco, CA September 10, 2016 Presenter Consensus Statement from the International Stereotactic Body Radiotherapy Consortium for Head and Neck Carcinoma -Patient Selection and Pre- and Post-SBRT Evaluation Poster Presentation American Society for Radiation Oncology (ASTRO) Annual Meeting 2016 Boston, MA September 26, 2016

Invited speaker (Together with Drs. Ben Slotman, Arjun Sahgal, Bin Teh) Strategies to mitigate complications from SBRT Educational session (SA-CME) American Society for Radiation Oncology (ASTRO) Annual Meeting 2016 Boston, MA September 27, 2016

Invited speaker Spinal SBRT Fundementals and Special Considerations: Current Research and Future Directions Educational session. American Society for Radiation Oncology (ASTRO) Annual Meeting 2016 Boston, MA September 28, 2016

Discussant CNS tumors: e-poster Scientific session American Society for Radiation Oncology (ASTRO) Annual Meeting 2016 Boston, MA September 25, 2016

Invited speaker Why SBRT and What is State of the Art Hands-On SBRT Workshop MTMI Seattle, WA October 14, 2016

Invited speaker Future Directions of SBRT Hands-On SBRT Workshop MTMI Seattle, WA October 16, 2016

Invited expert Meeting the expert session Hands-On SBRT Workshop MTMI, Seattle, WA October 16, 2016

Invited speaker Golden age of radiotherapy Hong Kong International Oncology Symposium Hong Kong SAR October 28, 2016

Invited speaker Brain and spine radiosurgery Hong Kong Neurosurgical Society Hong Kong SAR October 31, 2016

Course Director and Moderator BOOST: Lung Cancer Case-based Review RSNA 2016, Chicago, IL

Moderator and Speaker Controversy Session: Is It Time to Put Whole Brain Radiotherapy to Pasture? What's New in the Treatment of Limited Brain Metastases RSNA 2016, Chicago, IL

Moderator Imaging Evaluation, Target Delineation and Response Evaluation for Skull Base and Spinal Stereotactic Radiosurgery/Radiotherapy Refresher course (SA-CME) RSNA 2016, Chicago, IL

Moderator Stereotactic body radiotherapy for renal cell carcinoma (Webinar) Radiosurgery Society December 13, 2016

Speaker Optimal imaging for disease evaluation, target delineation and post-treatment response evaluation for stereotactic body radiotherapy for spinal metastases (Webinar) Radiosurgery Society January 17, 2017

Moderator Long-Term Outcome of SBRT in Non-Small Cell Lung Cancer: Indiana University Experience (Webinar) Radiosurgery Society October 10, 2017

Faculty eContouring for Pediatric and SBRT/Spine Cancer American Society for Radiation Oncology (ASTRO) Annual Meeting 2017 San Diego, CA September 24, 2017

Faculty Poster Walk with a Professor Radiosurgery Society Meeting Las Vegas, NV November 3, 2017

Moderator (with Markus Kufeld) SRS for Spine – Oral Abstract Session Radiosurgery Society Meeting Las Vegas, NV November 2, 2017

Moderator Lung BOOST- Case Based Review November 28, 2017 RSNA 2017, Chicago, IL

Moderator Lung BOOST- The Confluence of Diagnostic Radiology and Radiation Oncology November 28, 2017 RSNA 2017, Chicago, IL

Speaker Oncodiagnosis Panel- Spinal Metastasis November 26, 2017 RSNA 2017, Chicago, IL

Invited Speaker Radiosurgical Contouring and Dosimetry, and Complication Avoidance: Conventional Radiation vs. Radiosurgery 8th Annual Multimodal Treatment of Spinal Tumors- Seattle Science Foundation Seattle, WA April 14, 2018

Invited Speaker Spine Metastasis: Conventional Radiation Therapy vs. Radiosurgery 8th Annual Multimodal Treatment of Spinal Tumors- Seattle Science Foundation Seattle, WA April 14, 2018

Invited Panelist (Jens R. Chapman, M.D. and Ehud Mendel, M.D. vs. Simon S. Lo, M.D. and Peter Gerszten, M.D., M.P.H., FACS) Battle of the Surgeons: Case Discussions and Debates: Radiosurgery vs. Surgery 8th Annual Multimodal Treatment of Spinal Tumors- Seattle Science Foundation Seattle, WA April 14, 2018

Invited panelist/ speaker Expert Panel: Challenging Cases in Palliation Palliative Radiation in the Modern Era: Evidence and Strategies for Best Practice New York, NY April 20, 2018

Invited panelist/ speaker Expert Panel: Immunotherapy + Radiation: Issues of Safety, Synergy, and Pseudoprogression Palliative Radiation in the Modern Era: Evidence and Strategies for Best Practice New York, NY April 20, 2018

Moderator (Speakers: Stephanie Combs and Hany Soliman) Postoperative SRS/SRT to the Surgical Cavity after Brain Metastases Resection Radiosurgery Society Webinar June 5, 2018

Invited speaker SBRT for renal cell carcinoma Hong Kong SBRT study group (HKSBRTSG) Hong Kong SAR June 30, 2018

Invited speaker Safe Delivery of SBRT: Normal Tissue Tolerance and Interaction with Systemic Therapies Hong Kong SBRT study group (HKSBRTSG) Hong Kong SAR June 30, 2018

Discussant PRO Scientific Session ASTRO San Antonio, TX October 20, 2018

Invited faculty/speaker PRO- CNS Updates In The Management of Brain Metastases ASTRO San Antonio, TX October 21, 2018

Invited faculty/speaker CNS/ Spine SBRT E-contouring ASTRO San Antonio, TX October 22, 2018

Invited faculty/speaker Scientific Panel- Use of Advanced Technologies in Palliative Care: a Brave New World or a Costly Mistake? Appropriate use of advanced technologies ASTRO San Antonio, TX October 24, 2018

International Distinguished Speaker The changing landscape of brain metastasis management ICON Group Melbourne, Australia November 7, 2018 International Distinguished Speaker Plenary Session: Contemporary advances in the management of brain metastases Stereotactic Interest Group of Australasia (SIGA) Inaugural Meeting Melbourne, Australia November 8, 2018

International Distinguished Speaker Stereotactic Radiosurgery for large brain metastases – Should we do it alone, before surgery or after surgery? Stereotactic Interest Group of Australasia (SIGA) Inaugural Meeting Melbourne, Australia November 8, 2018

International Distinguished Speaker The role of spine SBRT in the palliative oncology Stereotactic Interest Group of Australasia (SIGA) Inaugural Meeting Melbourne, Australia November 8, 2018

Panelist What radiosurgery system will best suit the needs of my department? Stereotactic Interest Group of Australasia (SIGA) Inaugural Meeting Melbourne, Australia November 8, 2018

Special Guest Panelist (Together with Drs. Albert Koong, Brian Kavanagh, Paul Keall, and Fiona McDonald) Challenging cases in oligometastases Peter MacCallum Cancer Centre Stereotactic Ablative Radiotherapy (SABR) Symposium Melbourne, Australia November 9, 2018

Moderator Lung BOOST- Case-based review November 2018 RSNA 2018, Chicago, IL

Faculty (together with Drs. Kristin Redmond and William Yuh) CNS BOOST- E-contouring for spine SBRT November 2018 RSNA 2018, Chicago, IL

Speaker (Together with Dr. Kristin Redmond from Johns Hopkins University) Special ASTRO webinar E-contouring: Spine SBRT and CNS Invited by ASTRO February 26, 2019

Invited faculty (Declined due to time conflict) IAEA SBRT course Hiroshima, Japan February 2019

Invited speaker (Declined due to time conflict)

Langendorff Symposium Freiburg, Germany March 2019

Panel Co-Chair and Speaker SRS/SBRT: Building Collaborative Teams – Exchanging Experiences and Addressing Challenges Building a spine oncology program from scratch Radiosurgery Society Meeting 2019 San Diego, CA March 22, 2019

Co-Moderator with Dr. Najeeb Mohideen Showdown Debate between Dr. Shankar Siva and Dr. Robert Aboussaly SBRT vs Focal Ablation Therapy for Primary Renal Cell Carcinoma – Let the Madness Begin Radiosurgery Society Meeting 2019 San Diego, CA March 22, 2019

Speaker ARS Bone Metastasis Appropriate Use Committee Report American Radium Society Meeting Dana Point, CA April 7, 2019

Panel Chair and Speaker (Panelists: Simon S Lo, E Antonio Chiocca, Tracy Batchelor, Andrew Sloan) Bringing New and Innovative Glioblastoma Treatments to Daily Practice American Society of Clinical Oncology Annual Meeting ' Chicago, IL June 3, 2019

Faculty (Together with Drs. Kristin Redmond and Gregory Videtic) E-contouring for SBRT for oligometastases ASTRO Annual Meeting Chicago, IL September 15, 2019

Discussant Scientific Session: Palliative 1 ASTRO Annual Meeting Chicago, IL September 16, 2019

Panelist (Together with Drs. Yolanda Tseng, Candice Johnstone, Peter Hoskin, Alyssa Fairchild) Stat Rads! Turn the Beam On! Challenging Palliative Radiation Emergencies ASTRO Annual Meeting Chicago, IL September 17, 2019

Invited speaker SRS for large brain metastases- Preop, postop or no op? Hong Kong International Oncology Symposium Hong Kong SAR October 26, 2019

Moderator (Speakers: Shankar Siva and Rodney Ellis) SBRT for primary renal cell carcinoma Radiosurgery Society Webinar November 6, 2019

Panelist Oncodiagnosis Panel: Renal Cell Carcinoma RSNA Annual Meeting Chicago, IL December 2019

Panelist Lung BOOST- Case-based review RSNA Annual Meeting Chicago, IL December 2019

Panelist (Together with Tithi Biswas and Smith Apisarnthanarax) Post-Stereotactic-Radiation Imaging Assessment: Spine & GI SBRT Radiosurgery Society Webinar December 12, 2019

Invited speaker Post-SBRT imaging evaluation- Spinal metastasis Radiosurgery Society Meeting Washington, DC June 2020 (Virtual Meeting)

Invited panelist and speaker SRS/ SBRT in the time of COVID-19 Radiosurgery Society Webinar April 21, 2020 (743 attendees online)

Invited speaker (together with Dr. Kristin Redmond from Johns Hopkins) Strategies to Guide Safe Delivery of Spinal SBRT Radiosurgery Society Webinar June 30, 2020

Invited speaker Managing adverse effects of AVM radiosurgery 4th Biennial Radiosurgery Research and Education Meeting International Radiosurgery Research Foundation (IRRF) Uniondale, New York June 20, 2020

Invited speaker (together with Drs. Shankar Siva, Alexander Louie, and Nicholas Zaorsky) The Role of Stereotactic Radiotherapy in Renal Cell Carcinoma - Radioresistance or Resistance to Radiate? ASTRO Annual Meeting Miami Beach, FL October 2020

Invited speaker Stereotactic Body Radiotherapy for Spinal Metastases- The Current State of Affairs and Future Directions Hong Kong International Oncology Symposium November 8, 2020

Invited panelist (together with S. R. Digumarthy, J.D. Patel, D. Johnstone, F. Kong) BOOST: Lung, Mediastinum, Pleura—Case-based Multidisciplinary Review (Interactive Session) RSNA Annual Meeting Chicago, IL November 2020

Keynote Guest Speaker Singapore Radiation Oncology Annual Dinner Symposium December 17, 2020

Speaker Radiosurgery Society Distinction in Stereotactic Radiotherapy ACRO Annual Meeting February 27, 2021

Moderator (Speakers: Arjun Sahgal and Kristin Redmond) Webinar: The road to establishing SBRT as the standard of care for limited spinal metastases Radiosurgery Society April 27, 2021

Faculty speaker Sparing of the spinal cord: Limitations in treatment planning Spine Tumor Academy Switzerland (Virtual) April 24, 2021

Invited Faculty Postoperative Spine: Aim before your shoot- Don't shoot from the hips!/ Expert Wrap Up/ Panel Discussion RadoncAsia Singapore On-Target (SPOT) Contouring Workshop Series 2021 August 14-15, 2021

Moderator, Panel Chair and Speaker Judicious use of SRS/ HSRT for large brain metastasis ASTRO October 2021

Invited Speaker Advanced raditotherapy- A brave new world or a costly mistake? Hong Kong International Oncology Symposium November 2021

Panel Lead Lung Cancer: Case-Based Review RSNA November 29, 2021 Moderator Scientific Session: Radiation Oncology (CNS/ Head and Neck) RSNA November 30, 2021

Moderator Scientific Session: Radiation Oncology (Lung) RSNA December 2, 2021

Panelist (Co-Panelists: Stan Benedict, Brain Lally, Jaroslaw Hepel, and Fraser Cobb) Distinction in Practice in SRT Accreditation Program ACRO/ Radiosurgery Society Recorded on December 7, 2021

Invited Faculty Spinal Oncology- Radiation Oncology Perspective ASSR Spine Oncology webinar December 18, 2021

Co-moderator with Dr. Percy Lee Showdown debate: Single fraction vs Multifraction SBRT for Spinal Metastases Radiosurgery Society Meeting March 5, 2022 San Diego

Speaker Managing the Prevention of SRS/ SBRT-Induded Toxicities: SBRT for Spinal Lesions Radiosurgery Society Meeting March 5, 2022 San Diego

Speaker Performance and Quality Imporvement Session Radiosurgery Society Meeting March 6, 2022 San Diego

Invited Special Guest Speaker (Invited by Profeessor Yasushi Nagata) Japanese Society for Radiation Oncology November 10-12, 2022 Hiroshima, Japan

#### Courses/ educational sessions developed for national meetings:

## **Radiological Society of North America**

Radiological Society of North America 2015 (together with Dr. Stephanie Terezakis from Johns Hopkins University): 5 refresher courses in radiation oncology

1 refresher course in radiation safety (both diagnostic radiology and radiation oncology)

1 Controversy Session

1 Oncodiagnosis Panel 1 CNS BOOST contouring session Total: 9 courses

Radiological Society of North America 2016 (together with Dr. Stephanie Terezakis from Johns Hopkins University and Dr. Christina Tsien from Washington University):
5 refresher courses in radiation oncology
1 Controversy Session
1 Oncodiagnosis Panel
1 Lung BOOST Case Based Review

- 1 Lung BOOST Anatomy and Contouring
- Total: 9 courses

Radiological Society of North America 2017 (together with Dr. Christina Tsien from Washington University):
5 refresher courses in radiation oncology
2 Controversy Sessions
1 Oncodiagnosis Panel
1 Lung BOOST Case Based Review
1 Lung BOOST Anatomy and Contouring
1 Lung BOOST Didactic
Total: 11 courses

Radiological Society of North America 2018 (together with Dr. Christina Tsien from Washington University):
5 refresher courses in radiation oncology
One Controversy Session
1 Oncodiagnosis Panel
1 Lung BOOST Case Based Review
1 Lung BOOST Anatomy and Contouring
1 Lung BOOST Didactic
1 BOOST Spinal SBRT e-contouring
Total: 11 courses

Radiological Society of North America 2019 (together with Dr. Christina Tsien from Washington University):
5 refresher courses in radiation oncology
One Controversy Session
1 Oncodiagnosis Panel
1 Lung BOOST Case Based Review
1 Lung BOOST Anatomy and Contouring
1 Lung BOOST e-contouring
Total: 10 courses

Radiological Society of North America 2020 1 Oncodiagnosis Panel 1 Lung BOOST Case Based Review Total: 2 courses

Radiological Society of North America 2021 (together with Dr. Sung Kim, Anna Shapiro and Suresh Mukherji)
7 Case-Based reviews
3 Oncodiagnosis Panels
3 Scientific Sessions
Total: 13 courses

#### **American Society of Clinical Oncology**

American Society of Clinical Oncology 2019 CNS track 5 educational courses

American Society of Clinical Oncology 2020 CNS track 5 educational courses

American Society of Clinical Oncology 2021 CNS track 5 educational courses

#### **Radiosurgery Society**

Radiosurgery Meeting 2017 Member of Planning Committee

Radiosurgery Meeting 2019 Member of Planning Committee

Radiosurgery Meeting 2020 Member of Planning Committee

Radiosurgery Meeting 2021 Member of Planning Committee

Radiosurgery Meeting 2022 Member of Planning Committee

#### Invited and/or refereed local regional meetings (on behalf of co-authors and co-presenters, if applicable):

A comparison of stereotactic radiosurgery and fractionated stereotactic radiotherapy for the treatment of meningiomas: A seven year institutional experience at University of Minnesota (Oral presentation in the Minnesota Radiological Society Meeting on April 9, 2000).

Surgery impacts on survival of patients with low grade glioma treated with postoperative radiation therapy (Oral presentation in the Minnesota Radiological Society Meeting on April 8, 2001)

Is stereotactic radiotherapy adequate treatment for atypical and malignant meningiomas? (Oral presentation in Minnesota Radiological Society Meeting on April 6, 2002)

Radiosurgery for meningiomas and acoustic neuromas (Guest faculty at the VI Detroit Neurosurgery Symposium: The Renaissance of Neurosurgery, September 13-14, 2002)

Hodgkin Lymphoma- Role of Radiation Therapy (Invited speaker at Chicago Radiological Society, January 15, 2004)

Modern Radiation Techniques and their Application in Brain Tumors. 1<sup>st</sup> Annual Neuro-oncology Symposium: Recent Advances in high Grade Gliomas and the Impact on Treatment. Arthur G. James Cancer Hospital. September 16, 2006.

Extracranial Stereotactic Radiotherapy. Lung Cancer Care for 21<sup>st</sup> Century. Ohio State University Medical Center. March 3, 2007.

Advanced Radiation Therapy Modalities Invited speaker for OSU Medical Service Board October 18, 2007

Stereotactic Body Radiation Therapy. Southern Ohio Cancer Center Grand Round October 19, 2007

Spinal Metastases- Radiotherapy Perspective Spine and Bone Tumor Symposium OSU comprehensive cancer center May 3, 2008

American Society of Clinical Oncology review Arthur G. James Cancer Hospital June 21, 2008

Stereotactic Body Radiation Therapy for Early Stage Non-Small Cell Lung Cancer Today's Multidisciplinary Approach to Lung Cancer St. Vincent's Health System, Erie, PA An affiliate of University of Pittsburgh Medical Center May 16, 2009

American Society for Radiation Oncology (ASTRO) Review Case Comprehensive Cancer Center (Joint Cleveland Clinic Foundation/ UH Seidman Cancer Center) November 15, 2014

CNS and Radiosurgery Update Washington State Radiological Society Annual Meeting November 5, 2016

#### Media

https://www.dispatch.com/article/20080120/news/301209867

LA times- http://articles.latimes.com/2007/dec/31/health/he-unreal31

Reuters Health- <u>https://www.journalofclinicalpathways.com/news/stereotactic-radiosurgery-after-complete-resection-brain-mets-improves-outcomes</u>

https://www.doximity.com/doc\_news/v2/entries/19873369

https://physicsworld.com/a/how-has-covid-19-impacted-the-provision-of-radiation-therapy/

https://www.astro.org/ASTRO/media/ASTRO/AffiliatePages/arro/PDFs/ARROCase\_SpineSBRT.pdf?utm\_source=Magne tMail&utm\_medium=email&utm\_term=INSERT\_EMAIL&utm\_content=INSERT\_MESSAGE\_NAME&utm\_campaign= INSERT\_MESSAGE\_SUBJECT

#### Health Technology Clinical Committee Health Care Authority Conflict of Interest Disclosure

As stewards of public funds, the practicing clinicians who serve (or apply to serve) on the Committee strive to uphold the highest standards of transparency and impartiality. Identifying financial, professional, and other interests contribute to the effective management of perceived, potential, and/or real conflicts of interest/bias that could affect Committee determinations. (WAC 182-55)

This Conflict of Interest form must be completed by an applicant for appointment to the State of Washington Health Technology Clinical Committee (HTCC) or appointment to any of its subcommittees or work groups.

A member of the HTCC or any of its subcommittees or work groups may not participate in discussions or deliberations of any class of drugs, health technology, or any agenda item for which a conflict of interest is identified and may not vote on any such matter.

If a conflict of interest is so great as to make it difficult for any member to participate meaningfully in the work of the HTCC, that member may be asked to resign.

1	Applicant information	
First name:		Middle initial:
Last name:		
Phone number:	Email:	
2	Financial interests	

Disclose your financial interests and relationships occurring over the last twenty-four months.

**List amounts totaling** \$1,000 or more from a single source.

**Indicate the category** of financial interest/relationship by referring to the disclosure categories below. Select the letter corresponding to your financial interest(s). You may indicate multiple categories.

**Indicate the source and date** of the financial interest. For each chosen category, include date and if your activities are ongoing.

**Indicate the recipient.** Family: spouse, domestic partner, child, stepchild, parent, sibling (his/her spouse or domestic partner) currently living in your home.

#### **Financial interest categories**

Use these categories to indicate the nature of the financial interest:

- A. Payment from parties with a financial or political interest in the outcome of work as part of your appointment or activity.
- B. Employment including work as an independent contractor, consultant, whether written or unwritten.
- C. Ownership or owning stock (stock, options, warrants) or holding debt or other significant proprietary interests or investments in any third party that could be affected.
- D. Receiving a proprietary research grant or receiving patents, royalties, or licensing fees.
- E. Participating on a company's proprietary governing boards.
- F. Participating in a speakers bureau.
- G. Receiving honoraria.

Please list your financial interests on the next page. Attach additional sheets if necessary.

#### **Financial interest disclosures**

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Category (A-G)	Source of income and date	Amount	Recipient	
G	Japanese Society for Radiation Oncology	US\$1,000	✓ Self	Family
			Self	Family
			Self	Family
			Self	Family
			Self	Family
			Self	Family
			Self	Family

**Other interests** 

Please respond to the following questions. Disclose all interests that may apply to topics covered in upcoming meetings.

Have you authored, coauthored, or publicly provided an opinion, editorial, or publication related to any meeting topic? Topics(s):

Yes. I have published numerous papers and edited textbooks on SBRT for various sites including lung, liver, kidney, spine, prostate, adrenal, head and neck and gynecologic cancers.

#### Are you involved in formulating policy positions or clinical guidelines related to any meeting topic? Topics(s):

Yes, I have led or participated in multiple clinical guidelines for SBRT for various disease sites and have participated in the renewal of the American Society for Radiation Oncology (ASTRO) Model Policy for SBRT.

Could a coverage determination based on a Committee decision conflict with policies you have promoted or are obliged to follow? Topic(s):

No.

Signature

I have read the Conflict of Interest Disclosure form. I understand the purpose of the form and agree to the application of the information to determine conflicts of interest. The information provided is true and complete as of the date the form was signed. If circumstances change, I am responsible for notifying committee staff in order to amend this disclosure. I will complete this form annually by July 1st of each year of committee membership.

To sign this request, do not use the "Fill & Sign" function; instead, simply click in the signature field to add your signature.

Signature

Date

11/15/2022

Or mail to: Health Technology Assessment Program Washington State Health Care Authority P.O. Box 42712 Olympia, WA 98504-2712

or return form to shtap @hca.wa.gov

Submit

4

#### Health Technology Clinical Committee Application for Membership



1	Contact inf	ormation	
First name:			Middle initial:
Last name:			
Address:			
Phone number:		Best method, time to reach you:	
Email:		Today's date	
2	Personal in	formation (optional)	
Gender:			
Male Female	X/non-binary <sup>1</sup>		
Pronouns (select all that apply)			
She/her He/him	They/them	Other (subj./obj.):	
Race or Ethnicity			
American Indian or Alaska	Native	Asian or Pacific Islander America	n
Black/ African American		Latino, Hispanic, Spanish	
White/ Caucasian		Other:	
3	Profession	al training	
Education (list degrees):			
Health care practitioner license	s:		
Professional affiliations:			
Board certifications, formal trai	ning, or other designc	ations:	
Current position (title and empl	oyer):		
Current practice type and years	in practice:	Total years as an active practitioner:	
Location of practice (city):			

<sup>1</sup> Non-binary (X) is an umbrella term used to describe those who do not identify as exclusively male or female. This includes but is not limited to people who identify as genderqueer, gender fluid, agender, or bigender.

4

#### Experience

Provide a brief explanation (up to 150 words each) addressing the following:

1) Why you would like to serve on the clinical committee;

2) The value of informing health policy decisions with scientific evidence, including any examples incorporating new evidence into your practice;

3) How your training and experience will inform your role on the committee

4) Treating populations that may be underrepresented in clinical trials: women, children, elderly, or people with diverse ethnic and racial backgrounds, including recipients of Medicaid or other social safety net programs?

#### Ability to serve

Are you able to participate in all-day meetings, an estimated six times per year? Are you willing to commit to the responsibilities of a committee member, including:	Yes	No
<ul> <li>Attending meetings prepared for the topics of the day;</li> </ul>		
<ul> <li>Actively participating in discussions;</li> </ul>		
<ul> <li>Making decisions based on the evidence presented and the public interest<sup>1</sup>?</li> </ul>	Yes	No
Could you, or any relative, benefit financially from the decisions made by the HTCC?	Yes	No
6 References		

Provide three professional ref	erences:	
<b>1.</b> First name:	Last name:	
Relationship:	Title:	
Contact email:	Phone number:	
<b>2.</b> First name:	Last name:	
Relationship:	Title:	
Contact email:	Phone number:	
<b>3.</b> First name:	Last name:	
Relationship:	Title:	
Contact email:	Phone number:	

#### Please return:

Completed application

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curriculum vitae

conflict of interest disclosure

#### to send via email to: shtap@hca.wa.gov

OR mail to: Health Technology Assessment Program Washington State Health Care Authority P.O. Box 42712 Olympia, WA 98504-2712

<sup>1</sup> Detailed in Washington Administrative Code (WAC) and committee bylaws



## Washington State Health Care Authority

Agency medical director comments

### **Stereotactic Body Radiation Therapy**

**Sophie Cain Miller, MD MPH** Medical Officer for Medicaid WA Health Care Authority

May 19, 2023







## **Radiation Therapy**

#### • External beam radiation therapy

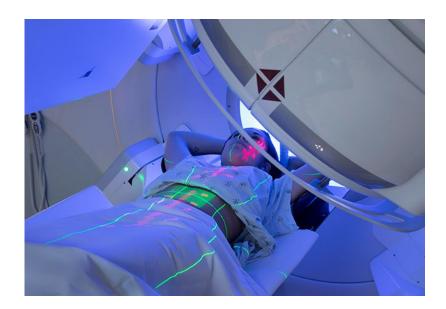
- Utilizes machine to aim high doses of radiation at cancer cells
- Local treatment to specific areas of tumor
- Internal radiation therapy
  - Source of radiation is put inside body (seeds, capsules)
  - Also known as "brachytherapy"



Washington State Health Care Authority

## Stereotactic Body Radiation Therapy (SBRT)

- Radiation treatment modality
- High anatomic targeting accuracy
  - Target lesion (tumor) is localized in a three dimensional system
  - Imaging modalities utilized for planning: CT, MRI, PET
- High doses of precise radiation in fewer fractions
  - One to five sessions (fractions)
  - 6-30Gy

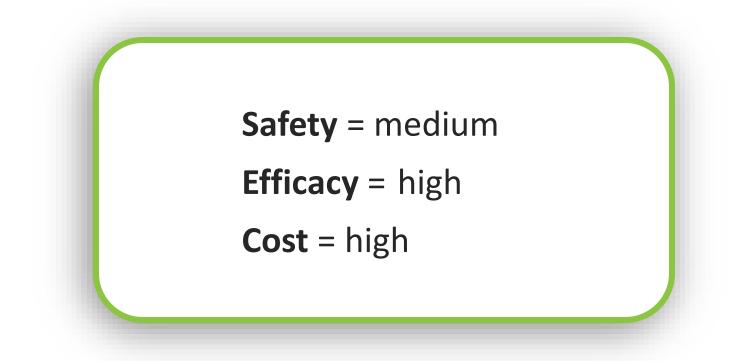








### Agency medical director concerns - overall



Note: concerns noted above from initial review in 2012







## **Previous HTCC decisions**

- 2012 evidence review of stereotactic radiosurgery (SRS) and SBRT
- Coverage for:
  - Central nervous system primary and metastatic tumors
  - Cancers of the spine and paraspinal structures
  - Inoperable Stage 1 non small cell lung cancer
- All evaluations should include a multidisciplinary team analysis including surgical input







## Current State Agency Policy on SBRT

- Uniform Medical Plan
- Medicaid
- Labor and Industries

Follow 2012 HTCC decision





## SBRT: HTCC Re-review Rationale

- Updated evidence search in 2016 and 2018
  - During those reviews, not sufficient evidence to review coverage policy decision
- Chosen for re-review in 2022/2023 due to new evidence that could prompt coverage policy changes

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Evidence for expanding use for other cancer types





### **Key Questions**

- What is the evidence of effectiveness for SBRT for patients with cancers not currently covered (CNS cancers and inoperable stage 1 NSCLC)?
- What are the harms of SBRT in patients with included cancers?
- What is the evidence that SBRT has differential efficacy or harms in subpopulations, including those defined by:
  - Sex, age, site and type of cancer, stage and grade of cancer
  - Setting, provider characteristics, equipment, quality assurance standards, procedures
- What is the evidence of cost and cost-effectiveness of SBRT?







### **Agency Experience Data**

#### Washington State – Combined Medicaid & PEBB/SEBB UMP

	2018	2019	2020	2021	Total (unique)
Individuals with at least one SBRT-related procedure/service	182	234	217	225	787
Female, count	92	116	114	135	421
Male, count	88	115	103	88	359
Number of encounters with SBRT	975	1,343	1,174	1,206	4,698
Amount paid, SBRT	\$800,066	\$1,223,446	\$1,196,542	\$908,197	\$4,128,250
Amount paid, SBRT and related procedures	\$1,164,151	\$1,682,401	\$1,657,914	\$1,375,493	\$5,879,959

Data notes: Small numbers suppressed from L&I to protect patient privacy. Claimant sex was not always reported. Annual members for Medicaid excludes members that are dually eligible for Medicaid and Medicare. Related procedures for amount paid reflects all claims submitted with the procedure code for the same date of service, and includes professional, facility, and ancillary claims (such as durable medical equipment). Managed care amount paid reflects an estimate of the amount paid for the procedure. UMP data does not reflect patient cost share. Individuals who had a procedure in more than one year are only counted once in the "Total" summary. Amounts paid of \$0 were excluded from amount paid table value calculations.

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### Cost by CPT code

Code	de Description		Medicaid FFS		L&I	
СРТ		Non-facility	Facility	Non-facility	Facility	
77373	Stereotactic body radiation therapy, treatment delivery, per fraction to 1 or more lesions, including image guidance, entire course not to exceed 5 fractions	\$692.98	\$692.98	\$2,102.93	\$2,102.93	
77435	Stereotactic body radiation therapy, treatment management, per treatment course, to 1 or more lesions, w/ image guidance, max 5 fractions	\$367.87	\$367.87	\$1,082.15	\$1,082.15	

Data notes: Medicaid FFS from October 1, 2021 Physician-Related Services <u>Fee Schedule</u> (accessed January 18, 2023; <u>webpage</u>). L&I from <u>2021 provider fee schedule</u> (accessed January 18, 2023). PEBB/UMP fees are confidential and not publicly available (proprietary).

#### Copyright Statement

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## **Evidence Considerations**

- For each type of cancer, different bodies of evidence
- Limited number of RCTs  $\rightarrow$  Lower certainty of evidence
  - 12 RCTs in total (in 21 publications) across all cancer types
- Comparison groups
  - Differ by study and cancer type
  - Examples: convention radiation, medication only
- How outcomes are measured
  - Survival disease free, progression free
  - Disease control
  - Quality of life







Safety

Precision and accuracy needed due to high radiation dose per fraction

- Location of Treatment
  - High doses delivered to target area
  - Margin in millimeters
  - Adjacent organs at risk (OARs)
- Equipment
- Staffing
  - Radiation oncologist and medical physicist
  - Specialized training is encouraged







- SBRT not associated with significantly higher rates of toxicity compared with other treatment modalities
- Type of toxicity varies by treatment site
- Severe and/or life-threatening toxicities were rare









- Limited studies on cost. Specific to cancer type and comparison group (cRT, medication, etc)
- Specific examples include:
  - Cost savings for low and intermediate risk prostate cancer, along with improved access to radiation treatments



Washington State Health Care Authority

### Payer Coverage

Condition	United	Aetna	Cigna
Prostate	$\checkmark$	$\checkmark$	$\checkmark$
Lung	$\checkmark$	$\checkmark$	$\checkmark$
Pancreatic adenocarcinoma	$\checkmark$	$\checkmark$	$\checkmark$
Liver/HCC	$\checkmark$	$\checkmark$	$\checkmark$
Oligometastatic	$\checkmark$	$\checkmark$	$\checkmark$
Kidney	Oligometastatic disease only	Х	Х
Adrenal	Х	Х	Х
Head and Neck	Х	Recurrent	Recurrent
Bone	Х	Mets to spine only	Х
A STATE OF A	lotos: V = covorod V =	wat as your al	

Notes:  $\checkmark$  = covered, X = not covered



## Professional Society and other Guidelines

- American Society for Therapeutic Radiation Oncology (ASTRO)
- National Comprehensive Cancer Network (NCCN)
- American Society of Clinical Oncology (ASCO)
- Society for Interventional Radiology (SIR)
- European Society for Medical Oncology (ESMO)
- National Institute for Health and Care Excellence (NICE)



Washington State Health Care Authority

### Professional Society Guidelines (1)

Condition	ASTRO	NCCN	Other
Prostate	Recommended for Low- or intermediate-risk	<ul> <li>Acceptable efficacy and toxicity for low, intermediate, high, very high-risk groups</li> </ul>	<ul> <li>AUA: strong recommendation, evidence level: Grade A</li> <li>Other organizations have less strong guidance</li> </ul>
Lung, NSCLC	Recommended for early stage inoperable NSCLC with evaluation by multidisciplinary team	<ul> <li>Medically inoperable early-stage NSCLC</li> <li>Improved local control and overall survival compared to cRT</li> <li>"Reasonable alternative" to operable but high risk</li> </ul>	<ul> <li>ASCO: Stage I or IIA (node negative), and in operable or patient deemed high risk</li> <li>SIR: Stage IA</li> <li>ESMO: Stage IV with limited synchronous metastases</li> <li>NICE: Stage I-IIA who decline surgery. Stage IIIA who decline or can't tolerate chemo</li> </ul>
Pancreatic adenocarcino ma	<ul> <li>Resectable, only recommended on clinical trial or multi-institutional registry</li> <li>Borderline resectable and select locally advanced appropriate for downsizing, conditionally recommended</li> <li>Locally advanced, not appropriate for downsizing, conditional recommendation</li> </ul>	<ul> <li>Option for first line therapy for locally advanced disease, good performance status</li> <li>Not candidates for induction chemo</li> <li>Treatment for recurrence</li> </ul>	<ul> <li>ASTRO caveats: adjuvant SBRT post operatively recommended in clinical trial setting</li> </ul>
	recommendation	Notes: See pages 117 – 130 c	of SBRT final report for complete guideline information



### Professional Society Guidelines continued (2)

Condition	ASTRO	NCCN	Other
Liver/HCC	<ul> <li>Candidates for OLT, conditionally recommended as a bridge to transplant or downstaging</li> <li>Liver confined HCC, recommended</li> </ul>	<ul> <li>Alternative to ablation/embolization when these therapies have failed/contraindicated</li> <li>Unresectable or inoperable</li> </ul>	<ul> <li>ACR: Hepatocellular cancer with solitary tumor 0-3cm "may be appropriate"</li> <li>ESMO: may be considered as alternative for ablation of tumors with high risk of local failure</li> </ul>
Oligometastatic		Based on cancer type	
Kidney		May be considered for medically inoperable stage I or stage II	<ul> <li>ESMO: Option for management in metastatic disease</li> <li>Limited other guidance</li> </ul>
Adrenal		Can consider	
Head and neck		Insufficient evidence to recommend	
Bone		Consider for oligometasteses	• ESMO: Patients unfit for surgery

Notes: See pages 117 – 130 of SBRT final report for complete guideline information



Washington State Health Care Authority

### **Cost-effectiveness**

Condition	Cost-effectiveness Considerations	Certainty
Prostate	May be lower in cost than IMRT May be cost effective for oligometastatic hormone resistant cancer	N/A
Lung	May be cost ineffective for maintenance therapy for oligometastatic disease	N/A
Pancreatic adenocarcinoma	May be higher cost than cRT or chemo - very lower certainty of evidence	Very low
Liver/HCC	May be cost ineffective compared to RFA, however is cost effective for salvage therapy- low to moderate certainty of evidence	Low to moderate
Oligometastatic	May be cost-effective, moderate certainty of evidence based on 2 modeling studies	Moderate
Kidney	N/A	N/A
Adrenal	N/A	N/A
Bone	N/A	N/A

#### Localized Prostate cancer: covered with conditions

SBRT is a covered benefit for:

- Very low, low and intermediate risk prostate cancer, as defined by NCCN based on stage, Gleason score and PSA level
- Evaluation includes multidisciplinary team analysis with surgical input



# Non Small Cell Lung Cancer (NSCLC): covered with conditions

- SBRT is a covered benefit for :
  - non small cell lung cancer
    - Stage I and Stage IIA (node negative)
       AND
  - Medically inoperable, or patient is deemed too high risk, or declines operative intervention
- Evaluation includes multidisciplinary team analysis with surgical input



ity

Pancreatic Adenocarcinoma: covered with conditions

- SBRT is a covered benefit for:
  - Locally advanced pancreatic adenocarcinoma,
- Non covered for those with evidence of direct invasion of bowel or stomach based on CT, MRI or endoscopy
- Evaluation includes multidisciplinary team analysis with surgical input

ity

### **Oligometastatic disease:** covered with conditions

- SBRT is a covered benefit for:
  - Oligometastatic disease as defined by:
    - Three or fewer metastatic lesions in synchronous setting
    - Appropriate imaging demonstrating no evidence of widespread metastatic disease
    - Karnofsky performance score greater than or equal to 60, or ECOG score less than or equal to 2
  - Evaluation includes multidisciplinary team analysis with surgical input

### Hepatocellular carcinoma: covered with conditions

- SBRT is a covered benefit for:
  - Hepatocellular carcinoma
    - 5 or fewer lesions, 6cm or smaller

AND

• Karnofsky performance score greater than or equal to 60, or ECOG score less than or equal to 2

OR

- Unresectable/inoperable disease, not a candidate for liver transplant
- Evaluation includes multidisciplinary team analysis with surgical input



SBRT is not a covered benefit for treatment of the *primary* tumor of the following cancer types:

- Renal
- Adrenal
- Bone
- Small cell lung cancer
- Melanoma
- Merkel Cell
- Biliary tract cancer
- Head and neck cancers
- Breast \*
- Ovarian\*
- Cervical \*
- Esophageal \*
- Colorectal \*

SBRT is a covered benefit for treatment of oligometastatic lesions.



### SUMMARY OF AMD COVERAGE RECOMMENDATIONS

Condition(s)	Recommendations
Prostate	SBRT is a covered benefit for very low, low and intermediate risk prostate cancer as defined by NCCN based on stage, Gleason score and PSA Level
Lung	SBRT is a covered benefit for non small cell lung cancer Stage I and Stage IIA, tumors considered medically inoperable, or patient is deemed too high risk, or declines operative intervention.
Pancreatic adenocarcinoma	SBRT is a covered benefit for locally advanced pancreatic adenocarcinoma. Non-covered for those with evidence of direct invasion of bowel or stomach based on CT, MRI or endoscopy
Liver/HCC	SBRT is a covered benefit for hepatocellular carcinoma in the following settings: 5 or fewer lesions, 6cm or smaller, and Karnofsky performance score greater than or equal to 60, or ECOG score less than or equal to 2 OR for unresectable/inoperable disease, not a candidate for liver transplant
Oligometastatic	SBRT is a covered benefit for oligometastatic disease as defined by three or fewer lesions in a synchronous setting, with appropriate imaging demonstrating no evidence of widespread metastatic disease. KS greater than or equal to 60, ECOG less than or equal to 2.
Renal, Adrenal, Bone, SCLC, Melanoma, Merkel Cell, Biliary tract cancer, head and neck cancer, breast, ovarian, cervical, esophageal, colorectal	SBRT is not a covered benefit.





## Questions?

More Information:

Sophie Cain Miller, MD, MPH Sophie.miller@hca.wa.gov







Cancer Type	Prostate cancer, localized
Evidence	<ul> <li>2023 upate-4 RCTs, 14 comparative studies, 18 noncomparative studies</li> <li>RCTs w adequate sample size</li> <li>Similarly or more effective than other options</li> <li>Low to moderate certainty of evidence</li> </ul>
Other payors	United, Aetna, Cigna
ASTRO	Recommended, low or intermediate risk
NCCN	"acceptable efficacy and toxicity" for low, intermediate, high, very high risk groups
Cost	May be lower in cost than IMRT May be cost effective for oligometastatic hormone resistant cancer
Other considerations	Toxicity- some evidence for increased urinary obstruction/retention

Washington State Health Care Authority

Cancer Type	Non Small Cell Lung Cancer
Evidence	<ul> <li>2023 update:- 1 RCT, 11 comparative studies, 11 non comparative studies</li> <li>Similarly or more effective than cRT</li> <li>Low to moderate certainty of evidence</li> </ul>
Other payors	United, Aetna, Cigna
ASTRO	Early stage medically inoperable
NCCN	<ul> <li>Medically inoperable early stage NSCLC</li> <li>Improved local control and overall survival compared to cRT</li> <li>"Reasonable alternative" to operable but high risk</li> </ul>
Cost	May be cost ineffective for maintenance therapy for oligometastatic disease
Other considerations	For pts at high surgical risk recommend multidisciplinary evaluation

Washington State Health Care Authority

Cancer Type	Liver cancer/HCC			
Evidence	<ul> <li>2023 update- 20 comparative studies</li> <li>May be as effective as radiofrequency ablation for early-stage liver cancer- results mixed</li> <li>Alone or in combo with other treatments for small liver cancers</li> <li>May be more effective than sorafenib for advanced liver cancer</li> <li>Low to very lower certainty of evidence</li> </ul>			
Other payors	United, Aetna, Cigna			
ASTRO	Recommended for coverage			
NCCN	Alternative to ablation/embolization when these therapies have failed/contraindicated Unresectable or inoperable			
Cost	May be cost ineffective compared to RFA, however is cost effective for salvage therapy- low to moderate certainty of evidence			
Other				



Cancer Type	Oligometastatic
Evidence	<ul> <li>2023 update- 3 RCTs, 3 comparative studies, 12 non comparative studies</li> <li>More effective than standard of care or observation</li> <li>Low to moderate certainty of evidence</li> <li>(except for prostate Ca)</li> </ul>
Other payors	United, Aetna, Cigna
ASTRO	Recommended for coverage
NCCN	Based on cancer type
Cost	May be cost-effective, moderate certainty of evidence based on 2 modeling studies
Other considerations	Detailed criteria for defined patient population





Cancer Type	Pancreatic adenocarcinoma
Evidence	<ul> <li>2023 update-3 comparative studies</li> <li>May be more effective than cRT</li> <li>Low certainty of evidence</li> </ul>
Other payors	United, Aetna, Cigna
ASTRO	Conditionally recommended for borderline resectable, locally advanced
NCCN	<ul> <li>Option in list of first line therapy- following chemo or in patients not candidates for chemo</li> <li>Treatment for recurrence</li> <li>Good performance status</li> </ul>
Cost	May be higher cost than cRT or chemo - very lower certainty of evidence
Other considerations	SBRT should be delivered at an "experienced, high-volume center" per NCCN





Cancer Type	Kidney cancer
Evidence	<ul> <li>2012- no eligible studies</li> <li>2023 update: 1 comparative study, N=90,000</li> <li>For stage I RCC, SBRT associated with worse overall survival compared to ablation or surgery, low certainty of evidence</li> </ul>
Other payors	United for oligometastatic disease only
ASTRO	
NCCN	May be considered for medically inoperable stage I or stage II
Other considerations	







Cancer Type	Adrenal
Evidence	<ul> <li>2023 update: 1 noncomparative study, harms only. High risk of bias due to lack of comparator</li> </ul>
Other payors	Not covered
ASTRO	
NCCN	"can consider"
Other considerations	







Cancer Type	Head and Neck Cancers
Evidence	<ul> <li>2023 update: 1 RCT, 4 comparative studies</li> <li>May be similarly effective to brachytherapy</li> <li>May be less effective than charged particle RT</li> <li>Similar in effectiveness to IMRT</li> </ul>
Other payors	Aetna, Cigna (previously irradiated sites)
ASTRO	
NCCN	<i>"currently insufficient evidence to recommend SBRT for treatment of H&amp;N cancers"</i>
Other considerations	
	35





Cancer Type	Bone
Evidence	In 2012 review, no eligible studies. 2023 review, single center RCT, N=160
Other payors	Cigna (mets to spine only)
ASTRO	
NCCN	
Other considerations	Spinal lesions already covered by 2012 HTCC decision





#### Stereotactic Body Radiation Therapy

#### Order of scheduled presentations:

	Name
1	<ul> <li>Providence Swedish Radiosurgery Center</li> <li>Christopher Loiselle, MD</li> <li>Robert Meier, MD</li> <li>Daniel Landis, MD</li> </ul>
2	UW Medicine Radiology group • Edward Kim, MD • John Kang, MD, PhD • Jing Zeng, MD • Smith "Jim" Apisarnthanarax, MD • Ramesh Rengan, MD, PhD
3	David Kantorowitz, MD

Providence SWEDISH

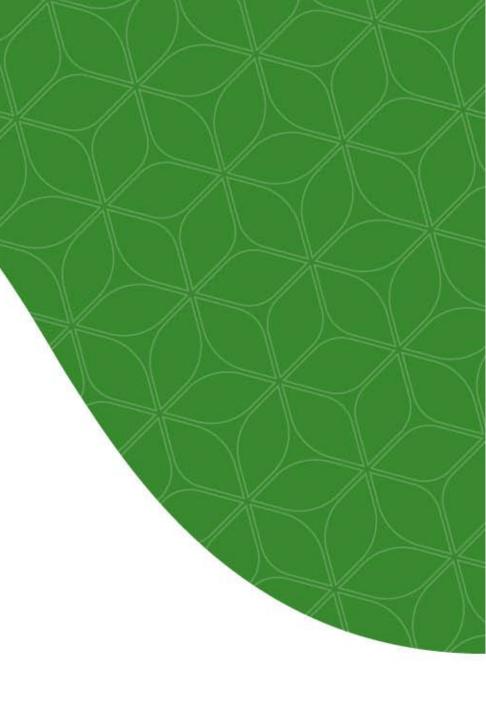
Christopher Loiselle, MD Executive Medical Director

Robert Meier, MD Radiation Oncology Medical Director

Daniel Landis, MD, PhD Radiation Oncology

Providence Swedish Radiosurgery Center CyberKnife |Gamma Knife

WA – HTA – Stereotactic radiation surgery (SRS) and stereotactic body radiation therapy (SBRT) – public comment meeting
May 19, 2023



- Prostate cancer
  - Page 114 of Evidence Report
    - "SBRT similarly or more effective than other options for men with localized prostate cancer"
    - Excellent example of need for modern policy update
    - SBRT has become a clinical standard of care for prostate cancers
- Oligometastatic prostate cancer
  - Page 115 of Evidence Report
    - "Appears to be more effective than standard of care or observation for oligometastatic cancer, however, for oligometastatic prostate cancer, elective nodal radiation..."
    - Context of care is necessary
      - Few oligometastatic prostate cancer patients are candidates for elective nodal radiation (node only pelvic disease)
      - Comparative studies of elective nodal radiation are not the same level of evidence as randomized trials for SBRT
- Oligometastatic cancer
  - Additional prospective randomized studies: CURB trial, Extend trial
    - Evidence for treatment of oligometastatic disease has rapidly increased
    - Based on current evidence, Medicare and commercial policies cover SBRT for oligometastatic disease
- Trigeminal Neuralgia/Essential Tremor
  - Missing in report
  - Standard of care
- Re-irradiation
  - Most policies cover SRS/SBRT in setting of prior radiation treatment
  - Standard of care



#### Health Technology Clinical Committee Health Conflict of Interest Disclosure



As stewards of public funds, the practicing clinicians who serve (or apply to serve) on the Committee strive to uphold the highest standards of transparency and impartiality. Identifying financial, professional, and other interests contribute to the effective management of perceived, potential, and/or real conflicts of interest/bias that could affect Committee determinations. (WAC 182-55)

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If a conflict of interest is so great as to make it difficult for any member to participate meaningfully in the work of the HTCC, that member may be asked to resign.

1	Applicant information	
First name:		Middle initial:
Christopher		
Last name:		
Loiselle		
Phone number:	Email:	

2

#### Financial interests

Disclose your financial interests and relationships occurring over the last twenty-four months. **List amounts totaling** \$1,000 or more from a single source.

**Indicate the category** of financial interest/relationship by referring to the disclosure categories below. Select the letter corresponding to your financial interest(s). You may indicate multiple categories.

**Indicate the source and date** of the financial interest. For each chosen category, include date and if your activities are ongoing.

**Indicate the recipient.** Family: spouse, domestic partner, child, stepchild, parent, sibling (his/her spouse or domestic partner) currently living in your home.

#### **Financial interest categories**

Use these categories to indicate the nature of the financial interest:

- A. Payment from parties with a financial or political interest in the outcome of work as part of your appointment or activity.
- B. Employment including work as an independent contractor, consultant, whether written or unwritten.
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- D. Receiving a proprietary research grant or receiving patents, royalties, or licensing fees.
- E. Participating on a company's proprietary governing boards.
- F. Participating in a speakers bureau.
- G. Receiving honoraria.

Please list your financial interests on the next page. Attach additional sheets if necessary.

Category (A-G)	Source of income and date	Amount	Recipient	
G	Accuray	\$15,000	✓ Self	Family
			Self	Family
			Self	Family
			Self	Family
			Self	Family
			Self	Family
			Self	Family

3

#### Other interests

Please respond to the following questions. Disclose all interests that may apply to topics covered in upcoming meetings.

Have you authored, coauthored, or publicly provided an opinion, editorial, or publication related to any meeting topic? Topics(s):

Yes, Treatment of Essential Tremor with Stereotactic Radiosurgery, IJROBP, Vol 90, Iss 1, 5167, 2014.

Are you involved in formulating policy positions or clinical guidelines related to any meeting topic? Topics(s):

No

Could a coverage determination based on a Committee decision conflict with policies you have promoted or are obliged to follow? Topic(s):

No

4

#### Signature

I have read the Conflict of Interest Disclosure form. I understand the purpose of the form and agree to the application of the information to determine conflicts of interest. The information provided is true and complete as of the date the form was signed. If circumstances change, I am responsible for notifying committee staff in order to amend this disclosure. I will complete this form annually by July 1st of each year of committee membership.

To sign this request, do not use the "Fill & Sign" function; instead, simply click in the signature field to add your signature.

Signature

Date

April 28, 2023

Or mail to: Health Technology Assessment Program Washington State Health Care Authority P.O. Box 42712 Olympia, WA 98504-2712

or return form to shtap @hca.wa.gov

Submit

#### Health Technology Clinical Committee Health Conflict of Interest Disclosure



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1	Applicant information	
First name:		Middle initial:
Daniel		М
Last name:		
Landis		
Phone number:	Email:	

2

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- F. Participating in a speakers bureau.
- G. Receiving honoraria.

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3

Category (A-G)	Source of income and date	Amount	Recipient	
			Self	Family

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Washington State Health Care Authority

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1	Applicant info	ormation	
First name:			Middle initial:
Robert			
Last name:			
Meier			
Phone number:		Email:	
	<b>—</b>		

2

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- G. Receiving honoraria.

Please list your financial interests on the next page. Attach additional sheets if necessary.

3

Category (A-G)	Source of income and date	Amount	Recipient	
g	Accuray 2022	\$2000.00	✓ Self	Family
			Self	Family
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			Self	Family

Other interests

Please respond to the following questions. Disclose all interests that may apply to topics covered in upcoming meetings.

Have you authored, coauthored, or publicly provided an opinion, editorial, or publication related to any meeting topic? Topics(s):

Yes, ESTRO 2021 OC-0509 10-year outcome of ultrahypofractionated stereotactic RT from two multicenter prostate cancer trials. Multicenter Trial of Stereotactic Body Radiation Therapy for Low-& Intermediate-Risk Prostate cancer: Int J Radiation Oncol Biol Phys, Vol. 102, No. 2, pp. 296-303, 2018

Are you involved in formulating policy positions or clinical guidelines related to any meeting topic? Topics(s):

No

Could a coverage determination based on a Committee decision conflict with policies you have promoted or are obliged to follow? Topic(s):

No			
4	Signature		

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Signature

Date

April 28, 2023

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Submit

# Ed Kim

### University of Washington Fred Hutchinson Cancer Center

May 19, 2023





## Sarcoma oligometastases

- Rare cancer with ~ 300 new patients in WA state per year
- Surgical resection of oligometastatic disease is standard of care for this uncommon malignancy
- Not all lesions are resectable
  - Medically inoperable
  - proximity to critical vascular structures
  - Location may require lobectomy with accompanying loss of lung function
- SBRT is endorsed as an appropriate treatment modality for oligometastases in the NCCN guidelines for sarcoma



### **UW** Medicine

## Recent Data supporting SBRT for sarcoma

- Retrospective series of 50 patients with 109 lesions treated with SBRT 2005 2021, 3 year local control rate of 88% and 3 year OS 50%. 1 year chemotherapy free survival was 85%. (Wafa Asha et al. Stereotactic Body Radiation Therapy for Sarcoma Pulmonary Metastases, Am J Clin Oncol. 2023 Mar 14. PMID: 36914598).
- Retrospective series of 26 oligometastases treated with SBRT 2010-2022. 2 yr LC 96%, 2 yr OS 74%. Median OS not met. (*Gutkin P, et al. Stereotactic body radiotherapy for metastatic sarcoma to the lung: adding to the arsenal of local therapy. Radiat Oncol. 2023 Mar 1;18(1):42. PMID: 36859309*)
- Retrospective series of 51 lesions treated with SBRT in 28 patients, with 5yr local control 96%, 2 yr OS 96% and 5yr OS 60.5%. (*Navarria, et al. Stereotactic body radiation therapy for lung metastases from soft tissue sarcoma. Eur J Cancer. 2015 Mar;51(5):668-74. PMID: 25686482*).
- Prospective phase 2 trial of 44 patients with 71 lesions. 1 yr LC 98.5%, median DFS 12 months, median OS 49 months. 1 yr OS 88.6% and 5yr OS 48.2%(Navarria et al. Stereotactic Body Radiation Therapy for Lung Metastases From Sarcoma in Oligometastatic Patients: A Phase 2 Study. Int J Radiat Oncol Biol Phys. 2022 Nov 15;114(4):762-770. PMID: 35987453.)





## Colorectal oligometastases

- Liver directed therapies have a well-established role in the management of oligometastatic disease in colorectal cancer
- ~ 20-25% of patients who undergo resection of liver metastases are long term survivors
- Typically managed in multidisciplinary fashion



**UW** Medicine

## Recent SBRT Data for colorectal disease

- 1. Prospective phase 2 trial of 61 patients with 76 liver metastases treated with SBRT. 1 yr LC 94%, 5 yr LC 78%. 1 yr OS 85%, 5 yr 18% with median OS 27.6 mos. *(Scorsetti et al. Phase II trial on SBRT for unresectable liver metastases: long-term outcome and prognostic factors of survival after 5 years of follow-up. Radiation Oncology volume 13, Article number: 234, 2018).*
- International multicenter registry study of 515 patients treated with SBRT with 1 yr LC 87% and OS 84%. (Mendez Romero et al. The Dutch–Belgian Registry of Stereotactic Body Radiation Therapy for Liver Metastases: Clinical Outcomes of 515 Patients and 668 Metastases. Int J Radiat Oncol Biol Phys 2021, 109, 5, 1377-86.)
- 3. Retrospective review of 99 liver metastases treated 2007-2015. Median OS 53 mos, 2yr OS 81.4%, 1 yr LC 87% and 2 yr LC 72%. (*Py et al. Long-term outcome of Stereotactic Body Radiation Therapy for patient with unresectable liver metastases from colorectal cancer. Cancer Radiother. 2021 Jun;25(4):350-357.*)





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First name:		Middle initial:
Edward		
Last name:		
Kim		
Phone number:	Email:	
	_	
2	Financial interests	

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3

Category (A-G)	Source of income and date	Amount	Recipient	
В	Employed by the University of Washington (ongoi		✓ Self	Family
			Self	Family
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			Self	Family

Other interests

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Have you authored, coauthored, or publicly provided an opinion, editorial, or publication related to any meeting topic? Topics(s):

My publications re: SBRT: ACR Appropriateness Criteria for non-spine bone metastases. J Palliat Med 2015. ACR Appropriateness Criteria for Metastatic Epidural Spinal Cord Compression and Recurrent Spinal metastasis J Palliat Med 2015, SBRT for HCC, Tech in Cancer Res Treat 2018, etc

Are you involved in formulating policy positions or clinical guidelines related to any meeting topic? Topics(s):

I am a member of the NCCN soft tissue sarcoma clinical practice guidelines panel

Could a coverage determination based on a Committee decision conflict with policies you have promoted or are obliged to follow? Topic(s):

No

Signature

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Signature

Date

Submit

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4/24/23

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#### Health Technology Clinical Committee Health Care Authority Conflict of Interest Disclosure

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1	Applicant information	
First name:		Middle initial:
John		
Last name:		
Kang		
Phone number:	Email:	
	_	

2

#### Financial interests

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Category (A-G)	Source of income and date	Amount	Recipient
В	Change Healthcare 2021-ongoing	350,000	Self 🖌 Family
			Self Family
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			Self Family
			Self Family

3

#### Other interests

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No

Are you involved in formulating policy positions or clinical guidelines related to any meeting topic? **Topics(s):** 

No

Could a coverage determination based on a Committee decision conflict with policies you have promoted or are obliged to follow? Topic(s):

No 4 Signature

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Signature Date Submit

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4/11/23

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1	Applicant information	
First name: DHULD		Middle initial: $\mathbf{A}$
Last name: Kartorow	172	
Phone number:	Email:	t and the second se
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HCA 13-0086 (10/21)

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Signature

Date

4624/23

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# Stereotactic Radiosurgery and Stereotactic Body Radiation Therapy: Final Evidence Report

Washington Health Technology Clinical Committee Meeting May 19, 2023 Presented by Beth Shaw, MSc, and Valerie King, MD, MPH



### Technology

- Radiation therapy (RT) uses high-energy X-ray or other particles to destroy cancer cells
  - Usually consists of a specific number of treatments over a set time period
  - Can be used in combination with other treatments, such as chemotherapy or surgery
- Most common type of RT is external-beam RT (EBRT)
  - 3-dimensional conformal radiation therapy (3D-CRT)
  - Intensity modulated radiation therapy (IMRT)
  - Proton beam therapy (PBT)
  - Image-guided radiation therapy (IGRT)
  - Stereotactic body radiation therapy (SBRT)

### Technology

- Stereotactic body radiation therapy (SBRT) or stereotactic ablative radiotherapy (SABR)
  - Typically delivered in 1 to 5 fractions
  - Can be primary treatment for early-stage cancers, treatment for discrete tumors in oligometastatic disease, for selected benign neoplasms in or near the central nervous system (CNS), or in recurrent cancer within previously irradiated regions
  - Other radiation-based options include implanted internal radiation therapy (or brachytherapy), intraoperative radiation therapy (IORT), systemic radiation therapy, radioimmunotherapy, and radiosensitizers or radioprotectors

### **Clinical Need**

- In 2019, 1,752,735 new invasive cancer cases were reported in the US
  - 863,830 in females and 888,905 in males
  - Incidence rate was 439 per 100,000
- While cancer affects people of all ages, races, ethnicities, and sexes, it does not affect all groups equally
  - Genetics, lifestyle, environmental exposures, and other factors can lead to differences in risk
  - For most cancers, age is the most important risk factor, with around 58% of cancers occurring in adults aged 65 years or older

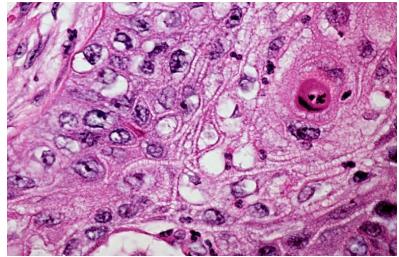


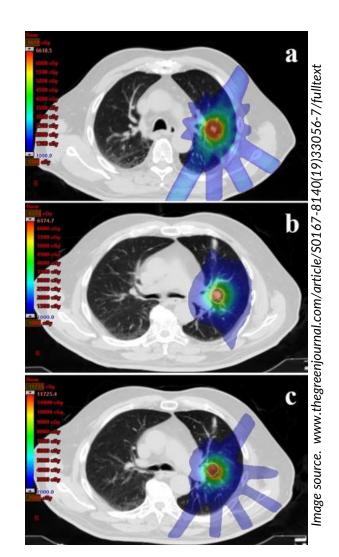
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### History

- In 2012, HTA on the effectiveness of stereotactic radiation surgery (SRS) and SBRT for treating various cancers was commissioned
- In 2013, coverage determination adopted
  - SRS for CNS primary and metastatic tumors is a covered benefit for adults and children when the following criteria are met:
    - Patient functional status score (i.e., Karnofsky score) is greater than or equal to 50; and
    - Evaluation includes multidisciplinary team analysis (e.g., tumor board), including surgical input
  - SBRT is covered for adults and children for the following conditions when the following criteria are met:
    - For cancers of spine/paraspinal structures; or
    - For inoperable non-small cell lung cancer (NSCLC), stage 1; and
    - Evaluation includes multidisciplinary team analysis, including surgical input
  - All other indications are noncovered

### History

- Since 2012, 3 signal search reports were conducted
- In 2022, SBRT was selected for rereview based on newly available published evidence that could change the original coverage decision
- Aim of the 2023 update
  - Conduct updated evidence searches
  - Produce an updated evidence review



### **Abbreviations**

- **ADT** androgen deprivation therapy
- **aHR** adjusted hazard ratio
- Cl confidence interval
- **cRT** conventional radiation therapy
- **CT** chemotherapy
- **EBRT** external beam radiation therapy
- **ENRT** elective nodal radiation therapy
- **FDA** U.S. Food and Drug Administration
- **GI** gastrointestinal
- **GRADE** Grading of Recommendations, Assessment, Development, and Evaluation

- **GU** genitourinary
- Gy Gray

٠

- **HFRT** hypofractionated radiotherapy
- **HIFU** high-intensity focused ultrasound
- **HR** hazard ratio
- ICER incremental cost-effectiveness ratio
  - **IMRT** intensity-modulated RT
    - IV intravenous
  - KQ key question

### **Abbreviations**

- LCNEC large-cell neuroendocrine carcinoma of the lung
- **MFRT** multifraction radiation therapy
- NCT US National Clinical Trial
- **NSCLC** non-small cell lung cancer
- NR not reported
- NRS nonrandomized study
- OR odds ratio
- **PFS** progression-free survival
- **QALY** quality-adjusted life year
- **RCT** randomized controlled trial

- **RFA** radiofrequency ablation
- **RT** radiation therapy
- **SABR** stereotactic ablative radiotherapy
- **SBRT** stereotactic body radiation therapy
- **TA** thermal ablation
- **TACE** transarterial chemoembolization
- **UHRT** ultrahypofractionated radiation therapy

### **Key Questions**

- Effectiveness in cancers not currently covered
- Harms in cancers not currently covered
- Differential efficacy or harms
  - Patient characteristics
  - Cancer type, site, grade, stage
  - Setting or provider characteristic
- Costs and cost-effectiveness



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# **Methods**

- Rescreened studies identified from the previous report and signal searches against criteria for this update
- Checked studies submitted during public comment on the PICOS and key questions
- Ran database searches in OVID Medline and Cochrane
- Searched websites and databases for clinical guidelines
- Searched FDA databases and ClinicalTrials.gov for harms and ongoing studies
- Assessed risk-of-bias (RoB) using standardized checklists
- Assessed certainty of evidence (CoE) using GRADE approach



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# **Risk-of-Bias Assessment**

#### • Low

Clear reporting of methods and mitigation of potential biases

#### Moderate

Incomplete information about methods that might mask important limitations

# High

Clear flaws that might introduce serious bias

# **GRADE Certainty of Evidence**

Outcomes rated: overall survival, progression, disease control, quality of life, and toxicity

#### • **High** (RCTs start here)

Very confident the estimate of effect of intervention on outcome lies close to the true effect

#### Moderate

Moderately confident in estimate of effect of intervention on outcome; true effect is likely close to estimate, but possibly different

#### Low (Nonrandomized studies start here)

Little confidence in estimate of effect of intervention on outcome; true effect may be substantially different from estimate

#### • Very Low

No confidence in estimate of effect of intervention on outcome; true effect is likely substantially different from estimate

#### **Outcomes**

Outcome Measure	Advantages	Disadvantages
Overall survival (OS)	<ul> <li>Easily and precisely measured</li> <li>Generally based on objective and quantitative assessment</li> </ul>	<ul> <li>May be affected by switch-over of control to treatment or subsequent therapies</li> <li>Needs longer follow-up</li> <li>Includes noncancer deaths</li> </ul>
Disease-free survival, event- free survival	<ul> <li>Generally assessed earlier and with smaller sample size than survival studies</li> <li>Generally based on objective and quantitative assessment</li> </ul>	<ul> <li>Potentially subject to assessment bias, particularly in open-label studies</li> <li>Definitions vary among studies</li> <li>Balanced timing of assessments among treatment arms is critical</li> <li>Includes noncancer deaths</li> </ul>

# **Outcomes (cont.)**

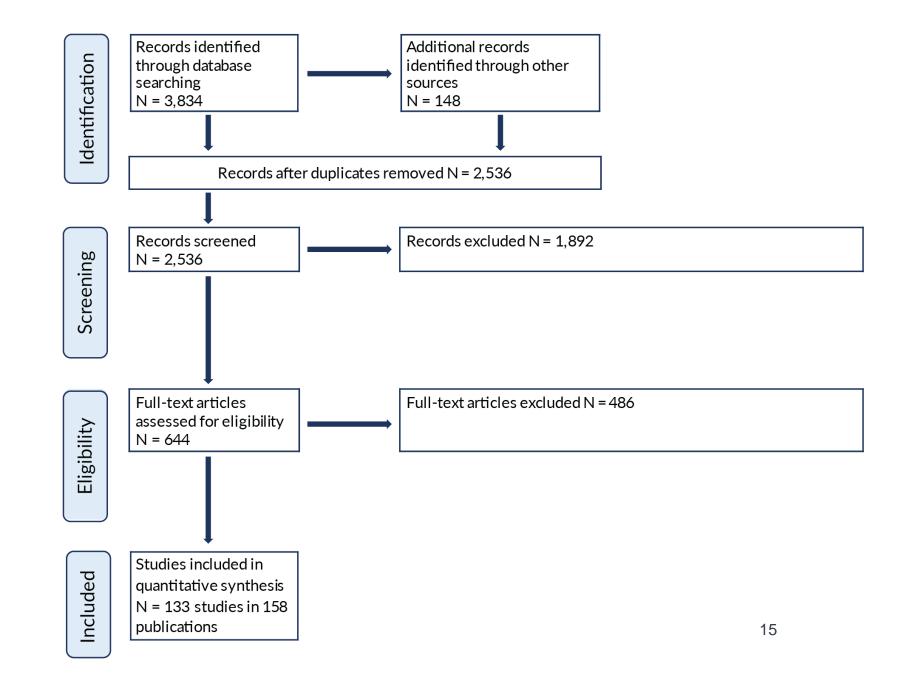
Outcome Measure	Advantages	Disadvantages
Progression- free survival (PFS), time to progression	<ul> <li>Generally assessed earlier and with smaller sample size than survival studies</li> <li>Measurement of stable disease included</li> <li>Generally based on objective and quantitative assessment</li> </ul>	<ul> <li>Potentially subject to assessment bias, particularly in open-label studies</li> <li>Definitions vary among studies</li> <li>Frequent radiological or other assessments</li> <li>Balanced timing of assessments among treatment arms is critical</li> <li>May not always correlate with survival</li> </ul>
Objective response rate (ORR) Complete response rate (CRR)	<ul> <li>Generally assessed earlier and with smaller sample size than survival studies</li> <li>Effect on tumor attributable to drug(s) or other treatment, not natural history</li> <li>Generally based on objective and quantitative assessment</li> </ul>	<ul> <li>Definitions vary among studies</li> <li>Frequent radiological or other assessments</li> <li>May not always correlate with survival</li> </ul>

# Findings

#### **Updated Evidence Review**



#### **Study Flow**



# Findings

#### Key Questions 1 to 3: Effectiveness

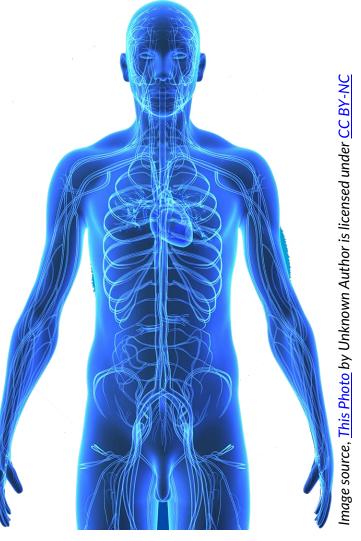


# **Findings by Cancer Site**

- Breast
- Prostate
- Lung
- Colorectal
- Uterine
- Melanoma
- Renal
- Pancreatic

Note: Bold indicates cancer sites for which we did not identify any eligible studies for this update.

- Head and neck
- Ovarian
- Liver
- Cervical
- Esophageal
- Oligometastatic
- Other



#### • 2012 report

 In the 2012 report, overall strength of evidence was assessed as very low for harms based on 4 case series

No comparative studies on the effectiveness of SBRT

### • 2023 update

- 4 RCTs
- 14 comparative nonrandomized studies (NRSs)
- **18** noncomparative studies (harms only)

Citation Setting NCT or Other Trial ID or Study Name	Duration Risk-of-bias	Patient Characteristics	Intervention	Comparator(s)
Randomized controll	1			
Brand et al, 2019	Followed up to	Total N = 874 men with low- to	• SBRT	<ul> <li>cRT or moderately</li> </ul>
37 centers in the	24 months	intermediate-risk localized prostate	$\circ$ 36.25 Gy in 5 fractions over 1	hypofractionated radiotherapy
UK, Ireland, and	Moderate risk-	cancer, comprising 433 in the SBRT	to 2 weeks (i.e., daily or	• PTV dose was 78 Gy in 39
Canada	of-bias	group and 441 in the control group	alternate days, at center	daily fractions or, following an
NCT01584258			discretion), with an additional secondary CTV dose target of	approved protocol amendment, 62 Gy in 20 daily
PACE-B			40 Gy	fractions
Kwan et al., 2022	At least 6	Total N = 80 men with	• SBRT	Moderate hypofractionation RT
2 sites in Canada	months	intermediate- to high-risk localized	• 36.25 Gy in 5 fractions weekly	<ul> <li>70 Gy in 28 fractions 5 times a</li> </ul>
NCT02594072	Moderate risk-	prostate cancer, comprising 42 in the SBRT group and 36 in the	<ul> <li>ADT (6 months in intermediate risk and 18 months in high-</li> </ul>	<ul> <li>week</li> <li>ADT (6 months in intermediate)</li> </ul>
ASSERT	of-bias	control group	risk) by either luteinizing	risk and 18 months in high-
			hormone-releasing hormone	risk) by either luteinizing
			agonists or antagonists	hormone-releasing hormone
				agonists or antagonists

Citation Setting NCT or Other Trial ID or Study Name	Duration Risk-of-bias	Patient Characteristics	Intervention	Comparator(s)
Randomized controlle	ed trials			
Lukka et al., 2018	Median follow-	Total N = 255 men with localized	• SBRT	• UHRT
37 sites, including	up of 3.8 years	T1 to T2 stage prostate cancer,	• 36.25 Gy in 5 fractions of	<ul> <li>51.6 Gy in 12 fractions of</li> </ul>
academic centers, in		comprising 127 in the SBRT in the group and 128 in the UHRT group	7.25 Gy • More than 2 weeks	4.3 Gy ₀ More than 2.5 weeks
the US and Canada	of-bias			
NCT01434290				
Widmark et al., 2019		Total N = 1,200 men with	• SBRT	• cRT
12 centers in	10 years	intermediate-to-high-risk localized	• 42.7 Gy in 7 fractions	• 78.0 Gy in 39 fractions
Sweden and	Moderate risk-	prostate cancer, comprising 598 in	<ul> <li>3 days over 2.4 weeks</li> </ul>	<ul> <li>5 days per week for 8 weeks</li> </ul>
Denmark	of-bias	the SBRT group and 602 in the cRT		
ISRCTN45905321		group		
HYPO-RT-PC				

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
	nediate-to-high-risk localized prostate cancer		
Overall survival			
N = 1,200 1 RCT	<ul> <li>In intermediate-to-high-risk localized prostate cancer:</li> <li>5-year overall survival: HR, 1.11; 95% CI, 0.73 to 1.69</li> </ul>	⊕⊕⊖⊖ Low	Downgraded 2 levels for imprecision (i.e., very wide CIs)
Progression-free surviv	val	•	
N = 1,200 1 RCT	<ul> <li>In intermediate-to-high-risk localized prostate cancer:</li> <li>5-year failure-free survival (biochemical or clinical failure: aHR, 1.00 (95% CI, 0.76 to 1.33)</li> </ul>	⊕⊕⊕⊖ MODERATE	Downgraded 1 level for imprecision (i.e., wide Cls)
Disease-control			
N = 1,200 1 RCT	<ul> <li>In intermediate-to-high-risk localized prostate cancer:</li> <li>Local failure: HR, 0.94; 95% Cl, 0.40 to 2.22</li> <li>Distant failure: HR, 0.99; 95% Cl, 0.63 to 1.54</li> <li>Use of ADT at 5 years: HR, 1.12; 95% Cl, 0.79 to 1.59</li> </ul>	⊕OOO VERY LOW	Downgraded 1 level for risk of bias and 2 levels for imprecision (i.e., very wide Cls)

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
	of RT for localized prostate cancer (all risk groups)		
Overall survival			
N = 75,749 5 comparative NRSs	Men with localized prostate cancer (all risk groups) treated with SBRT had similar or improved overall survival when compared with other treatment options, including cRT, IMRT and brachytherapy; studies reported at different times using different statistics, precluding any summary statistics (see detailed findings below).	⊕⊕OO low	Not downgraded
Disease-control		-	
N = 1,190 4 comparative NRSs	Men with localized prostate cancer (all risk groups) treated with SBRT had similar or improved disease control when compared with other treatment options, including cRT, IMRT and brachytherapy, with biochemical control rates of around 89% to 100% at 5 years.	⊕⊕⊖O low	Not downgraded

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale			
SBRT vs. other forms o	SBRT vs. other forms of RT for localized prostate cancer (all risk groups)					
Quality of life						
N = 2,154	Men with localized prostate cancer (all risk groups) treated	$\oplus \oplus \bigcirc \bigcirc$	Downgraded 1 level each for risk-			
3 RCTs	with SBRT had a similar quality of life to men treated with	LOW	of-bias and imprecision (i.e., not			
	other forms of RT; however, specific symptoms affecting		assessable)			
	quality of life may vary between treatments.					

- Based on the studies included in this review, we conclude that SBRT:
  - May be similarly or more effective than other options for individuals with localized prostate cancer (very low to moderate certainty of evidence [CoE], based on 4 RCTs and 14 comparative NRSs)

- Eligible guidelines made primarily conditional recommendations on the use of SBRT as an option for treating prostate cancer
  - 2022 American Society for Radiation Oncology and American Urological Association (ASTRO/AUA)
    - Strong recommendations on the use of SBRT for low- or intermediate-risk localized prostate cancer
    - Wording suggests a conditional approach with SBRT being offered as an option
  - 2022 Prostate Cancer Guidelines Panel (European Joint guidelines)
    - SBRT solely as a subject for future investigation
  - 2018 Australian and New Zealand Faculty of Radiation Oncology Genito-Urinary Group (FROGG)
    - SBRT an option for metastatic disease or as salvage therapy with inconsistent, low-quality evidence
    - Highlights the need for more clinical trials

- In 2013, the following coverage determination was adopted:
  - SBRT is covered for adults and children for the following conditions when these criteria are met:
    - For inoperable NSCLC, stage 1; and
    - Evaluation includes multidisciplinary team analysis, including surgical input
- 2012 report
  - Included 20 noncomparative NRSs in lung cancer, other than in inoperable NSCLC, stage 1 lung cancer
  - Populations in the studies tended to be mixed, and included both primary lung cancer and metastatic lung cancer
  - No coverage determination for other forms of lung cancer

- 2023 update
  - 2 RCTs
  - 11 comparative NRSs
  - 11 noncomparative studies (harms only)

Citation Setting NCT or Other Trial ID or Study Name	Duration Risk-of-bias	Patient Characteristics	Intervention	Comparator(s)
Randomized controlled tria	als			
Altorki et al., 2021	Followed up to 2	Total N = 60 people with	<ul> <li>SBRT plus durvalumab</li> </ul>	<ul> <li>Durvalumab</li> </ul>
Single center in the US	years	potentially resectable early-	<ul> <li>3 consecutive daily fractions of</li> </ul>	<ul> <li>2 cycles of</li> </ul>
NCT02904954		stage NSCLC (stages IA to IIIA), comprising 30 in durvalumab plus SBRT group and 30 in durvalumab group	<ul> <li>8 Gy</li> <li>2 cycles of durvalumab 3 weeks apart at a dose of 1.12 g by IV infusion over 60 min</li> </ul>	durvalumab 3 weeks apart at a dose of 1·12 g by IV infusion over 60 min
Theelen et al., 2019	Median follow-up	Total N = 78 people with	<ul> <li>SBRT plus pembrolizumab</li> </ul>	<ul> <li>Pembrolizumab</li> </ul>
3 centers in the Netherlands NCT02492568 PEMBRO-RT	of 24 months Moderate risk-of- bias	advanced NSCLC, comprising 38 in SBRT group and 40 in control group	<ul> <li>3 doses of 8 Gy delivered on alternate days to a single tumor site that did not overlap with biopsy site and was deemed most safe or convenient for patient</li> <li>Pembrolizumab administered IV at 200 mg every 3 weeks</li> </ul>	<ul> <li>Pembrolizumab administered IV at 200 mg every 3 weeks</li> </ul>

Number of Participants (N)	Findings	Certainty of Evidence	Rationale
Number of Studies			
	RT for operable early-stage NCSLC		
Overall survival		1	
5 comparative MK55	SBRT was associated with significantly worse outcomes than surgery for operable early-stage NCSLC; surgery was associated with around a 60 to 65% lower risk of mortality. However, 1 study did find that in patients who were medically operable, SBRT and lobectomy may be equally effective.	⊕OOO VERY LOW	Downgraded 1 level for inconsistency
Progression-free survival		-	
1 comparative NRS	In patients who were medically operable, SBRT and lobectomy may be equally effective (HR, 1.57; 95% CI, 0.68 to 3.64)	⊕OOO VERY LOW	Downgraded 1 level for risk- of-bias and 2 levels for imprecision (i.e., very wide Cls)
Disease-control			
1 RCT	In people with potentially resectable early-stage NCSLC, SBRT in combination with durvalumab was associated with significantly higher odds of having a major pathological response (OR, 16.0; 95% CI, 3.2 to 79.6) or a partial radiographic response (46.7% SBRT with durvalumab vs. 3.3% durvalumab; $P = .001$ ) than durvalumab alone.	⊕⊕⊕O MODERATE	Downgraded 1 level for risk- of-bias

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. RT for inoperable Overall survival	e stage II		
1 comparative NRS	$t_{\text{han}} = CDT (UD = 0.70, 0.50) CL = 0.71 t_{\text{han}} = 0.97) \text{ ar}$	⊕⊕OO low	Not downgraded

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale		
SBRT vs. no SBRT for adv	anced NCSLC				
Overall survival		-			
N = 78	People with advanced NSCLC treated with SBRT after	$\Theta \oplus \Theta O$	Downgraded 1 level for		
	pembrolizumab or pembrolizumab alone had a similar overall survival (median: 15.9 months SBRT vs. 7.6 months control; HR, 0.66; 95% CI, 0.37 to 1.18)	MODERATE	imprecision (i.e., wide CIs)		
	However, in subgroup analyses, men (HR, 0.42; 95%Cl, 0.19 to 0.96; <i>P</i> = .04) and smokers (HR, 0.48; 95% Cl, 0.25 to 0.93; <i>P</i> = .03) had significantly improved survival with SBRT compared with pembrolizumab alone.				
Progression-free survival					
	People with advanced NSCLC treated with SBRT after		Downgraded 1 level each for		
1 RCT	pembrolizumab or pembrolizumab alone had a similar PFS (HR, 0.71; 95% Cl, 0.42 to 1.18).	IOW	risk-of-bias and imprecision (i.e., wide Cls)		

Number of Participants (N)	Findings	Certainty of Evidence	Rationale				
Number of Studies							
	BRT vs. surgery or cRT for lung metastases						
Overall survival		r					
	In people with lung metastases, SBRT and surgery may be	$\oplus \oplus \bigcirc \bigcirc$	Not downgraded				
4 comparative NRSs	associated with similar overall survival (median survival at 2	LOW					
	years of around 68% to 77% in the SBRT group vs. 82% in the						
	surgery group); however, SBRT may be associated with						
	improved survival when compared with cRT (median survival						
	of 26 months in the SBRT group vs. 9 months in the cRT						
	group; P < .001).						
Progression-free survival		i					
	People with lung metastases treated with SBRT had	$\oplus 000$	Downgraded 1 level for				
3 comparative NRSs	significantly worse PFS than people treated with surgery	VERY LOW	inconsistency				
	(around 3 times more likely to have progression). However,						
	results were mixed with 1 study showing no difference						
	between SBRT and surgery.						
Disease-control							
	Results were mixed with SBRT being associated with both	$\oplus 000$	Downgraded 1 level for				
	similar and lower levels of local control than surgery for lung	VERY LOW	inconsistency				
	metastases. SBRT, however, was significantly associated with						
	improved local control when compared with cRT. Studies						
	reported at different times using different statistics,						
	precluding any summary statistics (see detailed findings						
	below).						

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale		
SBRT vs. surgery or cRT for	SBRT vs. surgery or cRT for LCNEC of the lung				
Overall survival					
N = 3,963	In people with LCNEC of the lung, SBRT may be associated	$\oplus \oplus \bigcirc \bigcirc$	Not downgraded		
	with improved survival when compared with cRT (HR, 0.83;	LOW			
	95% CI, 0.68 to 1.00) <sup>b</sup> , but worse outcomes when compared				
	with surgery (HR, 1.61; 95% Cl, 1.36 to 1.92).				

- Based on the studies included in this review, we conclude that SBRT:
  - May be similarly or more effective than radiation therapy for inoperable stage II non-small cell lung cancer (NSCLC; low CoE, based on 1 comparative NRS) or in combination with pembrolizumab compared with pembrolizumab alone for advanced NSCLC (low to moderate CoE, based on 1 RCT)
  - Also appears to be similarly or more effective than conventional radiation therapy (cRT) for people with lung metastases (very low to low CoE, based on 4 comparative NRSs) or large cell neuroendocrine carcinoma (LCNEC) of the lung (low CoE, based on 2 comparative NRSs)
  - May be less effective than surgery for resectable lung cancer (very low to low CoE, based on 10 comparative NRSs)

- Overall, the guidelines indicate there is some evidence for the use of SBRT for lung cancer
  - Evidence quality is generally moderate-to-low
  - Usually recommended for patients who refuse or who are at high risk for surgery, lobectomy, or chemotherapy
  - For patients who are operable, SBRT is still considered a therapy under investigation in the current guidelines

- 2012 report
  - No eligible studies identified
- 2023 update
  - 1 comparative NRS
  - **1** noncomparative study (harms only)

Number of Participants (N) Number of Studies		Certainty of Evidence	Rationale		
SBRT vs. cRT in stage I RCC					
Overall survival					
N = 91,965 1 comparative NRS	<ul> <li>In people with stage I RCC, SBRT was associated with a significantly worse overall survival than people treated with ablation or surgery:</li> <li>Partial nephrectomy vs. SBRT: HR, 0.29 (95% CI, 0.19 to 0.46)</li> <li>Cryoablation vs. SBRT: HR, 0.40 (95% CI, 0.26 to 0.60)</li> <li>Radiofrequency ablation or microwave ablation vs. SBRT: HR, 0.46 (95% CI, 0.31 to 0.67)</li> </ul>	⊕⊕OO Low	Not downgraded		

- Based on the studies included in this review, we conclude that SBRT:
  - May be less effective than ablation (RFA, microwave, or cryoablation) or surgery for stage 1 renal cell carcinoma (low CoE, based on 1 comparative NRS)

- Overall, the guidelines make conditional recommendations on the use of SBRT for certain clinical situations
  - Particularly for metastatic disease or when patients are considered unsuitable for surgery
  - Based on low- to moderate-quality evidence
- However, guidelines also highlight the need for future clinical trials on the use of SBRT in renal cancer

- 2012 report
  - Overall strength of evidence assessed as very low for effectiveness and harms, based on 1 systematic review and 4 case series
- 2023 update
  - 3 comparative NRSs

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale			
SBRT vs. CT or IMRT for unresected pancreatic cancer						
Overall survival						
N = 14,331 1 comparative NRS	In people with unresected pancreatic cancer treated with SBRT had significantly better overall survival than people treated with CT (13.9 months SBRT vs. 10.2 months CT; $P < .001$ ) or IMRT (13.9 months SBRT vs. 12.2 months IMRT; $P = .049$ ). However, there was no difference in overall survival between SBRT with multi-agent CT and multi-agent CT alone (14.8 months SBRT with multi-agent CT vs. 12.9 months multi-agent CT alone; $P = .09$ ).	⊕⊕OO LOW	Not downgraded			

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale		
SBRT vs. cRT for locally advanced pancreatic cancer					
Overall survival					
N = 8,450 1 comparative NRS	People with locally advanced pancreatic cancer treated with SBRT had significantly better overall survival than people treated with cRT at 2 years (HR, 0.84; 95% Cl, 0.75 to 0.93), with a significantly longer median survival.	⊕⊕⊖O Low	Not downgraded		

- Based on the studies included in this review, we conclude that SBRT:
  - May be more effective than chemotherapy or intensitymodulated radiation therapy for unresected pancreatic cancer (low CoE, based on 1 comparative NRS)
  - May be more effective than conventional RT for pancreatic cancer (low CoE, based on 3 comparative NRSs)

- 2019 ASTRO
  - Conditional recommendations on the use of SBRT as an option for treating pancreatic cancer
  - Based on very low- to low-quality evidence
  - Following surgical resection, SBRT should only be used in the context of research

- 2012 report
  - Overall strength of evidence assessed as very low for harms for head and neck cancers (specifically, ocular and glomus jugulare), based on 1 systematic review and 7 case series
  - No comparative effectiveness studies were identified
- 2023 update
  - 1 RCT
  - 4 comparative NRSs

Citation Setting NCT or Other Trial ID	Risk-of-bias	Patient Characteristics	Description of Intervention	Description of Comparator(s)
Randomized controlled trials				
McBride et al., 2021	RCT	Total N = 62 people with	SBRT in combination	<ul> <li>Nivolumab</li> </ul>
Single center in the US	Median follow-up of	metastatic or recurrent head and neck squamous cell carcinoma,	with nivolumab 。 9 Gy in 3 fractions	
NCT02684253	20 months	comprising 32 in SBRT group and	delivered every other	
	Moderate risk-of-bias	30 in control group	day	

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale	
SBRT vs. brachytherapy	/ in early-stage oropharyngeal cancer			
Overall survival		-		
N = 250 1 comparative NRS	SBRT boost or brachytherapy boost after cRT were associated with a similar overall survival at 3 years (81% SBRT vs. 83% BT; P = .83).	⊕⊕⊖O low	Not downgraded	
<b>Progression-free surviv</b>	al			
N = 250 1 comparative NRS	SBRT boost or brachytherapy boost after cRT were associated with a similar disease-free survival at 3 years (92% SBRT vs. 86% BT; $P = .15$ ).	⊕⊕⊖⊖ Low	Not downgraded	
Disease-control				
N = 250 1 comparative NRS	SBRT boost or brachytherapy boost after cRT were associated with a similar local control rate at 3 years (97% SBRT vs. 94% BT; P = .33).	⊕⊕⊖⊖ low	Not downgraded	
Quality of life				
N = 250 1 comparative NRS	No significant difference in quality of life in patients with early- stage oropharyngeal cancer boosted with SBRT or brachytherapy after cRT.	⊕⊕○○ Low	Downgraded 1 level for imprecision (i.e., not assessable)	

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. other treatmer	nt options for recurrent or metastatic head and neck cancer		
Overall survival			
1 RCT	in the second seco	MODERATE	Downgraded 1 level for imprecision (i.e., not assessable)
3 comparative NRSs	SBRT appears to be associated with a significantly worse overall survival than charged particle RT, (HR, 0.35; 95% CI, 0.13 to 0.94), but a similar cancer-specific survival to IMRT (HR, 0.88; 95% CI, 0.70 to 1.10) and conformal RT (at 2 years, 64% SBRT vs. 47% conformal RT; <i>P</i> = .40).		Not downgraded

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. other treatmer	nt options for recurrent or metastatic head and neck cancer		
Progression-free surviva	al		
N = 62	No difference between nivolumab in combination with SBRT or	$\Theta \oplus \Theta \bigcirc$	Downgraded 1
1 RCT	nivolumab alone (at 12 months, 16.8% SBRT vs. 32.2% control; P = .79)	MODERATE	level for imprecision (i.e., not assessable)
Disease-control			
N = 62	No difference between nivolumab in combination with SBRT or	$\oplus \oplus \bigcirc \bigcirc$	Downgraded 2
1 RCT	nivolumab alone (at 12 months, OR, 0.80; 95% Cl, 0.24 to 2.61)		levels for imprecision (i.e., very wide Cls)ª
N = 641	SBRT appears to be associated with similar levels of disease	$\oplus \oplus \bigcirc \bigcirc$	Not downgraded
3 comparative NRSs	control to IMRT (HR, 1.15; 95% CI, 0.89 to 1.50), conformal RT (at 2 years, 82% SBRT; 80% conformal RT; <i>P</i> = .57), and charged particle RT (at 1 year, 67% SBRT vs. 67% charged particle RT; <i>P</i> value not reported)	LOW	

- Based on the studies included in this review, we conclude that SBRT:
  - May be similarly effective to brachytherapy, when used as a boost treatment after cRT for early-stage oropharyngeal cancer (low CoE, based on 1 comparative NRS)
  - May be less effective than charged particle RT for recurrent or metastatic head and neck cancer, but similar in effectiveness to intensity-modulated RT (IMRT) and conformal RT (low to moderate CoE, based on 1 RCT and 3 comparative NRSs)

• No eligible guidelines identified

- 2012 report
  - Evidence assessed as being of very low certainty, with any conclusions about benefit and harms being uncertain, based on 2 systematic reviews of case series and 7 case series
- 2023 update
  - 20 comparative NRSs
  - **11** noncomparative studies (harms only)

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. RFA for early-stag	e HCC	•	
Overall survival			
N = 4,892	In people with early-stage HCC, results were	$\oplus 000$	Downgraded 1 level for
4 comparative NRSs	mixed. SBRT may be associated with similar overall survival to RFA (at 5 years, 78.4% vs. 46.3%; P = .09 over the 5 years); however, 1 study showed that SBRT may be associated with worse survival than RFA at 5 years (HR, 0.67; 95% CI, 0.55 to 0.81).	VERY LOW	inconsistency (i.e., mixed results)
Progression-free survival	•	•	
N = 98	In people with early-stage HCC, SBRT after RFA	$\oplus 000$	Downgraded 1 level for
1 comparative NRS	may be associated with similar PFS to repeated RFA (at 2 years, $31.4\%$ vs. $28.6\%$ ; $P = .31$ ).	VERY LOW	imprecision (i.e., not assessable)
Disease-control			
N = 472	In people with early-stage HCC, SBRT may be	$\oplus \oplus \bigcirc \bigcirc$	Not downgraded
2 comparative NRSs	associated with similar rates of intrahepatic recurrence (at 3 years, 59.3% RT vs. 57.6% RFA; P = .64) and local recurrence (0 SBRT vs. 25.7% RFA; $P = .06$ ).	LOW	

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. TACE and RFA in s	small HCCs		
Overall survival	In people with small LICCs SPPT along or in	<b>#</b>	Downgradad 1 lovel for
	In people with small HCCs, SBRT, alone or in combination with TACE is associated with a similar overall survival to TACE alone, TACE in combination with TACE, or to RFA. Studies reported at different times using different statistics, precluding any summary statistics (see detailed findings below).	⊕OOO VERY LOW	Downgraded 1 level for imprecision (i.e., wide CIs)
Progression-free survival			
N = 615 3 comparative NRSs	In people with small HCCs, SBRT is associated with a similar PFS to RFA. SBRT in combination with TACE is associated with similar or improved PFS to TACE alone or SBRT alone. Studies reported at different times using different statistics, precluding any summary statistics (see detailed findings below).	⊕⊕OO Low	Not downgraded

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. TACE and RFA in s	mall HCCs		
Disease-control			
N = 683 4 comparative NRSs	In people with small HCC, SBRT added to TACE appears to be associated with improved local control, but results are mixed. Studies reported at different times using different statistics, precluding any summary statistics (see detailed findings below).		Downgraded 1 level for inconsistency (i.e., mixed results)

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. other treatments f	or unresectable HCC		
Overall survival			
	a such in a line with TACE and a such a large state of	⊕⊕OO Low	Not downgraded
Progression-free survival		1	
N = 889 5 comparative NRSs	In people with unresectable HCC, SBRT, alone or in combination with TACE, appears to be associated with similar or improved PFS compared with TACE alone, RFA, or SIRT. When compared with TA, SBRT appears to be associated with a lower PFS. Studies reported at different times using different statistics, precluding any summary statistics (see detailed findings below).	⊕⊕⊖⊖ low	Not downgraded

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. other treatments f	for unresectable HCC		
Disease-control			
N = 3,149	In people with unresectable HCC, SBRT, alone or	$\oplus \oplus \bigcirc \bigcirc$	Not downgraded
8 comparative NRSs	in combination with TACE, may have similar or improved rates of disease control and recurrence when compared with RFA or TACE alone. When compared with TA, results are mixed, with 1 study showing no difference and 1 showing a significant decrease in local control with SBRT. Studies reported at different times using different statistics, precluding any summary statistics (see detailed findings below).	LOW	

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. sorafenib for adva	nced HCC		
Overall survival		-	
1 comparative NRS	In people with advanced HCC, SBRT was associated with improved survival when compared with sorafenib (HR, 0.53; 95% CI, 0.36 to 0.77)	⊕⊕OO low	Not downgraded <sup>a</sup>
Progression-free survival		-	
1 comparative NRS	In people with advanced HCC, SBRT was associated with improved PFS when compared with sorafenib (HR, 0.59; 95% CI, 0.42 to 0.86)		Downgraded 1 level imprecision (i.e., wide CIs) <sup>a</sup>

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
	s bridging therapy for people on waiting list for liver	transplantation	due to HCC
Overall survival			
2 comparative NRSs	SBRT, as bridge therapy, appears to be associated with a similar overall survival to other options for bridge therapy (TACE, RFA, or HIFU; at around 61% to 73% at 3 years).	⊕⊕⊖O low	Not downgraded
Progression-free survival		•	
1 comparative NRS	SBRT, as bridge therapy, appears to be associated with improved PFS when compared with TACE or HIFU (progression at 3 years, 18.5% SBRT vs. 54.9% TACE vs. 62.8% HIFU; <i>P</i> < .001).	VERY LOW	Downgraded 1 level for imprecision (i.e., not assessable) <sup>a</sup>
Disease-control	· · · ·		
2 comparative NRSs	SBRT, as bridge therapy, appears to be associated with a better disease control than other options for bridge therapy (TACE or HIFU) but may be associated with worse disease control than RFA. Studies reported at different times using different statistics, precluding any summary statistics (see detailed findings below).		Downgraded 1 level for imprecision (i.e., not assessable) <sup>a</sup>

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale		
SBRT vs. TARE or cRT for unresectable intrahepatic cholangiocarcinoma Overall survival					
1 comparative NRS	SBRT was associated with improved survival compared with TARE (HR, 0.40; 95% CI, 0.22 to 0.74) or cRT (HR, 0.37; 95% CI, 0.20 to 0.68)	⊕⊕OO low	Not downgraded		

- Based on the studies included in this review, we conclude that SBRT:
  - May be as effective as radiofrequency ablation (RFA) for early-stage liver cancer; however, results were mixed (very low to low CoE, based on 4 comparative NRSs)
  - Alone, or in combination with transarterial chemoembolization (TACE), may be as effective as RFA or TACE alone for small liver cancers (very low to low CoE, based on 4 comparative NRSs) and for unresectable liver cancer (low CoE, based on 8 comparative NRSs)
  - May be more effective than sorafenib for advanced liver cancer (very low to low CoE, based on 1 comparative NRS)

- Based on the studies included in this review, we conclude that SBRT:
  - May be similarly or more effective than other options (RFA, TACE, high-intensity focused ultrasound [HIFU]) when used as a bridging therapy for people on the waiting list for liver transplantation due to liver cancer (very low to low CoE, based on 2 comparative NRSs)
  - May be more effective than sorafenib for advanced liver cancer (very low to low CoE, based on 1 comparative NRS)
  - May be more effective than transarterial radioembolization (TARE) for unresectable intrahepatic cholangiocarcinoma (low CoE, based on 1 comparative NRS)

- For biliary tract cancer
  - SBRT is conditionally recommended in specific situations
  - Based on low to moderate evidence quality
- For hepatocellular carcinoma
  - SBRT is an alternative for treatment of local failure in certain tumors
  - Based on low to moderate evidence quality
- 2022 American College of Radiology (ACR) appropriateness criteria
  - SBRT may be appropriate for a range of specific hepatocellular cancer types
  - Strength of evidence is not reported

- 2012 report
  - Included 2 noncomparative studies
  - No specific coverage determinations were made
- 2023 update
  - 3 RCTs
  - 3 comparative NRSs
  - **12** noncomparative studies (harms only)

NCT or Other Trial ID or Study Name	Duration Risk-of-bias	Patient Characteristics	Intervention	Comparator(s)
6 centers, including academic centers, in		Total N = 62 men with recurrent oligometastatic prostate cancer, comprising 31 in treatment arm (majority received SBRT; remainder underwent surgery) and 31 in active surveillance arm	<ul> <li>SBRT         <ul> <li>Total dose of 30 Gy (80% of maximal dose) delivered in 3 fractions</li> <li>25 (81%)</li> </ul> </li> <li>Metastectomy         <ul> <li>6 (19%)</li> </ul> </li> </ul>	• Surveillance
Palma et al., 2019 10 hospitals in Canada, the Netherlands, Scotland, and Australia NCT01446744 SABR-COMET	Followed up to 10 years Moderate risk-of- bias	Total N = 99 people with a controlled primary tumor and 1 to 5 oligometastatic lesions, comprising 66 in SBRT group and 33 in control group Primary sites were mostly adrenal, bone, liver, and lung	<ul> <li>SBRT         <ul> <li>Doses ranged from 30 to 60 Gy in 3 to 8 fractions, depending on target size and location</li> <li>Single fractions of 16 to 24 Gy permitted for targets in brain and vertebrae</li> <li>Concurrent chemotherapy or targeted therapy was not permitted within 4 weeks before SBRT</li> </ul> </li> <li>Standard of care, tailored to individual clinical circumstance</li> </ul>	<ul> <li>Standard of care, tailored to individual clinical circumstance</li> <li>Radiotherapy delivered according to standard principles of palliative radiation, with goal of alleviating symptoms or preventing anticipated complications of progression</li> </ul>

Citation Setting NCT or Other Trial ID or Study Name Randomized controlled	Risk-of-bias	Patient Characteristics	Intervention	Comparator(s)
Phillips et al., 2020 3 academic centers in	Followed up to 24 months Moderate risk-of-	Total N = 54 men with oligometastatic prostate cancer, comprising 36 in group and 18 in observation group	<ul> <li>SBRT         <ul> <li>Dose and fractionation based on size and location of each lesion, with prescription doses ranging from 19.5 to 48.0 Gy in 3 to 5 fractions</li> <li>Salvage RT was allowed</li> <li>Patients were allowed to have received ADT or other systemic therapy during initial management or salvage treatment but not within 6 months of enrollment</li> </ul> </li> </ul>	<ul> <li>Observation         <ul> <li>Salvage RT was allowed</li> <li>Patients were allowed to have received ADT or other systemic therapy during initial management or salvage treatment but not within 6 months of enrollment</li> </ul> </li> </ul>

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale		
	d of care for oligometastatic cancer (primaries mostly adrenal, bone, liv	ver, and lung)			
Overall survival					
N = 99 1 RCT	1.10)	⊕⊕⊕⊖ MODERATE	Downgraded 1 level for imprecision (i.e., wide Cls)		
	At 6 years, improved survival with SBRT (HR, 0.47; 95% Cl, 0.27 to 0.81)				
<b>Progression-free</b>					
N = 99	At 5 years, improved PFS with SBRT (HR, 0.47; 95% CI, 0.30 to 0.76)		Downgraded 1 level for risk-of-		
1 RCT	At 6 years, improved PFS with SBRT (HR, 0.48; 95% CI, 0.31 to 0.76)	MODERATE	bias		
Disease-control	Disease-control				
N = 99	SBRT is associated with improved disease control (absence of	$\oplus \oplus \bigcirc \bigcirc$	Downgraded 1 level each for risk-		
1 RCT	progression, 75% SBRT vs. 49% standard of care; <i>P</i> = .001; lesional control by location).	LOW	of-bias and imprecision (i.e., not assessable)		
Quality of life					
N = 99	People with a controlled primary tumor and 1 to 5 oligometastatic	$\oplus \oplus \bigcirc \bigcirc$	Downgraded 1 level each for risk-		
1 RCT	lesions treated with SBRT or standard of care had a similar quality of life at each subsequent follow-up.	LOW	of-bias and imprecision (i.e., not assessable)		

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale	
SBRT vs. observa	tion for oligometastatic prostate cancer			
Overall survival				
N = 116 2 RCTs	In a pooled analysis of 2 RCTs, the median for overall survival was not reached in either group, with similar overall survival between groups (HR, 0.53; 95% CI, 0.13 to 2.11).	⊕⊕OO low	Downgraded 2 levels for imprecision (i.e., very wide CIs)	
<b>Progression-free</b>	survival	-		
N = 116 2 RCTs	SBRT may be associated with similar or improved PFS (11.9 months MDT vs. 5.9 months surveillance; HR, 0.44; 95% CI, 0.29 to 0.66), and other measures of disease-related survival (ADT-free and castration-resistant prostate cancer-free survival).	⊕⊕⊖⊖ low	Downgraded 1 level each for risk- of-bias and imprecision (i.e., wide Cls)	
Disease-control				
N = 54 1 RCT	Men treated with SBRT had higher complete response (28% SBRT vs. 8% observation) and partial response rates (43% vs. 39% observation) at 6 months; however, no formal statistical testing was reported.	⊕⊕⊖⊖ low	Downgraded 1 level each for risk- of-bias and imprecision (i.e., not assessable)	
Quality of life				
N = 116 2 RCTs	No difference between groups.	⊕⊕OO low	Downgraded 1 level each for risk- of-bias and imprecision (i.e., not assessable)	

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale	
	SBRT for oligometastatic prostate cancer			
Overall survival				
N = 506	Men with oligometastatic or oligorecurrent prostate cancer treated	$\oplus \oplus \bigcirc \bigcirc$	Not downgraded <sup>a</sup>	
	with SBRT or cRT had a similar overall survival at 2 years (87.7% SBRT vs. 87.3% cRT; <i>P</i> = .91)	LOW		
<b>Progression-free</b>				
N = 682	SBRT appears to be associated with a worse metastasis-free survival	$\oplus 000$	Downgraded 1 level for	
2 comparative	when compared with elective nodal RT but similar or improved PFS	VERY LOW	imprecision (i.e., not assessable)	
NRSs	when compared with cRT.			
Disease-control				
N = 239	SBRT appears to be associated with worse outcomes (local and	$\oplus \bigcirc \bigcirc \bigcirc \bigcirc$	Downgraded 1 level for	
2 comparative	lymph node progression, relapse) when compared with elective	VERY LOW	imprecision (i.e., not assessable)	
NRSs	nodal RT (68% SBRT vs. 77% with elective nodal RT; P = .01) but			
	similar or improved outcomes (time to ADT or castration-resistance)			
	when compared with no SBRT.			

- Based on the studies included in this review, we conclude that SBRT:
  - Appears to be more effective than standard of care or observation for oligometastatic cancer (low to moderate CoE, based on 3 RCTs)
  - For oligometastatic prostate cancer, elective nodal radiation therapy may be more effective than SBRT (very low to low CoE, based on 3 comparative NRSs)

• No eligible guidelines identified

### **Adrenal Cancer**

- 2012 report
  - Overall strength of evidence was assessed as very low for effectiveness and harms for adrenal metastases, based on 2 case series
- 2023 update
  - 1 noncomparative study (harms only)
- No eligible guidelines identified

#### • 2012 report

- No included studies other than for spine cancers (a covered indication)
- 2023 update
  - □ 1 RCT

Citation Setting NCT or Other Trial ID	Duration Risk of Bias	Patient Characteristics	Description of Intervention	Description of Comparator(s)
Nguyen et al., 2019	Followed up to 24 months	Total N = 160 people with	• SBRT	Standard MFRT
Single academic center in the US <b>NCT02163226</b>	Moderate risk of bias	radiologically confirmed painful bone metastases, comprising 81 in the SBRT group and in the 79 control group	<ul> <li>Single-fraction 12 Gy for lesions &gt; 4 cm or 16 Gy for lesions ≤4 cm</li> <li>Standard concurrent chemotherapy, immunotherapy, or targeted therapy was allowed</li> </ul>	<ul> <li>30 Gy delivered in 10</li> <li>3-Gy fractions</li> <li>Standard concurrent chemotherapy, immunotherapy, or targeted therapy was allowed</li> </ul>

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale		
SBRT vs. cRT for bone metastas		Evidence			
Overall survival	565				
N = $160$ 1 RCT	People with radiologically confirmed painful bone metastases (mostly nonspine) treated	⊕⊕⊕O MODERATE	Downgraded 1 level imprecision (i.e., not		
	with SBRT or MFRT had a similar overall survival (median, 6.7 months in both groups).		assessable)		
Disease-control		-			
N = 160	When compared with MFRT, SBRT was	$\oplus 000$	Downgraded 1 level for risk of		
1 RCT	found to be noninferior for both local failure (HR, 0.18; 95% Cl, 0.02 to 1.47).	VERY LOW	bias and 2 levels for imprecision (i.e., very wide Cls)		
Quality of life					
N = 160 1 RCT	No significant difference in quality of life for patients treated with SBRT or with MFRT.	⊕⊕⊖O Low	Downgraded 1 level each for risk of bias and imprecision (i.e., not assessable)		

- Based on the studies included in this review, we conclude that SBRT:
  - May be as effective as multifraction RT for painful bone metastases (moderate CoE, based on 1 RCT)

- For nonspine bone cancer, guidelines recommend
  - SBRT is considered as an option, particularly for localized or metastatic disease
  - Evidence quality, when provided, is low

# Subgroups

- Few studies reported on clinical subgroups of interest, but there was some indication specific populations (by cancer site) may be more likely to benefit from SBRT compared with other populations
  - Subgroups varied by cancer type and treatment site and were often only reported in single studies

# Findings

#### Key Questions 2 and 3: Toxicity



#### **Prostate Cancer**

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. other forms	of RT for localized prostate cancer (all risk groups)		
Toxicity			
N = 2,409 4 RCTs	Rates of toxicities of grade 3 or higher were relatively infrequent in SBRT for localized prostate cancer (around 1% to 2%), and were similar to those of other RTs.	⊕⊕⊕○ MODERATE	Downgraded 1 level for risk-of- bias
N = 67,968 5 comparative NRSs	Overall, grade 3 toxicities were rare (up to 6% depending on the specific toxicity and the time point) and no grade 4 or 5 events were reported when SBRT was used for localized prostate cancer (all risk groups). There may be some evidence SBRT is associated with increased urinary retention or obstruction, urinary fistula, and more GI and GU toxicity than IMRT, and greater GI toxicity than brachytherapy.	⊕⊕⊖O Low	Not downgraded

## Lung Cancer

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. surgery and othe Toxicity	er RT for any lung cancer		
N = 138 2 RCTs	Grade 3 and higher events occurred in around 3% to 11% of SBRT group; most common were dyspnea and pneumonia, pancreatitis, and fatigue.		Downgraded 1 level for risk- of-bias
N = 221 2 comparative NRSs	Grade 3 toxicities were not common with SBRT, and	⊕⊕○○ low	Not downgraded

### **Pancreatic Cancer**

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. CT or cRT for pancrea	tic cancer		
Toxicity			
N = 5,624 1 comparative NRS	In people with nonmetastatic, unresectable pancreatic cancer, SBRT was associated with significantly more GI bleeds than CT alone (HR, 4.13; 95% CI, 2.58 to 6.61) and GI strictures (HR, 1.58; 95% CI, 1.18 to 2.21). However, risk varied by age, with SBRT being associated with similar rates of GI complications to cRT in younger people.	⊕⊕⊖O Low	Not downgraded

### Head and Neck Cancers

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. other options f	or early and recurrent head and neck cancers		
Toxicity			
1 RCT	windows also also a long de 2 and bish an 0.70/ CDDT va 1.2.20/	MODERATE	Downgraded 1 level for imprecision (i.e., not assessable)
4 comparative NRSs	SBRT had a favorable toxicity profile, with similar or fewer toxicities than other treatment options (brachytherapy, conformal RT, IMRT, charged particle RT); however, grade 5 events were relatively high, with 1 study reporting 12.5% grade 5 events in the SBRT group	LOW	Not downgraded

## Liver Cancer

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. other treatments	for HCC		
Toxicity			
N = 6,071 16 comparative NRSs	Rates of toxicities of grade 3 or higher were relatively infrequent in SBRT, and were similar to those of other RTs or treatment options. SBRT may be associated with some increased toxicities, but it is also associated with some decreased toxicities when compared with other options. Rates of toxicities varied by type of toxicity and time frame.	⊕⊕○○ LOW	Not downgraded

### **Oligometastatic Cancers**

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. other tre	eatment for oligometastatic cancer		
(primary sites incl	uded prostate, breast, lung, and other sites)		
Toxicity			
N = 215 3 RCTs	No grade 3 and higher toxicities were seen in 2 of the 3 trials, but some SBRT-related deaths were observed.	⊕⊕⊖⊖ low	Downgraded 1 level each for risk- of-bias and imprecision (i.e., not assessable)
N = 745 3 comparative NRSs	No grade 3 and higher toxicities were reported, lower than those experienced with elective nodal RT (up to 2%).	⊕OOO VERY LOW	Downgraded 1 level for imprecision (i.e., not assessable)

### **Bone Cancer**

Number of Participants (N) Number of Studies Toxicity	Findings	Certainty of Evidence	Rationale
N = 160 1 RCT	No significant difference in toxicities for patients treated with SBRT or with MFRT. SBRT was associated with around 1% grade 3 or higher toxicities, and up to 10% for fatigue grade 3 and higher.	⊕⊕⊖O Low	Downgraded 1 level each for risk of bias and imprecision (i.e., not assessable)

### **Toxicity and Harms**

- Overall, SBRT was not associated with significantly higher rates of toxicity than other treatment options
  - Types of toxicity varied by treatment site
  - Events classed as grade 4 and 5 toxicities were rare
- No comparative toxicity for adrenal or renal cancers
- Very few reports in the FDA's Medical Device Recall database were classified as Class 1 (defined as a situation where there is a reasonable chance that a product will cause serious health problems or death)

Related to software and placement issues

• Similar safety issues were reported to the FDA's MAUDE database

## Subgroups

- Few studies reported on clinical subgroups of interest, but there was some indication specific populations (by cancer site) may be more at risk of SBRT-related toxicity compared with other populations
  - Varied by cancer type and treatment site and were often only reported in single studies

### **Clinical Practice Guidelines**

- Recommendations on the use of SBRT and payer policies varied in approach to the use of SBRT, with some guidelines or policies being more supportive of the use of SBRT depending on the cancer site
- Guidelines and payer policies often noted the limited evidence base, but also highlighted that SBRT may be preferred by patients because of fewer treatment fractions, and favorable safety profile

# Findings

Key Questions 4: Costs and Cost-Effectiveness



### **Prostate Cancer**

Study ID Study Risk-of-bias	Population	Intervention	Comparators	Economic Analytic Method
Pan et al., 2018 Low risk-of-bias	Men with localized prostate cancer	• SBRT	• IMRT	Cost comparison
Parikh et al., 2020 Low risk-of-bias	Men with oligorecurrent hormone-sensitive prostate cancer	• SBRT	<ul> <li>Abiraterone acetate plus prednisone and ADT</li> <li>Docetaxel and ADT</li> </ul>	Cost-effectiveness analysis (Markov state transition model)

### **Prostate Cancer**

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. IMRT for prostate c	ancer	•	
Outcome: cost-effectiveness	-		
N = 3 hypothetical cohorts 1 economic modelling study	Upfront SBRT may be a cost-effective option for people who wish to avoid systemic therapy; however, it was the cost-effective strategy in only 53.6% of microsimulations at a WTP of \$100,000 per QALY	⊕⊕⊖O Low	Downgraded 1 level for indirectness (i.e., oligometastatic hormone-resistant prostate cancer) and for imprecision (i.e., wide Cls)
Outcome: costs			· ·
N = 12,128 1 comparative NRS	SBRT had lower costs for both the payer (\$49,504 for SBRT and \$57,244 for IMRT; <i>P</i> < .001) and patient than IMRT (\$1,015 for SBRT and \$1,560 for IMRT; <i>P</i> < .001) No difference between treatments in complication costs or overall health care costs at 2 years	⊕OOO VERY LOW	Downgraded 1 level for indirectness (i.e., localized prostate cancer in younger men with private insurance)



Study ID Study Risk-of-bias	Population	Intervention	Comparators	Economic Analytic Method
Kim et al., 2019 Low risk-of-bias	People with oligometastatic stage IV NSCLC, grouped by mutation status	<ul> <li>SBRT plus maintenance therapy</li> </ul>		Cost-effectiveness analysis (Markov state transition model)

## Lung Cancer

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT plus maintenance therapy	vs maintenance therapy for lung cancer		
Outcome: cost-effectiveness			
N = 3 hypothetical cohorts	SBRT was assessed as not being cost-	$\oplus \oplus \oplus \bigcirc$	Downgraded 1 level for
1 economic modelling study	effective at a WTP threshold of \$100,000 when added to maintenance therapy for people with oligometastatic NSCLC.	MODERATE	indirectness (i.e., oligometastatic NSCLC only, by mutation status)

### **Pancreatic Cancer**

Study ID Study Risk-of-bias	Population	Intervention	Comparators	Economic Analytic Method
Moningi et al., 2022 Low risk-of-bias	People with non- metastatic, unresectable pancreatic cancer	• SBRT	<ul><li> cRT</li><li> Chemotherapy</li></ul>	Cost comparison

### **Pancreatic Cancer**

Number of Participants (N)	Findings	Certainty of Evidence	Rationale
Number of Studies			
SBRT vs. cRT or chemoth	erapy for pancreatic cancer		
Outcome: cost-effectiver	ness		
Not reported			
Outcome: costs			
N = 5,624	Healthcare payments were greatest for SBRT when	$\oplus 000$	Downgraded 1 level
1 comparative NRS	compared with cRT or chemotherapy under US Medicare ( $P < .001$ ) and employer-based insurance ( $P < .001$ ).	VERY LOW	for indirectness (i.e., nonmetastatic, unresectable pancreatic cancer)

### Head and Neck Cancers

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT plus maintenance therapy vs salvage therapies for head and neck cancer			
Outcome: cost-effectivenes	S		
N = 1 hypothetical cohort 1 economic modelling study	None of treatment strategies were cost-effective. However, SBRT-based reirradiation has potential to be cost-effective, as model was sensitive to median survival.	⊕⊕⊕O MODERATE	Downgraded 1 level for indirectness <sup>a</sup> (i.e., locoregional previously irradiated head and neck cancer)



Study ID Study Risk-of-bias	Population	Intervention		Economic Analytic Method
Parikh et al., 2018	People with early-	• SBRT	• RFA	Cost-effectiveness
Low risk-of-bias	stage liver cancer			analysis

### **Liver Cancer**

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. RFA for liver cancer			
Outcome: cost-effectiveness			
N = 440 1 comparative NRS, including a cost-effectiveness analysis	SBRT was not cost-effective compared with RFA in overall population of people with early-stage HCC; however, 85.5% of bootstrap ICER estimates were lower than WTP threshold of \$100,000.	⊕⊕⊕O MODERATE	Downgraded 1 level for indirectness (i.e., early-stage HCC only)

### **Oligometastatic Cancers**

Study ID Study Risk-of-bias	Population	Intervention	Comparators	Economic Analytic Method
Kumar et al., 2021 Low risk-of-bias	People with oligometastatic disease	• SBRT	Standard care	Cost-effectiveness analysis (Markov model)
Mehrens et al., 2021 Low risk-of-bias	People with oligometastatic disease	• SBRT	Standard care	Cost-effectiveness analysis (partitioned survival model)

### **Oligometastatic Cancers**

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. standard care for oligome	tastatic cancer		
Outcome: cost-effectiveness			
N = 2 hypothetical cohorts based on SABR-COMET data 2 economic modelling studies	Addition of SBRT increased costs and improved quality adjusted survival, overall leading to a cost- effective treatment strategy for patients with oligometastatic cancer.	⊕⊕⊕O MODERATE	Downgraded 1 level for indirectness (i.e., based on a single trial for patient outcomes)



Study ID	Population	Intervention	Comparators	Economic Analytic
Study Risk-of-bias				Method
Santos et al., 2021	People with bone	• SBRT	• EBRT	Cost comparison
Low risk of bias	metastases		• IMRT	

### **Bone Cancer**

Number of Participants (N) Number of Studies	Findings	Certainty of Evidence	Rationale
SBRT vs. other forms of RT fo	r bone cancer		
Outcome: costs			
N = 40,993 cases 1 comparative NRS	For people with bone metastases, the cost of SBRT was significantly higher for both professional and technical fees (\$679 lower provider costs and \$6,422 lower technical costs for external beam RT; \$36 lower provider costs and \$2,534 lower technical costs for IMRT; <i>P</i> < .001).	⊕OOO VERY LOW	Downgraded 1 level for indirectness (i.e., no indication how many were nonspine metastases)

### **Costs and Cost-Effectiveness**

- While the economic literature was sparse, SBRT appears to be:
  - Possibly cost-effective for oligometastatic hormone-resistant prostate cancer (low CoE, based on 1 economic modeling study)
  - Lower in costs than IMRT for prostate cancer (very low CoE, based on 1 comparative NRS)
  - Cost-ineffective when compared with maintenance therapy for oligometastatic lung cancer (moderate CoE, based on 1 economic modeling study)
  - Higher in costs than cRT or chemotherapy for pancreatic cancer (very low CoE, based on 1 comparative NRS)

### **Costs and Cost-Effectiveness**

- While the economic literature was sparse, SBRT appears to be:
  - Cost-ineffective as reirradiation when compared with other salvage therapies, including IMRT with chemotherapy, for head and neck cancers (moderate CoE, based on 1 economic modeling study)
  - Cost-ineffective when compared with RFA for liver cancer (low CoE, based on 1 economic modeling study)
  - Cost-effective when compared with standard of care for oligometastatic cancer (moderate CoE, based on 2 economic modeling studies)
  - More expensive than EBRT and IMRT for bone cancer (very low CoE, based on 1 comparative NRS)
- No identified economic evidence in adrenal cancer or renal cancer

# Conclusion



### Conclusion

- For some cancer sites, evidence shows SBRT has the potential to be an effective option when compared with other treatment options
  - Varies by specific type of cancer and comparative care
- However, SBRT for other cancers remains unsupported with limited or no comparative evidence of effectiveness
- Some guidelines are more supportive of the use of SBRT, but most note the limited evidence base, highlighting it may be preferred by patients because of the fewer treatment fractions and the favorable safety profile of SBRT when compared with other treatment options







#### HTCC Coverage and Reimbursement Determination Analytic Tool

### HTA's goal is to achieve *better health care outcomes* for enrollees and beneficiaries of state programs by paying for proven health *technologies that work*.

To find best outcomes and value for the state and the patient, the HTA program focuses on three questions:

- 1. Is it safe?
- 2. Is it effective?
- 3. Does it provide value (improve health outcome)?

The principles HTCC uses to review evidence and make determinations are:

#### Principle One: Determinations are evidence-based

HTCC requires scientific evidence that a health technology is safe, effective and cost-effective<sup>1</sup> as expressed by the following standards<sup>2</sup>:

- Persons will experience better health outcomes than if the health technology was not covered and that the benefits outweigh the harms.
- The HTCC emphasizes evidence that directly links the technology with health outcomes. Indirect evidence may be sufficient if it supports the principal links in the analytic framework.
- Although the HTCC acknowledges that subjective judgments do enter into the evaluation of evidence and the weighing of benefits and harms, its recommendations are not based largely on opinion.
- The HTCC is explicit about the scientific evidence relied upon for its determinations.

#### Principle Two: Determinations result in health benefit

The outcomes critical to HTCC in making coverage and reimbursement determinations are health benefits and harms<sup>3</sup>:

- In considering potential benefits, the HTCC focuses on absolute reductions in the risk of outcomes that people can feel or care about.
- In considering potential harms, the HTCC examines harms of all types, including physical, psychological, and non-medical harms that may occur sooner or later as a result of the use of the technology.
- Where possible, the HTCC considers the feasibility of future widespread implementation of the technology in making recommendations.

The principles and standards are based on USPSTF Principles at: <u>http://www.ahrq.gov/clinic/ajpmsuppl/harris3.htm</u>

Based on Legislative mandate: RCW 70.14.100(2).

- The HTCC generally takes a population perspective in weighing the magnitude of benefits against the magnitude of harms. In some situations, it may make a determination for a technology with a large potential benefit for a small proportion of the population.
- In assessing net benefits, the HTCC subjectively estimates the indicated population's value for each benefit and harm. When the HTCC judges that the balance of benefits and harms is likely to vary substantially within the population, coverage or reimbursement determinations may be more selective based on the variation.
- The HTCC considers the economic costs of the health technology in making determinations, but costs are the lowest priority.

#### Using evidence as the basis for a coverage decision

Arrive at the coverage decision by identifying for Safety, Effectiveness, and Cost whether (1) evidence is available, (2) the confidence in the evidence, and (3) applicability to decision.

#### 1. Availability of evidence:

Committee members identify the factors, often referred to as outcomes of interest, that are at issue around safety, effectiveness, and cost. Those deemed key factors are ones that impact the question of whether the particular technology improves health outcomes. Committee members then identify whether and what evidence is available related to each of the key factors.

#### 2. Sufficiency of the evidence:

Committee members discuss and assess the evidence available and its relevance to the key factors by discussion of the type, quality, and relevance of the evidence<sup>4</sup> using characteristics such as:

- Type of evidence as reported in the technology assessment or other evidence presented to committee (randomized trials, observational studies, case series, expert opinion);
- The amount of evidence (sparse to many number of evidence or events or individuals studied);
- Consistency of evidence (results vary or largely similar);
- Recency (timeliness of information);
- Directness of evidence (link between technology and outcome);
- Relevance of evidence (applicability to agency program and clients);
- Bias (likelihood of conflict of interest or lack of safeguards).

Sufficiency or insufficiency of the evidence is a judgment of each clinical committee member and correlates closely to the GRADE confidence decision.

Not Confident	Confident
Appreciable uncertainty exists. Further information is needed or further information is likely to change confidence.	Very certain of evidentiary support. Further information is unlikely to change confidence

<sup>4</sup> Based on GRADE recommendation: <u>http://www.gradeworkinggroup.org/FAQ/index.htm.</u>

#### 3. Factors for Consideration - Importance

At the end of discussion a vote is taken on whether sufficient evidence exists regarding the technology's safety, effectiveness, and cost. The committee must weigh the degree of importance that each particular key factor and the evidence that supports it has to the policy and coverage decision. Valuing the level of importance is factor or outcome specific but most often include, for areas of safety, effectiveness, and cost:

- Risk of event occurring;
- The degree of harm associated with risk;
- The number of risks; the burden of the condition;
- Burden untreated or treated with alternatives;
- The importance of the outcome (e.g. treatment prevents death vs. relief of symptom);
- The degree of effect (e.g. relief of all, none, or some symptom, duration, etc.);
- Value variation based on patient preference.

#### **Clinical committee findings and decisions**

#### Efficacy considerations

- What is the evidence that use of the technology results in more beneficial, important health outcomes? Consider:
  - Direct outcome or surrogate measure
  - o Short term or long term effect
  - o Magnitude of effect
  - o Impact on pain, functional restoration, quality of life
  - Disease management
- What is the evidence confirming that use of the technology results in a more beneficial outcome, compared to no treatment or placebo treatment?
- What is the evidence confirming that use of the technology results in a more beneficial outcome, compared to alternative treatment?
- What is the evidence of the magnitude of the benefit or the incremental value?
- Does the scientific evidence confirm that use of the technology can effectively replace other technologies or is this additive?
- For diagnostic tests, what is the evidence of a diagnostic tests' accuracy?
  - Does the use of the technology more accurately identify both those with the condition being evaluated and those without the condition being evaluated?
- Does the use of the technology result in better sensitivity and better specificity?
- Is there a tradeoff in sensitivity and specificity that on balance the diagnostic technology is thought to be more accurate than current diagnostic testing?
- Does use of the test change treatment choices?

#### Safety

- What is the evidence of the effect of using the technology on significant morbidity?
  - Frequent adverse effect on health, but unlikely to result in lasting harm or be life-threatening, or;
  - o Adverse effect on health that can result in lasting harm or can be life-threatening?
- Other morbidity concerns?
- Short term or direct complication versus long term complications?
- What is the evidence of using the technology on mortality does it result in fewer adverse non-fatal outcomes?

#### Cost impact

• Do the cost analyses show that use of the new technology will result in costs that are greater, equivalent or lower than management without use of the technology?

#### **Overall**

- What is the evidence about alternatives and comparisons to the alternatives?
- Does scientific evidence confirm that use of the technology results in better health outcomes than management without use of the technology?

#### Next step: Cover or no cover

If not covered, or covered unconditionally, the chair will instruct staff to write a proposed findings and decision document for review and final adoption at the following meeting.

#### Next step: Cover with conditions

If covered with conditions, the committee will continue discussion.

- 1) Does the committee have enough information to identify conditions or criteria?
  - Refer to evidence identification document and discussion.
  - Chair will facilitate discussion, and if enough members agree, conditions and/or criteria will be identified and listed.
  - Chair will instruct staff to write a proposed findings and decision document for review and final adoption at next meeting.
- 2) If not enough or appropriate information, then Chair will facilitate a discussion on the following:
  - What are the known conditions/criteria and evidence state
  - What issues need to be addressed and evidence state

The chair will delegate investigation and return to group based on information and issues identified. Information known but not available or assembled can be gathered by staff; additional clinical questions may need further research by evidence center or may need ad hoc advisory group; information on agency utilization, similar coverage decisions may need agency or other health plan input; information on current practice in community or beneficiary preference may need further public input. Delegation should include specific instructions on the

task, assignment or issue; include a time frame; provide direction on membership or input if a group is to be convened.

#### **Clinical committee evidence votes**

#### First voting question

The HTCC has reviewed and considered the technology assessment and information provided by the administrator, reports and/or testimony from an advisory group, and submissions or comments from the public. The committee has given greatest weight to the evidence it determined, based on objective factors, to be the most valid and reliable.

**Discussion document:** What are the key factors and health outcomes and what evidence is there? (Applies to the population in the PICO for this review)

Safety outcomes	Importance of outcome	Safety evidence/ confidence in evidence
Toxicity		

Efficacy – effectiveness outcomes	Importance of outcome	Efficacy / Effectiveness evidence
Overall survival		
Progression-free survival		
Disease-free survival		
Quality of life		
Objective response rate		
Complete response rate		

Cost outcomes	Importance of outcome	Cost evidence
Cost		
Cost-effectiveness		

Special population / Considerations outcomes	Importance of outcome	Special populations/ Considerations evidence
Age		
Sex		
Comorbidity		
Adolescents		
Pregnant individuals		

#### For safety:

Is there sufficient evidence that the technology is safe for the indications considered?

Unproven	Less	Equivalent	More in some	More in all
(no)	(yes)	(yes)	(yes)	(yes)

#### For efficacy/ effectiveness:

Is there sufficient evidence that the technology has a meaningful impact on patients and patient care?

Unproven	Less	Equivalent	More in some	More in all
(no)	(yes)	(yes)	(yes)	(yes)

#### For cost outcomes/ cost-effectiveness:

Is there sufficient evidence that the technology is cost-effective for the indications considered?

Unproven	Less	Equivalent	More in some	More in all
(no)	(yes)	(yes)	(yes)	(yes)

Discussion

Based on the evidence vote, the committee may be ready to take a vote on coverage or further discussion may be warranted to understand the differences of opinions or to discuss the implications of the vote on a final coverage decision.

- Evidence is insufficient to make a conclusion about whether the health technology is safe, efficacious, and cost-effective;
- Evidence is sufficient to conclude that the health technology is unsafe, ineffectual, or not cost-effective
- Evidence is sufficient to conclude that the health technology is safe, efficacious, and cost-effective for all indicated conditions;
- Evidence is sufficient to conclude that the health technology is safe, efficacious, and cost-effective for some conditions or in some situations

A straw vote may be taken to determine whether, and in what area, further discussion is necessary.

#### Second Vote

Based on the evidence about the technologies' safety, efficacy, and cost-effectiveness, it is:

Not covered	Covered unconditionally	Covered with conditions

#### **Discussion item**

Is the determination consistent with identified Medicare decisions and expert guidelines, and if not, what evidence is relied upon.

The report "identified no Medicare national coverage determination on the use of SBRT or any local coverage determinations that apply to the state of Washington."

#### **Medicare Coverage**

No Medicare National Coverage Determination (NCD)

#### **Selected Payer Coverage Determinations**

We identified no Medicare national coverage determination on the use of SBRT or any local coverage determinations that apply to the state of Washington.

Each of the 3 private payers that we reviewed, Aetna, Cigna, and, Regence, had coverage policies on the use of SBRT.<sup>310</sup>

Aetna considered SBRT as medically necessary in the following clinical conditions<sup>310</sup>:

- Stereotactic body radiation therapy with a CyberKnife, gamma knife, or linear accelerator (LINAC) is considered medically necessary for localized malignant conditions within the body where highly precise application of high-dose radiotherapy is required and clinically appropriate, including:
  - Hepatocellular carcinoma in individuals with unresectable disease considered extensive and not suitable for liver transplantation, or for individuals with local disease only with a good performance status (a score between 80 and 100 on the Karnofsky Performance Scale) but who are not amenable to surgery due to comorbidities
  - Prostate cancer in individuals with organ-confined prostate cancer with Gleason score less than or equal to 8 and prostate-specific antigen (PSA) less than 20

- o Non-small cell lung cancer for inoperable stage I or II tumors
- Oligometastatic colorectal cancer (1 to 3 metastases to the lung or liver) not amenable to surgery
- Inoperable primary spinal tumors with compression or intractable pain
- Recurrent metastatic disease in a previously irradiated area
- Recurrent localized head and neck cancer
- Metastatic lesions to the liver when the sole site of disease and cannot be surgically resected or undergo accepted ablation techniques
- Metastatic disease to the lung when clinically appropriate and on a case-by-case basis
- All other clinical sites or indications are considered experimental and investigational but will be considered on a case-by-case basis.
- Fractionated stereotactic radiotherapy is considered medically necessary when criteria for stereotactic radiosurgery are met. Fractionated stereotactic radiotherapy is useful for treatment of tumors in hard-to-reach locations, tumors with very unusual shapes, or for tumors located in such close proximity to a vital structure (e.g., optic nerve or hypothalamus) that even a very accurate high-dose single fraction of stereotactic radiosurgery could not be tolerated.

#### Aetna's coverage policy is due to be reviewed in early 2023.<sup>310</sup>

CIGNA has a series of recommendations on the use of the SBRT, reviewed in December of 2022<sup>311</sup>:

- Adrenal cancer
  - SBRT is considered not medically necessary in the adjuvant (post-operative) curative treatment of primary adrenocortical carcinoma.
- Bone metastases
  - SBRT using up to 5 fractions is considered not medically necessary for the treatment of bone metastases except in either of the following clinical scenarios:
    - Treatment to a portion of the spine that has been previously irradiated
    - Treatment of sarcoma, melanoma, and renal cell carcinoma that have metastasized to the spine. SBRT, as a complete course of therapy, must be completed in 5 fractions in a single episode of care.
  - SBRT is considered not to be medically necessary for all other bone metastases.
- Cervical cancer
  - SBRT as an alternative to brachytherapy is considered experimental, investigational, or unproven for the definitive treatment of cervical cancer.
  - SBRT is considered medically necessary based on a history of previous radiation to the same or abutting region and inability to deliver therapeutic doses of radiation with other techniques for locoregional recurrence of cervical cancer in an individual without evidence of distant metastases.
- Head and neck cancer
  - SBRT (up to 5 fractions) may be medically necessary for retreatment in an individual with head and neck cancer who has no evidence of metastatic disease. SBRT, as a complete course of therapy, must be completed in 5 fractions in a single episode of care.
- Liver and hepatobiliary cancer
  - The use of 3 to 5 fractions of SBRT is considered medically necessary to definitely treat concurrently 1 or more tumors in primary hepatocellular carcinoma when there is evidence of the ability to protect an adequate volume of uninvolved liver. SBRT, as a complete course of therapy, must be completed in 5 fractions in a single episode of care.
  - The use of up to 5 fractions of SBRT is considered medically necessary for the definitive treatment of intrahepatic bile duct cancer (cholangiocarcinoma).
  - The use of SBRT is considered not medically necessary for the definitive treatment of extrahepatic bile duct cancer (cholangiocarcinoma).
  - The use of SBRT is considered not medically necessary for adjuvant (postoperative) treatment of extrahepatic bile duct cancer (cholangiocarcinoma).
  - The use of SBRT is considered not medically necessary for definitive treatment of gall bladder cancer.

- The use of SBRT is considered not medically necessary for adjuvant (postoperative) treatment of gall bladder cancer.
- Renal cell carcinoma
  - The use of 3-dimensional conformal radiation therapy (3DCRT), intensity modulated radiation therapy (IMRT), or SBRT is considered not medically necessary in the definitive treatment of kidney cancer.
- Lung cancer
  - For stage I, node-negative stage IIA or T3N0 (T3 based on size) non-small cell lung cancer (NSCLC), the following regimens are considered medically necessary:
    - Definitive external beam radiation therapy to a dose of 60-70 Gy in 30-35 fractions using 3dimensional conformal radiation therapy (3DCRT)
    - Up to 5 fractions of stereotactic body radiation therapy (SBRT). SBRT, as a complete course of therapy, must be completed in 5 fractions in a single episode of care
  - For stage I or node-negative stage IIA limited-stage small-cell lung cancer (LSSCLC), the following regimens are considered medically necessary:
    - 3D conformal radiation therapy to a dose of 60-70 Gy in 30-35 fractions or 45 Gy delivered twice daily
    - Up to 5 fractions of stereotactic body radiation therapy (SBRT). SBRT, as a complete course of therapy, must be completed in 5 fractions in a single episode of care.
- Oligometastatic cancer
  - Up to 5 fractions of stereotactic body radiotherapy (SBRT) for extra-cranial oligometastases is considered medically necessary in the following clinical situations:
    - For an individual with non-small cell lung cancer who meets all of the following criteria:
      - Has had or will undergo curative treatment of the primary tumor (based on T and N stage)
      - Has 1 to 3 metastases in the synchronous setting
    - For an individual with colorectal cancer who meets all of the following criteria:
      - Has had or will undergo curative treatment of the primary tumor
      - Presents with 1 to 3 metastases in the lung or liver in the synchronous setting
      - For whom surgical resection is not possible
    - For an individual who meets the following criteria:
      - A clinical presentation of 1 to 3 adrenal gland, lung, liver, or bone metastases in the metachronous setting when ALL of the following criteria are met:
        - Histology is non-small cell lung, colorectal, breast, sarcoma, renal cell, melanoma, or prostate
        - Disease free interval of > 1 year from the initial diagnosis
        - Primary tumor received curative therapy and is controlled
        - No previous evidence of metastatic disease (cranial or extracranial)
        - All metastatic lesions present on imaging will be treated concurrently in a single episode of care
      - SBRT used to stimulate the abscopal effect is considered experimental, investigational, or unproven.
    - For an individual with oligoprogression (progression of a limited number of metastatic sites while other metastatic disease sites remain controlled), SBRT is considered not medically necessary.
    - SBRT, as a complete course of therapy, must be completed in 5 fractions in a single episode of care.
- Pancreatic cancer
  - SBRT using up to 5 fractions is considered medically necessary for curative treatment of unresectable/locally advanced cases and as preoperative treatment in borderlines resectable cases.
  - SBRT is considered not medically necessary in the palliative setting, postoperative setting, or for planned neoadjuvant treatment when the primary tumor is otherwise fully resectable.

- Prostate cancer
  - The following treatments are considered medically necessary for treatment of low-risk prostate cancer:
    - Hypofractionation 20-28 fractions of IMRT in up to 2 phases
    - Up to 5 fractions of SBRT alone (i.e., not as a boost)
  - The following treatments are considered medically necessary for treatment of intermediate-risk prostate cancer:
    - Hypofractionation 20-28 fractions of IMRT in up to 2 phases
    - Up to 5 fractions of SBRT alone (i.e., not as a boost)
  - The following treatments are considered medically necessary for treatment of high-risk prostate cancer when not treating the pelvic lymph nodes:
    - Hypofractionation 20-28 fractions of IMRT in up to 2 phases
    - Up to 5 fractions of SBRT alone (i.e., not as a boost)
- Melanoma

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- The use of SBRT to induce the abscopal effect is considered experimental, investigational, or unproven.
- Soft-tissue sarcoma
  - Up to 5 fractions of SBRT is considered medically necessary in the treatment of recurrent softtissue sarcoma located within a previously irradiated area.

Regence includes the following in its coverage policy for SBRT<sup>312</sup>:

- SRS and SBRT, also known as SABR, may be considered medically necessary for initial treatment or treatment of recurrence for any of the following indications:
  - Head and neck cancers outside of intracranial, skull base, and orbital sites, when there is documented previous radiation treatment to the planned target volume
  - o Hemangioblastoma of the spine
  - Hemangiopericytoma outside of intracranial, skull base, or orbital sites
  - Hepatic tumor (excluding hepatocellular carcinoma; primary or metastatic) as palliative or curative treatment when both of the following are met:
    - Absence or minimal extra hepatic disease; and
    - Karnofsky performance score greater than or equal to 60 or an ECOG score less than or equal to 2
  - Hepatocellular carcinoma (hepatoma) when all of the following criteria are met:
    - 5 or fewer hepatic lesions; and
    - Size of largest lesion is 6 cm diameter or less; and
    - Karnofsky performance score greater than or equal to 60 or an ECOG score less than or equal to 2
  - Lung metastases when both of the following criteria are met:
    - 5 or fewer metastatic lung lesions; and
    - Karnofsky performance score greater than or equal to 60 or an ECOG score less than or equal to 2
  - Non-small cell lung cancer (NSCLC), primary (node negative, tumor stage T1 and T2)
    - Oligometastases when the following criteria are met:
    - 5 or fewer metastatic lesions; and
    - Primary is controlled, stable, or expectation of the same; and
    - Metastases are limited to one to three organs; and
    - Karnofsky performance score greater than or equal to 60 or an ECOG score less than or equal to 2
  - O Osteosarcoma, metastatic when all of the following criteria are met:
    - 5 or fewer metastatic lesions; and
    - Karnofsky performance score greater than or equal to 60 or an ECOG score less than or equal to 2

- Pancreatic adenocarcinoma, locally advanced, borderline resectable, inoperable, or local recurrence after resection
- Paraganglioma
- Prostate cancer, very low- to intermediate-risk
- o Renal cell cancer, inoperable primary, when a urological surgeon has documented inoperability
- o Schwannomas
- Spinal or paraspinal tumors (primary or metastatic)
- SRS and SBRT (also known as SABR) are considered investigational when the first criterion is not met and for all other indications outside of intracranial, skull base, or orbital
- sites, including but not limited to:
  - Tumors, primary, of the cervix, endometrium, esophagus, hemangiomas, large bowel, ovaries, rectum, and small bowel

#### **Clinical Practice Guidelines**

#### **Prostate Cancer**

Table 1. Excerpted Clinical Practice Recommendations on Stereotactic Body Radiation Therapy for Prostate Cancer

Organization and Year Title	Recommendations	
Methodological Quality		
American Society for Radiation Oncology and American Urological Association (ASTRO/AUA),	Clinicians may offer ultrahypofractionated EBRT for patients with low- or intermediate- risk prostate cancer who elect EBRT.	Strong recommendation; evidence level: grade A
2022	In patients with low- or favorable intermediate-risk prostate cancer electing	Strong recommendation; evidence level: grade B
Clinically localized prostate cancer: AUA/ASTRO guideline, part I <sup>281</sup> ; part II <sup>282</sup> ; and part III <sup>29</sup>	radiation therapy, clinicians should offer dose-escalated hypofractionated EBRT (moderate or ultra), permanent low-dose rate (LDR) seed implant, or temporary high-	
Good methodological quality	dose rate (HDR) prostate implant as equivalent forms of treatment.	
Prostate Cancer Guidelines Panel, 2022	SBRT was discussed in literature review but panel concluded there was not enough	Available evidence is of low quality; strong
EAU - EANM - ESTRO - ESUR - ISU <i>P</i> - SIOG guidelines on prostate cancer <sup>283</sup>	evidence to make recommendations on its use.	recommendations cannot be made.
Good methodological quality		
Australian and New Zealand Faculty of Radiation Oncology Genito-Urinary Group (FROGG), 2018 Radiotherapy for recurrent prostate cancer: 2018 Recommendations of the	3.4. Recurrence limited to pelvic lymph nodes after curative local treatment: SBRT alone to involved node(s) may be considered in selected patients, but these patients should be informed that they are at high risk of relapse which may be harder to treat with curative intent	5, D • Expert opinion without explicit critical appraisal, or based on physiology, bench research or "first principles"

Organization and Year	Recommendations	
Title		
Methodological Quality		
Australian and New Zealand Radiation Oncology Genito- Urinary group <sup>284</sup>		• Level 5 evidence or troublingly inconsistent or inconclusive studies
Poor methodological quality	Radiotherapy management of oligometastases: 4.3. Patients should be encouraged to enter clinical trials where available to ascertain potential benefit of SBRT in addition to standard of care systemic therapy	of any level 4, D • Case-series (and poor- quality cohort and case-control studies • Level 5 evidence or troublingly inconsistent or inconclusive studies of any level
	5.4. If considering salvage local therapy, options include salvage prostatectomy, brachytherapy, stereotactic radiotherapy, HIFU, cryotherapy and salvage electroporation. Local salvage treatments are associated with significant toxicity and an individualized-treatment approach is recommended.	<ul> <li>4, C</li> <li>Case-series (and poor- quality cohort and case-control studies</li> <li>Level 4 studies or extrapolations from level 2 or 3 studies</li> </ul>
	Suitable patients should be considered for clinical trials.	

Abbreviations. EANM: European Association of Nuclear Medicine; EAU: European Association of Urologists; EBRT: external beam radiation therapy; ESTRO: European SocieTy for Radiotherapy and Oncology; ESUR: European Society of Urogenital Radiotherapy; ISUP: International Society of Urological Pathology; SBRT: stereotactic body radiation therapy; SIOG: International Society of Geriatric Oncology.

#### Lung Cancer

 Table 2. Excerpted Clinical Practice Recommendations on Stereotactic Body Radiation Therapy

 for Lung Cancer

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
American Society of Clinical Oncology (ASCO), 2021 Radiation therapy for small-cell lung cancer: ASCO guideline endorsement of an ASTRO guideline <sup>285</sup> Good methodological quality	<ul> <li>Recommendation 2.1. For patients with stage I or II node-negative LS- SCLC who are medically inoperable, either SBRT or conventional fractionation is recommended.</li> <li>Ultracentral tumors (ASCO clarifying comment: meaning those with the planning target</li> </ul>	Strength of recommendation: strong Quality of evidence: moderate

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
	volume touching or overlapping the proximal bronchial tree, esophagus, or trachea) may be more appropriately treated with conventional fractionation schema.	
Society of Interventional Radiology (SIR), 2021 Society of Interventional Radiology multidisciplinary position statement on percutaneous ablation of non- small cell lung cancer and metastatic disease to the lungs: endorsed by the Canadian Association for Interventional Radiology, the Cardiovascular and Interventional Radiological Society of Europe, and the Society of Interventional Oncology <sup>287</sup>	In patients with stage IA NSCLC, image-guided thermal ablation is a safe and effective treatment with minimal complications and acceptable long-term oncological and survival outcomes that are comparable to SBRT and sublobar resection. Thermal ablation should be considered alongside surgical resection and SBRT in patients who require preservation of lung parenchyma function.	Level C, moderate quality <ul> <li>Nonrandomized studies</li> <li>Supported by moderate quality evidence for or against recommendation; new research may be able to provide additional context</li> </ul>
Moderate methodological quality		
European Society for Medical Oncology (ESMO), 2020 (update of 2018) Metastatic non-small cell lung cancer: ESMO Clinical Practice Guidelines for diagnosis, treatment and follow-up <sup>289</sup> and Metastatic Non-Small-Cell Lung Cancer: ESMO Clinical Practice Guidelines for diagnosis, treatment and follow-up 2020 Update <sup>286</sup> Moderate methodological quality	Stage IV patients with limited synchronous metastases at diagnosis may experience long- term disease-free survival (DFS) following systemic therapy and local consolidative therapy [LCT: high-dose RT including stereotactic ablative body RT (SABR) or surgery]	<ul> <li>Level IIIB</li> <li>Prospective cohort studies</li> <li>Strong or moderate evidence for efficacy but with a limited clinical benefit, generally recommended</li> </ul>
National Institute for Health and Care	1.6.5. For people with stage I–IIA	Sublobar resection and
Excellence (NICE), 2018 Lung cancer: diagnosis and management [NG122] <sup>288</sup> Good methodological quality	(T1a–T2b, N0, M0) NSCLC who decline lobectomy or in whom it is contraindicated, offer radical radiotherapy with stereotactic ablative radiotherapy (SABR) or sublobar resection.	SABR [] not clear which is better.
	1.6.8. For people with stage I–IIA (T1a–T2b, N0, M0) NSCLC who decline surgery or in whom any surgery is contraindicated, offer	SABR provides better survival outcomes [] people often prefer it

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
	SABR. If SABR is contraindicated, offer either conventional or hyperfractionated radiotherapy.	because it involves fewer hospital visits.
	1.6.9. For eligible people with stage IIIA NSCLC who cannot tolerate or who decline chemoradiotherapy (with or without surgery), consider radical radiotherapy (either conventional or hyperfractionated).	Evidence was not strong enough to recommend conventional radiotherapy over hyperfractionated regimens or vice versa.

Abbreviations. LS-SCLC: limited-stage small-cell lung cancer; NSCLC: non-small cell lung cancer; SBRT: stereotactic body radiation therapy.

# **Gynecological Cancers**

 Table 3. Excerpted Clinical Practice Recommendations on Stereotactic Body Radiation Therapy

 for Gynecologic Cancers

Organization and Year	Recommendations	Strength of
Title		Recommendation
Methodological Quality		
European Society of Gynaecological Oncology (ESGO), 2018 European Society of Gynaecological Oncology/European Society for Radiotherapy and Oncology/European Society of Pathology Guidelines for management of patients with cervical cancer <sup>293</sup> Good methodological quality	<ul> <li>Pelvic sidewall recurrence after primary surgery</li> <li>Definitive radiotherapy or chemoradiotherapy followed by a stereotactic ablative boost/image-guided interstitial brachytherapy/particle beam therapy is an emerging option.</li> <li>Central pelvic or pelvic sidewall recurrence after radiotherapy or chemoradiotherapy</li> <li>Management of isolated organ metastases (lung, liver, etc.) should be discussed in a multidisciplinary team involved in treatment of specific organ affected by metastasis and should be treated according to preferred method for that organ involving local resection, radiofrequency ablation, interventional brachytherapy, or stereotactic ablative</li> </ul>	Not provided

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
European Society of Gynaecological Oncology (ESGO), 2020 ESGO/ESTRO/ESP guidelines for management of patients with endometrial carcinoma <sup>294</sup> Good methodological quality	radiotherapy according to size and anatomical position. Radiotherapy pretreated patients with locoregional recurrence • If surgery is not feasible, radical re-irradiation options include stereotactic body radiotherapy targeting recurrence, permanent seed implants, or proton therapy. In selected cases, limited volume re-irradiation with EBRT and brachytherapy boost may be an option (especially if longer interval from first irradiation).	<ul> <li>IV, C</li> <li>Retrospective cohort studies or case-control studies</li> <li>Insufficient evidence for efficacy or benefit does not outweigh risk or disadvantages (adverse events, costs, etc.), optional</li> </ul>
	Oligometastatic recurrent disease • Treatment options include: surgery, radiation therapy including stereotactic radiotherapy, and local ablating techniques	<ul> <li>IV,5 B</li> <li>Retrospective cohort studies or case-control studies</li> <li>Insufficient evidence for efficacy or benefit does not outweigh risk or disadvantages (adverse events, costs, etc.), optional</li> </ul>

Abbreviations. ESP: European Society of Pathology; ESTRO: European SocieTy for Radiotherapy and Oncology.

### Melanoma

Table 4. Excerpted Clinical Practice Recommendations on Stereotactic Body Radiation Therapy for Melanoma

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
European Society for Medical Oncology (ESMO), 2019 Cutaneous melanoma: ESMO clinical practice guidelines for diagnosis, treatment and follow-up Approved by the ESMO Guidelines Committee: February 2002, last update September 2019 <sup>295</sup>	Surgical removal or stereotactic irradiation of locoregional recurrence or single distant metastasis should be considered in fit patients, as a therapeutic option, offering potential for long-term disease control.	<ul> <li>III, C</li> <li>Prospective cohort studies</li> <li>Insufficient evidence for efficacy or benefit does not outweigh risk or disadvantages (adverse events, costs, etc.), optional</li> </ul>

Organization and Year Title	Recommendations	Strength of Recommendation
Methodological Quality		
Moderate methodological quality		

# **Renal Cancer**

# Table 5. Excerpted Clinical Practice Recommendations on Stereotactic Body Radiation Therapy for Renal Cancer

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
National Comprehensive Cancer Network (NCCN), 2022 Kidney Cancer, Version 3.2022 Moderate methodological quality	Resection is preferred over locally ablative procedures (e.g., image- guided ablation or SBRT However, these local techniques can be considered for liver or lung oligometastases	Not reported
	In patients with a limited number of liver or lung metastases, ablative radiotherapy to the metastatic site can be considered in highly selected cases or in the setting of a clinical trial. Radiotherapy should not be used in the place of surgical resection. Radiotherapy should be delivered in a highly conformal manner. The techniques can include 3D conformal radiation therapy, intensity-modulated radiation therapy (IMRT), or SBRT	
	SBRT may be considered for medically inoperable patients with Stage I kidney cancer (category 2B), with Stage II/III kidney cancer (both category 3)	
European Association of Urology (EAU), 2022	Local therapy of metastases in metastatic RCC • "Offer stereotactic radiotherapy for clinically	Weak

Organization and Year Title	Recommendations	Strength of Recommendation
Methodological Quality		
EAU guidelines on renal cell carcinoma <sup>297</sup> Good methodological quality	relevant bone- or brain metastases for local control and symptom relief."	
	<ul> <li>Local ablative therapy</li> <li>"Although early results of [SBRT] are encouraging, more evidence from randomised trials is needed."</li> </ul>	
American Urology Association (AUA), 2021	"Non-extirpative methods, eg, stereotactic-body-radiation-	Not applicable
Renal mass and localized renal cancer: evaluation, management, and follow-up: AUA guideline: Part I <sup>296</sup> and Part II <sup>299</sup>	therapy or high-intensity-focused- ultrasound, are still investigational." (Pt 1)	
Good methodological quality		
European Society for Medical Oncology (ESMO), 2019	Management of advanced/metastatic disease	IV, B • Retrospective cohort studies
Renal cell carcinoma: ESMO Clinical Practice Guidelines for diagnosis, treatment and follow- up	<ul> <li>RT can be used to treat unresectable local or recurrent disease and in patients unsuitable for surgery due to poor PS or</li> </ul>	or case-control studies • Strong or moderate evidence for efficacy but with a limited clinical benefit, generally recommended
Approved by ESMO Guidelines Committee: September 2008, last update January 2019. This publication supersedes previously published version— Ann Oncol 2016; 27 (Suppl 5): v58 to v68 <sup>298</sup>	unsuitable clinical condition. RT is an alternative if radioablation is not appropriate. Image-guided RT techniques such as VMAT or SBRT are needed to enable a high dose to be delivered.	
Moderate methodological quality		

Abbreviations. PS: performance status; RCC: renal cell carcinoma; RT: radiation therapy; SBRT: stereotactic body radiotherapy; VMAT: volumetric-modulated arc therapy.

# **Pancreatic Cancer**

Organization and Year	Recommendations	Strength of Recommendation
Title		Recommendation
Methodological Quality		
American Society for Radiation Oncology (ASTRO), 2019	a. Following surgical resection of pancreatic cancer, adjuvant SBRT is only recommended on a clinical trial or multi-institutional registry.	Strong recommendation
Radiation Therapy for Pancreatic Cancer:	that of mater institutional registry.	Very low quality of evidence; 100% consensus
Executive Summary of an ASTRO Clinical Practice Guideline <sup>301</sup> Good methodological quality	b. For patients with borderline resectable pancreatic cancer and select locally advanced pancreatic cancer appropriate for downstaging prior to surgery, a neoadjuvant therapy regimen of systemic chemotherapy followed by multifraction SBRT is conditionally recommended.	Conditional recommendation Low quality of evidence; 77% consensus
	c. For patients with locally advanced pancreatic cancer not appropriate for downstaging to eventual surgery, a definitive therapy regimen of systemic chemotherapy followed by either (1) conventionally fractionated RT with chemotherapy, (2) dose- escalated chemoradiation, or (3) multifraction SBRT without chemotherapy is conditionally recommended.	Conditional recommendation Low quality of evidence; 77% consensus

# Table 6. Excerpted Clinical Practice Recommendations on Stereotactic Body Radiation Therapy for Pancreatic Cancer

Abbreviations. RT: radiation therapy; SBRT: stereotactic body radiation therapy.

# Liver and Biliary Tract Cancer

Table 7. Excerpted Clinical Practice Recommendations on Stereotactic Body Radiation Therapy for Liver and Biliary Tract Cancer

Organization and Year	Recommendations	Strength of Recommendation
Title		
Methodological Quality		
American College of Radiology (ACR), 2022	<i>Note. The recommendations list it as "EBRT" but discussion shows that they mean SBRT.</i>	Not provided
American College of Radiology ACR appropriateness criteria management of liver cancer <sup>302</sup>	<ul> <li>Hepatocellular cancer</li> <li>Solitary tumor less than 3 cm, cirrhotic - may be appropriate</li> <li>Solitary tumor 3 to 5 cm, cirrhotic - may be appropriate</li> </ul>	

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
Good methodological quality	<ul> <li>Multifocal, bilobar disease, at least 1 tumor greater than 5 cm, cirrhotic - may be appropriate</li> <li>Solitary or multifocal disease with vascular invasion, cirrhotic - may be appropriate</li> </ul>	
	<ul> <li>Intrahepatic cholangiocarcinoma</li> <li>Peripheral hepatic lobar cholangiocarcinoma, less than 3 cm; no biliary ductal dilatation, macroscopic vascular invasion, regional lymphadenopathy, or distant metastases - may be appropriate</li> </ul>	
	<ul> <li>Ductal cholangiocarcinoma</li> <li>Hilar cholangiocarcinoma, greater than 3 cm with poorly defined margins, vascular invasion, and periportal lymphadenopathy - may be appropriate</li> </ul>	
	<ul> <li>Metastatic liver disease</li> <li>Multifocal metastatic neuroendocrine tumor (includes carcinoid tumors as well as islet cell tumors of the pancreas) - may be appropriate</li> <li>Solitary colorectal liver metastasis - may be appropriate</li> <li>Multifocal bilobar colorectal carcinoma (liver dominant or isolated) - usually not appropriate</li> </ul>	
American Society for Radiation Oncology (ASTRO), 2022 External beam radiation therapy for primary liver cancers: an ASTRO clinical practice guideline <sup>303</sup> Good methodological quality	Note: SBRT is described as ultrahypofractionated EBRT. a. For patients with HCC who are potential candidates for OLT, ultra- or moderately hypofractionated EBRT is conditionally recommended as a bridge to transplant or as a downstaging intervention.	Strength of recommendation: conditional Quality of evidence: low
	b. For patients with liver-confined HCC, for whom EBRT is recommended, dose-escalated ultra- or moderately hypofractionated EBRT is recommended, with choice of regimen based on tumor location, underlying liver function, and available technology.	Strength of recommendation: strong Quality of evidence: moderate

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
	c. For patients with unresectable IHC receiving EBRT, dose-escalated ultra- or moderately hypofractionated EBRT is conditionally recommended with fractionation based on tumor location, underlying liver function, and available technology. Implementation remark: Concurrent systemic therapy should not be used with ultrahypofractionated EBRT.	Strength of recommendation: conditional Quality of evidence: low
European Society for Medical Oncology (ESMO), 2022 Biliary tract cancer: ESMO clinical practice guideline for diagnosis, treatment and follow-up <sup>304</sup> Moderate methodological quality	SBRT can be considered for patients with IHC in case of contraindication to surgery for liver- limited disease in palliative setting.	<ul> <li>III, C</li> <li>Prospective cohort studies</li> <li>Insufficient evidence for efficacy or benefit does not outweigh the risk or the disadvantages (adverse events, costs, etc.), optional</li> </ul>
European Society for Medical Oncology (ESMO), 2018 Hepatocellular carcinoma: ESMO clinical practice guidelines for diagnosis, treatment and follow-up <sup>305</sup> Moderate methodological quality	High conformal HDR radioablation and SBRT may be considered as alternatives for ablation of tumors with a high risk of local failure after thermal ablation due to location.	<ul> <li>III, C</li> <li>Prospective cohort studies</li> <li>Insufficient evidence for efficacy or benefit does not outweigh risk or disadvantages (adverse events, costs, etc.), optional</li> </ul>

Abbreviations. EBRT: external beam radiation therapy; HCC: hepatocellular carcinoma; HDR: high dose rate; IHC: intrahepatic cholangiocarcinoma; OLT: orthotopic liver transplantation; SBRT: stereotactic body radiation therapy.

## **Bone Cancers**

 Table 8. Excerpted Clinical Practice Recommendations on Stereotactic Body Radiation Therapy

 for Bone Cancer

Organization and Year	Recommendations	Strength of Recommendation
Title Methodological Quality		Recommendation
European Society for Medical Oncology (ESMO), 2021 Bone sarcomas: ESMO- EURACAN-GENTURIS- ERN PaedCan clinical practice guideline for diagnosis, treatment, and follow-up <sup>306</sup> Moderate methodological quality	a. "For lung metastases, stereotactic RT, radiofrequency ablation (RFA) or cryotherapy might be used as alternative options in patients unfit for surgery [IV, B]. Some groups also consider RFA and stereotactic RT as potentially alternative local treatment options for bone metastases." <i>Note. Lung metastases from primary bone cancer.</i>	<ul> <li>IV, B</li> <li>Retrospective cohort studies or case-control studies</li> <li>Strong or moderate evidence for efficacy but with a limited clinical benefit, generally recommended</li> </ul>
	b. "For oligometastatic disease, surgery, RFA, cryotherapy or stereotactic RT can be considered in selected cases."	<ul> <li>V, B</li> <li>Studies without control group, case reports, expert opinions</li> <li>Strong or moderate evidence for efficacy but with a limited clinical benefit, generally recommended</li> </ul>
	c. "RFA and stereotactic RT are potential alternative local treatment options in patients unfit for surgery and for small lung or bone metastases."	<ul> <li>V, B</li> <li>Studies without control group, case reports, expert opinions</li> <li>Strong or moderate evidence for efficacy but with a limited clinical benefit, generally recommended</li> </ul>
Spanish Society of Radiation Oncology (SEOR), 2022 SEOR SBRT-SG	"it is not possible to clearly differentiate between patients who are candidates for SBRT and those who should undergo prophylactic surgery."	Not provided
stereotactic body radiation therapy consensus guidelines for nonspine bone metastasis <sup>307</sup>	"The initial evaluation of patients with NSBM who are potential candidates for SBRT must take into account the performance status of patients"	
Poor methodological quality	"The use of SBRT in polymetastatic patients in whom not all lesions are susceptible to radical local treatment (SBRT or surgery) has been published but is not the standard of care."	

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
	"The authors recommended a single fraction as the first treatment option because this scheme requires fewer hospital resources and a shorter hospital stay, an important benefit, especially in the context of the current pandemic. Notwithstanding that recommendation, the most widely accepted fractionation schedules to ensure a BED $\geq$ 60 Gy are a single fraction of 20–24 Gy, three fractions of 10 Gy each, or five fractions of 7–10 Gy."	
	"Dose de-escalation—defined as more fractions with a lower dose per fraction—should be performed if the lesion has previously been treated with SBRT or EBRT (provided that > 3 months have elapsed between treatments), or if the lesions involve weight-bearing bones, or in patients with moderate-severe (≥ 30%) cortical erosion."	
	"Dose escalation should be considered in metastases with a radiation-resistant histology (e.g., colon, kidney, melanoma, and sarcoma) or if bulky mass or extraosseous involvement is present."	

Abbreviations. BED: biologically equivalent dose; EBRT: external beam radiation therapy; EURACAN: European Reference Network for rare adult solid cancers; GENTURIS ERN: European Reference Network for all patients with one of the rare genetic tumor risk syndromes; NSBM: nonspine bone metastases; RT: radiation therapy.

## **Testicular Cancer**

Table 9. Excerpted Clinical Practice Recommendations on Stereotactic Body Radiation Therapy for Testicular Cancer

Organization and Year Title Methodological Quality	Recommendations	Strength of Recommendation
European Society for Medical Oncology (ESMO), 2022 Testicular seminoma and non- seminoma: ESMO-EURACAN clinical practice guideline for diagnosis, treatment and follow- up <sup>308</sup>	<ul> <li>Salvage treatment</li> <li>"Principally, all ablative therapies, including stereotactic RT and radiofrequency ablation, should be considered within a multidisciplinary approach with an expert centre."</li> </ul>	Not provided

Moderate methodological quality		
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Abbreviations. EURACAN: European Reference Network for rare adult solid cancers; RT: radiation therapy.

#### Next step: proposed findings and decision and public comment

At the next public meeting the committee will review the proposed findings and decision and consider any public comments as appropriate prior to a vote for final adoption of the determination.

- 1) Based on public comment was evidence overlooked in the process that should be considered?
- 2) Does the proposed findings and decision document clearly convey the intended coverage determination based on review and consideration of the evidence?

#### Next step: final determination

Following review of the proposed findings and decision document and public comments:

#### **Final vote**

Does the committee approve the Findings and Decisions document with any changes noted in discussion?

If yes, the process is concluded.

If no, or an unclear (i.e., tie) outcome chair will lead discussion to determine next steps.



#### **Final Key Questions and Background**

# **Use of Stereotactic Body Radiation Therapy**

## Background

#### **Technology of Interest**

Radiation therapy is a cancer treatment that uses high-energy X-ray or other particles to destroy cancer cells<sup>1</sup>. A radiation therapy regimen, or schedule, usually consists of a specific number of treatments given over a set period, and can be used to treat different types of cancer.<sup>1</sup> Radiation therapy can also be used in combination with other cancer treatments, such as chemotherapy or surgery.<sup>1</sup>

The most common type of radiation therapy is external-beam radiation therapy (EBRT), which delivers radiation from outside the body.<sup>1</sup> The different types of external-beam radiation therapy are<sup>1</sup>:

- 3-dimensional conformal radiation therapy (3D-CRT)
- Intensity modulated radiation therapy (IMRT)
- Proton beam therapy
- Image-guided radiation therapy (IGRT
- Stereotactic body radiation therapy (SBRT)

SBRT is defined as extracranial stereotactic ablative treatment delivery (which can include the spine) typically delivered in 1 to 5 fractions and is also referred to as stereotactic ablative radiotherapy (SABR).<sup>2</sup> SBRT can be used for a variety of clinical indications as primary treatment for selected early-stage cancers, as treatment for discrete tumors in patients with oligometastatic disease, for selected benign neoplasms in or near the central nervous system (CNS), or in recurrent cancer within previously irradiated regions.<sup>2</sup>

Other radiation-based therapies include implanted internal radiation therapy (or brachytherapy), intraoperative radiation therapy (IORT), systemic radiation therapy, radioimmunotherapy, and may also involve the use of radiosensitizers or radioprotectors.<sup>1</sup>

#### **Clinical Need and Target Populations**

In 2019, a total of 1,752,735 new invasive cancer cases were reported in the US: 863,830 among females and 888,905 among males.<sup>3</sup> For all cancers combined, the incidence rate was 439 per 100,000 standard population overall.<sup>3</sup> While cancer affects people of all ages, races, ethnicities, and sexes, it does not affect all groups equally.<sup>3</sup> Differences in genetics, healthy choices, environmental exposures, and other factors can lead to differences in risk among groups of people.<sup>3</sup> For most cancers, increasing age is the most important risk factor, with around 58% of cancers occurring in adults aged 65 years or older.<sup>3</sup>

#### **Policy Context**

The use of SBRT for various cancers is increasing in the US<sup>4-6</sup>; however, its effectiveness and safety in routine clinical practice for most cancers are unclear. This topic was originally selected

for the 2012 review because of medium-level concerns about the safety and efficacy of CMRA and high-level concern about costs. This topic was selected for re-review based on new evidence that could prompt potential coverage policy changes.

In 2012 the Washington State HTCC commissioned an evidence review on the effectiveness of stereotactic radiosurgery (SRS) and SBRT for treating various cancers.<sup>7</sup> On March 22, 2013, using that evidence review to guide decision making, the committee adopted the following coverage determination<sup>8</sup>:

- SRS for central nervous system (CNS) primary and metastatic tumors is a covered benefit for adults and children when the following criteria are met:
  - Patient functional status score (i.e., Karnofsky score) is greater than or equal to 50; and
  - Evaluation includes multidisciplinary team analysis (e.g., tumor board), including surgical input.
- SBRT is covered for adults and children for the following conditions when the following criteria are met:
  - For cancers of spine/paraspinal structures; or
  - For inoperable non-small cell lung cancer (NSCLC), stage 1; and
  - Evaluation includes multidisciplinary team analysis, including surgical input.
- All other indications are noncovered.

The objective of the health technology assessment (HTA) is to evaluate the effectiveness, safety, and cost-effectiveness of SBRT in adults and children with cancers not currently covered by the 2012 coverage decision (CNS and a subset of lung cancers). This evidence review will help inform Washington's independent Health Technology Clinical Committee as the committee determines coverage regarding the use of SBRT in adults and children with cancers not currently covered.

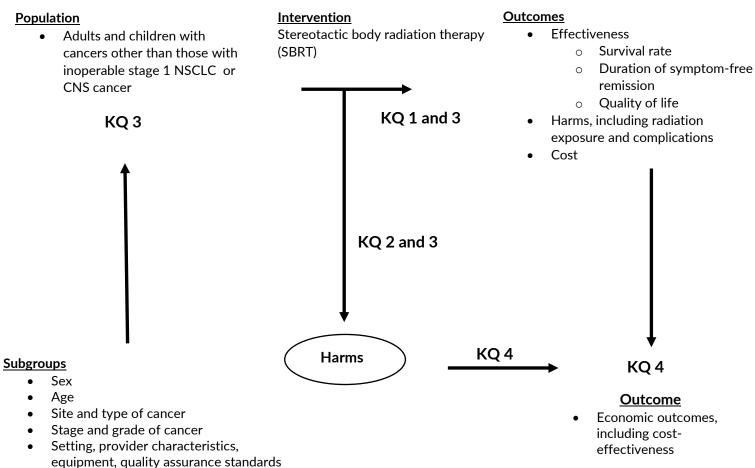
## **Key Questions**

- KQ1. What is the evidence of effectiveness for SBRT for patients with cancers not currently covered (CNS cancers and inoperable stage 1 NSCLC)?
- KQ2. What are the harms of SBRT in patients with included cancers?
- KQ3. What is the evidence that SBRT has differential efficacy or harms in subpopulations, including those defined by:
  - a. Sex
  - b. Age
  - c. Site and type of cancer
  - d. Stage and grade of cancer
  - e. Setting, provider characteristics, equipment, quality assurance standards and procedures
- KQ4. What is the evidence of cost and cost-effectiveness of SBRT?



## **Analytic Framework**

#### Figure 1. Analytic Framework



# **Detailed Inclusion and Exclusion Criteria**

Study Component	Inclusion	Exclusion
Populations	<ul> <li>Adults and children with non-CNS and non-NSCLC (inoperable, stage 1) malignancies where treatment by radiation therapy is appropriate</li> </ul>	<ul> <li>Studies in people with noncancer conditions (e.g., trigeminal neuralgia)</li> </ul>
Interventions	• SBRT, with devices such as Gamma Knife, CyberKnife, TomoTherapy, delivered in 10 or fewer fractions	<ul> <li>Treatments delivered in 11 or more fractions</li> <li>Interventions used for treatment planning or treatment delivery assessment only</li> </ul>
Comparators	<ul> <li>Conventional (conformal) external beam radiation therapy (EBRT)</li> <li>Other forms of radiation (e.g., brachytherapy)</li> <li>Chemotherapy</li> <li>Surgery</li> <li>No treatment</li> </ul>	• Comparators other than those stated
Outcomes	<ul> <li>Effectiveness         <ul> <li>Survival rate</li> <li>Duration of symptom-free remission</li> <li>Quality of life</li> </ul> </li> <li>Harms, including radiation exposure and complications</li> <li>Cost</li> <li>Cost-effectiveness</li> </ul>	<ul> <li>Studies that do not report outcomes of interest</li> <li>Data for treatment planning (e.g., dosing) or treatment delivery (e.g., accuracy)</li> <li>Economic outcomes from studies performed in non-US countries</li> <li>Economic outcomes from studies performed in the US that were published more than 5 years ago</li> </ul>
Timing Setting	<ul> <li>Any point in the treatment pathway</li> <li>Any outpatient or inpatient clinical setting in countries categorized as very high on the UN Human Development Index</li> </ul>	<ul> <li>None stated</li> <li>Emergency use settings</li> <li>Nonclinical settings (e.g., studies in healthy volunteers, animal models of disease)</li> <li>Countries categorized other than very high on the UN Human Development Index</li> </ul>

Study Component	Inclusion	Exclusion
Study Design	<ul> <li>For KQ1, KQ2, and KQ3         <ul> <li>Comparative study designs (prospective, retrospective, and randomized or controlled clinical trials)</li> </ul> </li> <li>For KQ2         <ul> <li>Comparative study designs</li> <li>Noncomparative study designs (≥ 100 participants)</li> </ul> </li> <li>For KQ4         <ul> <li>Comparative cost data and relevant economic evaluations</li> <li>Cost-effectiveness analyses</li> <li>Economic simulation modeling studies</li> </ul> </li> </ul>	<ul> <li>Abstracts, conference proceedings, posters, editorials, letters</li> <li>Studies without a comparator (unless for harms only)</li> <li>Proof-of-principle studies (e.g., technology development or technique modification)</li> <li>Studies without extractable data</li> </ul>
Sample Size	<ul> <li>Minimum sample size of 50 participants for comparative study designs</li> <li>Minimum sample size of 100 participants for noncomparative study designs</li> </ul>	• Studies that do not meet the minimum sample size
Publication	<ul> <li>Published, peer-reviewed, English- language articles</li> </ul>	<ul> <li>Studies reported only as abstracts that do not allow study characteristics to be determined</li> <li>Studies that cannot be located</li> <li>Duplicate publications of the same study that do not report different outcomes or follow-up times, or single site reports from published multicenter studies</li> <li>Studies published in languages other than English</li> </ul>

Abbreviations. CNS: central nervous system; KQ: key question; NSCLC: non-small cell lung cancer; SBRT: stereotactic body radiation therapy; UN: United Nations.

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