

BEYİN METASTAZLARINDA STEREOTAKTİK RADYOTERAPİ

Ela DELİKGÖZ SOYKUT¹

GİRİŞ

Beyin metastazları santral sinir sisteminde en sık görülen tümörlerdir. Son yıllarda primer hastalığın ve ekstrakranial metastazların etkin bir şekilde tedavi edilmesiyle uzayan sağkalım sürelerine bağlı olarak beyin metastazı gelişme sıklığı daha artmıştır. Beyin metastazlarının tedavisinde radyoterapi önemli rol almaktadır. Düşük performans durumu nedeniyle cerrahi tedavinin mümkün olmaması, kemoterapi ajanlarının sıklıkla kan beyin bariyerini geçemeyip etki edememesi, immünoterapi ajanları ile yapılmış çalışmaların henüz çok yeni olması sebebiyle beyin metastazı tedavisinde etkinlik açısından en büyük paya sahip olan radyoterapi halen yerini korumaktadır.

Klasik olarak beyin metastazı tedavisinde tüm beyin radyoterapisi (TBRT) uygulanmaktadır. Onkolojik tedavilerde amaç metastatik tümörlerde de öncelikle sağkalım sürelerini uzatabilmek ve/veya palyasyon sağlayıp yaşam kalitesini artırıbmaktır. Bu bağlamda stereotaktik radyoterapi (SRT) uygulamaları giderek artan sıklıkta kullanılarak beyin metastazı tedavi yönetiminde yer bulmuştur.

Stereotaktik radyoterapi, hedef dokuya yüksek doz radyasyon verilirken, çevre dokularda hızlı doz düşüşü sağlanarak sağlıklı dokuların radyasyondan en az şekilde etkilendiği radyoterapi yöntemidir. Günümüzde, radyoterapi planlama ve uygulamasındaki teknolojik gelişmeler ışığında SRT'nin uygulanıldığı çok çeşitli teknik ve tedavi cihazı mevcuttur. Tüm teknikler ile görüntü kılavuzluğunda, immobilizasyon gereçleri kullanılarak tedavi güvenle uygulanabilmektedir. Yüksek doz radyasyon tek fraksiyonda verildiğinde stereotaktik radyocerrahi (SRC), 2-5 fraksiyonda verildiğinde fraksiyonel stereotaktik radyoterapi (FSRT) tanımı kullanılır. Tedavinin tek günde veya fraksiyonel verilip verilmemesi kararı klinisyenin

¹ Uzman Doktor, SBÜ Samsun Eğitim ve Araştırma Hastanesi, eladelikgoz@gmail.com

KAYNAKLAR

- Andrews DW, Scott CB, Sperduto PW, et al. (2004). Whole brain radiation therapy with or without stereotactic radiosurgery boost for patients with one to three brain metastases: phase III results of the RTOG 9508 randomised trial. *Lancet*, 363 (9422), 1665-1672.
- Aoyama H, Shirato H, Tago M, et al. (2006). Stereotactic radiosurgery plus whole-brain radiation therapy vs stereotactic radiosurgery alone for treatment of brain metastases: a randomized controlled trial. *JAMA*, 295 (21), 2483-2491.
- Aoyama H, Tag M, Shirato H. (2015). Stereotactic Radiosurgery With or Without Whole-Brain Radiotherapy for Brain Metastases: Secondary Analysis of the JROSG 99-1 Randomized Clinical Trial. Japanese Radiation Oncology Study Group 99-1 (JROSG 99-1) Investigators. *JAMA Oncol*, 1 (4), 457-464. doi: 10.1001/jamaoncol.2015.1145.
- Brown PD, Pugh S, Laack NN, et al. (2013). Memantine for the prevention of cognitive dysfunction in patients receiving whole-brain radiotherapy: a randomized, double-blind, placebo-controlled trial. *Neuro Oncol*, 15 (10), 1429-1437. doi: 10.1093/neuonc/not114.
- Brown PD, Ballman KV, Cerhan JH, et al. (2017). Postoperative stereotactic radiosurgery compared with whole brain radiotherapy for resected metastatic brain disease (NCCTG N107C/CEC-3): a multicentre, randomised, controlled, phase 3 trial. *Lancet Oncol*, 18 (8), 1049-1060. doi: 10.1016/S1470-2045(17)30441-2. Epub 2017 Jul 4.
- Brown PD, Jaekle K, Ballman KV, et al. (2016). Effect of Radiosurgery Alone vs Radiosurgery With Whole Brain Radiation Therapy on Cognitive Function in Patients With 1 to 3 Brain Metastases: A Randomized Clinical Trial. *JAMA*, 316 (4), 401-409. doi: 10.1001/jama.2016.9839.
- Chang EL, Wefel JS, Hess KR, et al. (2009). Neurocognition in patients with brain metastases treated with radiosurgery or radiosurgery pluswhole-brain irradiation: a randomised controlled trial. *Lancet Oncol*, 10 (11), 1037-1044. doi: 10.1016/S1470-2045(09)70263-3. Epub 2009 Oct 2.
- Churilla TM, Handorf E, Collette S, et al. (2017). Whole brain radiotherapy after stereotactic radiosurgery or surgical resection among patients with one to three brain metastases and favorable prognoses: a secondary analysis of EORTC 22952-26001. *Ann Oncol*, 28 (10), 2588-2594. doi: 10.1093/annonc/mdx332.
- Churilla TM, Chowdhury IH, Handorf E, et al. (2019). Comparison of Local Control of Brain Metastases With Stereotactic Radiosurgery vs Surgical Resection: A Secondary Analysis of a Randomized Clinical Trial. *JAMA Oncol* 5 (2), 243-247. doi: 10.1001/jamaoncol.2018.4610.
- ClinicalTrials (2019). Memantine hydrochloride and whole-brain radiotherapy with or without hippocampal avoidance in reducing neurocognitive decline in patients with brain metastases. (30/08/2019 tarihinde <https://clinicaltrials.gov/ct2/show/study/NCT02360215> adresinden ulaşılmıştır).
- Grabber JJ, Cobbs CS, Olson JJ. (2019). Congress of Neurological Surgeons Systematic Review and Evidence-Based Guidelines on the Use of Stereotactic Radiosurgery in the Treatment of Adults With Metastatic Brain Tumors. *Neurosurgery*, 84 (3), E168-E170. doi: 10.1093/neurology/nyy543.
- Gondi V, Pugh SL, Tome WA, et al. (2014). Preservation of memory with conformal avoidance of the hippocampal neural stem-cell compartment during whole-brain radiotherapy for brain metastases (RTOG 0933): a phase II multi-institutional trial. *J Clin Oncol*, 32 (34), 3810-3816. doi: 10.1200/JCO.2
- Kocher M, Soffietti R, Abacioglu U, et al. (2011). Adjuvant whole-brain radiotherapy versus observation after radiosurgery or surgical resection of one to three cerebral metastases: results of the EORTC 22952-26001 study. *J Clin Oncol*, 29 (2), 134-141. doi: 10.1200/JCO.2010.30.1655.
- Kondziolka D, Patel A, Lunsford LD, et al. (1999). Stereotactic radiosurgery plus whole brain radiotherapy versus radiotherapy alone for patients with multiple brain metastases. *Int J Radiat Oncol Biol Phys*, 45 (2), 427-434.
- Mahajan A, Ahmed S, McAleer MF, et al. (2017). Post-operative stereotactic radiosurgery ver-

- sus observation for completely resected brain metastases: a single-centre, randomised, controlled, phase 3 trial. *Lancet Oncol*, 18 (8), 1040-1048. doi: 10.1016/S1470-2045(17)30414-X. Epub 2017 Jul 4.
- 16. Mulvenna P, Nankivell M, Barton R, et al. (2016). Dexamethasone and supportive care with or without whole brain radiotherapy in treating patients with non-small cell lung cancer with brain metastases unsuitable for resection or stereotactic radiotherapy (QUARTZ): results from a phase 3, non-inferiority, randomised trial. *Lancet*, 388 (10055), 2004-2014. doi: 10.1016/S0140-6736(16)30825-X.
 - 17. NCCN guidelines (2019). Central nervous system cancers. Limited and extensive brain metastases. (30/08/2019 tarihinde https://www.nccn.org/professionals/physician_gls/pdf/cns_blocks.pdf adresinden ulaşılmıştır).
 - 18. Rusthoven CG, Doebele RC. (2016). Management of Brain Metastases in ALK-Positive Non-Small-Cell Lung Cancer. *J Clin Oncol*, 34 (24), 2814-2819. doi: 10.1200/JCO.2016.67.2410.
 - 19. Sahgal A, Aoyama H, Kocher M, et al. (2015). Phase 3 trials of stereotactic radiosurgery with or without whole-brain radiation therapy for 1 to 4 brain metastases: individual patient data meta-analysis. *Int J Radiat Oncol Biol Phys*, 91 (4), 710-717. doi: 10.1016/j.ijrobp.2014.10.024.
 - 20. Shaw E, Scott C, Souhami L, et al. (1996). Radiosurgery for the treatment of previously irradiated recurrent primary brain tumors and brain metastases: initial report of radiation therapy oncology group protocol (90-05). *Int J Radiat Oncol Biol Phys*, 34 (3), 647-654.
 - 21. Sneed PK, Lamborn KR, Forstner JM, et al. (1999). Radiosurgery for brain metastases: is whole brain radiotherapy necessary? *Int J Radiat Oncol Biol Phys*, 43 (3), 549-558.
 - 22. Soltys SG, Adler JR, Lipani JD, et al. (2008). Stereotactic radiosurgery of the postoperative resection cavity for brain metastases. *Int J Radiat Oncol Biol Phys*, 70 (1), 187-193.
 - 23. Sperduto PW, Shanley R, Luo X, et al. (2014). Secondary analysis of RTOG 9508, a phase 3 randomized trial of whole-brain radiation therapy versus WBRT plus stereotactic radiosurgery in patients with 1-3 brain metastases; poststratified by the graded prognostic assessment (GPA). *Int J Radiat Oncol Biol Phys*, 90 (3), 526-531. doi: 10.1016/j.ijrobp.2014.07.002.
 - 24. Yamamoto M, Serizawa T, Shuto T, et al. (2014). Stereotactic radiosurgery for patients with multiple brain metastases (JLGK0901): a multi-institutional prospective observational study. *Lancet Oncol*, 15 (4), 387-395. doi: 10.1016/S1470-2045(14)70061-0. Epub 2014 Mar 10.