

# Boron neutron capture therapy

Boron neutron capture therapy is based on a [nuclear reaction](#) between the nonradioactive isotope boron-10 and either low-energy thermal neutrons or high-energy epithermal neutrons, which generate high linear energy transfer  $\alpha$  particles and a recoiled lithium nucleus ( $^7\text{Li}$ ) that selectively destroys the DNA helix in tumor cells. Boron neutron capture therapy is an emerging procedure aimed at improving the therapeutic ratio for the traditional treatment of various malignancies, which has been studied clinically in a variety of diseases, including glioblastoma, head and neck cancer, cutaneous melanoma, hepatocellular carcinoma, lung cancer, and extramammary Paget's disease. However, boron neutron capture therapy has not been clinically performed for urological cancers, excluding genital extramammary Paget's disease that appeared at the scrotum to penis area. In this review, we aimed to provide an updated summary of the current clinical literature of patients treated with boron neutron capture therapy and to focus on the future prospects of boron neutron capture therapy for urological cancers <sup>1)</sup>.

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Boron Neutron capture therapy (NCT) is a noninvasive therapeutic modality for treating locally invasive malignant tumors such as primary brain tumors and recurrent head and neck cancer.

It is a two step procedure: first, the patient is injected with a tumor localizing drug containing a non-radioactive isotope that has a high propensity or cross section ( $\sigma$ ) to capture slow neutrons. The cross section of the capture agent is many times greater than that of the other elements present in tissues such as hydrogen, oxygen, and nitrogen. In the second step, the patient is radiated with epithermal neutrons, which after losing energy as they penetrate tissue, are absorbed by the capture agent which subsequently emits high-energy charged particles, thereby resulting in a biologically destructive nuclear reaction.

All of the clinical experience to date with NCT is with the non-radioactive isotope boron-10, and this is known as boron neutron capture therapy (BNCT).

A boron delivery system with high therapeutic efficiency and low adverse effects is crucial for a successful boron neutron capture therapy (BNCT).

Boron cluster-containing redox nanoparticles (BNPs) are promising for enhancing the BNCT performance <sup>2)</sup>.

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The clinical results of BNCT in patients with [Glioblastoma](#) are similar to those of recent conventional treatments based on radiotherapy with concomitant and adjuvant temozolomide <sup>3)</sup>.

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Of the 180 patients with malignant brain tumors whom we treated with boron neutron capture therapy (BNCT) since 1968, only one (0.56%) developed multiple radiation-induced meningiomas. The parasagittal meningioma that had received 42Gy (w) for BNCT showed more rapid growth on Gd-enhanced MRI scans and more atypical features on histopathologic studies than the temporal convexity tumor that had received 20Gy (w). Long-term follow up MRI studies are necessary in long-survivors of malignant brain tumors treated by BNCT <sup>4)</sup>.

- 1: Takahara K, Miyatake SI, Azuma H, Shiroki R. Boron neutron capture therapy for urological cancers. *Int J Urol.* 2022 Mar 3. doi: 10.1111/iju.14855. Epub ahead of print. PMID: 35240726.
- 2: Kong Z, Li Z, Chen J, Liu S, Liu D, Li J, Li N, Ma W, Feng F, Wang Y, Yang Z, Liu Z. Metabolic characteristics of [<sup>18</sup>F]fluoroboronotyrosine (FBY) PET in malignant brain tumors. *Nucl Med Biol.* 2022 Jan 10;106-107:80-87. doi: 10.1016/j.nucmedbio.2022.01.002. Epub ahead of print. PMID: 35091195.
- 3: Hattori Y, Ishimura M, Ohta Y, Takenaka H, Kawabata S, Kirihata M. Dodecaborate Conjugates Targeting Tumor Cell Overexpressing Translocator Protein for Boron Neutron Capture Therapy. *ACS Med Chem Lett.* 2021 Dec 1;13(1):50-54. doi: 10.1021/acsmmedchemlett.1c00377. PMID: 35059123; PMCID: PMC8762747.
- 4: Kanygin V, Kichigin A, Zaboronok A, Kasatova A, Petrova E, Tsygankova A, Zavjalov E, Mathis BJ, Taskaev S. In Vivo Accelerator-Based Boron Neutron Capture Therapy for Spontaneous Tumors in Large Animals: Case Series. *Biology (Basel).* 2022 Jan 14;11(1):138. doi: 10.3390/biology11010138. PMID: 35053138; PMCID: PMC8773183.
- 5: Shirakawa M, Zaboronok A, Nakai K, Sato Y, Kayaki S, Sakai T, Tsurubuchi T, Yoshida F, Nishiyama T, Suzuki M, Tomida H, Matsumura A. A Novel Boron Lipid to Modify Liposomal Surfaces for Boron Neutron Capture Therapy. *Cells.* 2021 Dec 5;10(12):3421. doi: 10.3390/cells10123421. PMID: 34943929; PMCID: PMC8699917.
- 6: Yoshimura K, Kawabata S, Kashiwagi H, Fukuo Y, Takeuchi K, Futamura G, Hiramatsu R, Takata T, Tanaka H, Watanabe T, Suzuki M, Hu N, Miyatake SI, Wanibuchi M. Efficacy of Boron Neutron Capture Therapy in Primary Central Nervous System Lymphoma: In Vitro and In Vivo Evaluation. *Cells.* 2021 Dec 2;10(12):3398. doi: 10.3390/cells10123398. PMID: 34943904; PMCID: PMC8699713.
- 7: Kanygin V, Razumov I, Zaboronok A, Zavjalov E, Kichigin A, Solovieva O, Tsygankova A, Guselnikova T, Kasatov D, Sycheva T, Mathis BJ, Taskaev S. Dose- Dependent Suppression of Human Glioblastoma Xenograft Growth by Accelerator- Based Boron Neutron Capture Therapy with Simultaneous Use of Two Boron- Containing Compounds. *Biology (Basel).* 2021 Nov 2;10(11):1124. doi: 10.3390/biology10111124. PMID: 34827117; PMCID: PMC8615214.
- 8: Sasaki A, Tanaka H, Takata T, Tamari Y, Watanabe T, Hu N, Kawabata S, Kudo Y, Mitsumoto T, Sakurai Y, Suzuki M. Development of an irradiation method for superficial tumours using a hydrogel bolus in an accelerator-based BNCT. *Biomed Phys Eng Express.* 2021 Dec 7;8(1). doi: 10.1088/2057-1976/ac3d73. PMID: 34823226.
- 9: Kashiwagi H, Kawabata S, Yoshimura K, Fukuo Y, Kanemitsu T, Takeuchi K, Hiramatsu R, Nishimura K, Kawai K, Takata T, Tanaka H, Watanabe T, Suzuki M, Miyatake SI, Nakamura H, Wanibuchi M. Boron neutron capture therapy using dodecaborated albumin conjugates with maleimide is effective in a rat glioma model. *Invest New Drugs.* 2021 Nov 24. doi: 10.1007/s10637-021-01201-7. Epub ahead of print. PMID: 34816337.
- 10: Nuez-Martínez M, Pedrosa L, Martínez-Rovira I, Yousef I, Diao D, Teixidor F, Stanzani E, Martínez-Soler F, Tortosa A, Sierra À, Gonzalez JJ, Viñas C. Synchrotron-Based Fourier-Transform Infrared Micro-Spectroscopy (SR-FTIRM) Fingerprint of the Small Anionic Molecule Cobaltabis(dicarbollide) Uptake in Glioma Stem Cells. *Int J Mol Sci.* 2021 Sep 14;22(18):9937. doi: 10.3390/ijms22189937. PMID: 34576098; PMCID: PMC8466526.

- 11: Zaboronok A, Taskaev S, Volkova O, Mechetina L, Kasatova A, Sycheva T, Nakai K, Kasatov D, Makarov A, Kolesnikov I, Shchudlo I, Bykov T, Sokolova E, Koshkarev A, Kanygin V, Kichigin A, Mathis BJ, Ishikawa E, Matsumura A. Gold Nanoparticles Permit In Situ Absorbed Dose Evaluation in Boron Neutron Capture Therapy for Malignant Tumors. *Pharmaceutics*. 2021 Sep 16;13(9):1490. doi: 10.3390/pharmaceutics13091490. PMID: 34575566; PMCID: PMC8466622.
- 12: Kawabata S, Suzuki M, Hirose K, Tanaka H, Kato T, Goto H, Narita Y, Miyatake SI. Accelerator-based BNCT for patients with Glioblastoma recurrence: a multicenter phase II study. *Neurooncol Adv*. 2021 May 20;3(1):vdab067. doi: 10.1093/noajnl/vdab067. PMID: 34151269; PMCID: PMC8209606.
- 13: Takai S, Wanibuchi M, Kawabata S, Takeuchi K, Sakurai Y, Suzuki M, Ono K, Miyatake SI. Reactor-based boron neutron capture therapy for 44 cases of recurrent and refractory high-grade meningiomas with long-term follow-up. *Neuro Oncol*. 2022 Jan 5;24(1):90-98. doi: 10.1093/neuonc/noab108. PMID: 33984146; PMCID: PMC8730746.
- 14: Chen YW, Lee YY, Lin CF, Pan PS, Chen JK, Wang CW, Hsu SM, Kuo YC, Lan TL, Hsu SPC, Liang ML, Chen RH, Chang FC, Wu CC, Lin SC, Liang HK, Lee JC, Chen SK, Liu HM, Peir JJ, Lin KH, Huang WS, Chen KH, Kang YM, Liou SC, Wang CC, Pai PC, Li CW, Chiek DQS, Wong TT, Chiou SH, Chao Y, Tanaka H, Chou FI, Ono K. Salvage Boron Neutron Capture Therapy for Malignant Brain Tumor Patients in Compliance with Emergency and Compassionate Use: Evaluation of 34 Cases in Taiwan. *Biology (Basel)*. 2021 Apr 15;10(4):334. doi: 10.3390/biology10040334. PMID: 33920984; PMCID: PMC8071294.
- 15: Li Z, Kong Z, Chen J, Li J, Li N, Yang Z, Wang Y, Liu Z. <sup>18</sup>F-Boramino acid PET/CT in healthy volunteers and glioma patients. *Eur J Nucl Med Mol Imaging*. 2021 Sep;48(10):3113-3121. doi: 10.1007/s00259-021-05212-7. Epub 2021 Feb 15. PMID: 33590273.
- 16: Aydin HE, Gunduz MK, Kizmazoglu C, Kandemir T, Arslantas A. Cytotoxic Effect of Boron Application on Glioblastoma Cells. *Turk Neurosurg*. 2021;31(2):206-210. doi: 10.5137/1019-5149.JTN.30316-20.1. PMID: 33372254.
- 17: Fukuo Y, Hattori Y, Kawabata S, Kashiwagi H, Kanemitsu T, Takeuchi K, Futamura G, Hiramatsu R, Watanabe T, Hu N, Takata T, Tanaka H, Suzuki M, Miyatake SI, Kirihata M, Wanibuchi M. The Therapeutic Effects of Dodecaborate Containing Boronophenylalanine for Boron Neutron Capture Therapy in a Rat Brain Tumor Model. *Biology (Basel)*. 2020 Dec 1;9(12):437. doi: 10.3390/biology9120437. PMID: 33271972; PMCID: PMC7759915.
- 18: Shirakawa M, Nakai K, Sato Y, Nakamura S, Harada M, Ishihara K, Yoshida F, Matsumura A, Tomida H. Optimization of preparation methods for high loading content and high encapsulation efficiency of BSH into liposomes. *Appl Radiat Isot*. 2021 Mar;169:109260. doi: 10.1016/j.apradiso.2020.109260. Epub 2020 Jul 1. PMID: 33160809.
- 19: Kondo N, Hikida M, Nakada M, Sakurai Y, Hirata E, Takeno S, Suzuki M. Glioma Stem-Like Cells Can Be Targeted in Boron Neutron Capture Therapy with Boronophenylalanine. *Cancers (Basel)*. 2020 Oct 19;12(10):3040. doi: 10.3390/cancers12103040. PMID: 33086625; PMCID: PMC7603373.
- 20: Kanygin V, Zaboronok A, Taskaeva I, Zavjalov E, Mukhamadiyarov R, Kichigin A, Kasatova A, Razumov I, Sibirtsev R, Mathis BJ. In Vitro and In Vivo Evaluation of Fluorescently Labeled Borocaptate-Containing Liposomes. *J Fluoresc*. 2021 Jan;31(1):73-83. doi: 10.1007/s10895-020-02637-5. Epub 2020 Oct 19. PMID: 33078252.
- 21: Kamano S, Matsuyama M, Minamimura K. A Case of Diffuse Astrocytoma with 32-year Survival

- after Boron Neutron Capture Therapy. *NMC Case Rep J.* 2020 Sep 14;7(4):211-215. doi: 10.2176/nmccrj.cr.2019-0228. PMID: 33062571; PMCID: PMC7538456.
- 22: Nakai K, Endo K, Yoshida F, Koka M, Yamada N, Satoh T, Tsurubuchi T, Matsumura A, Matsumoto Y, Sakurai H. Boron analysis and imaging of cells with 2-hr BPA exposure by using micro-proton particle-induced gamma-ray emission (PIGE). *Appl Radiat Isot.* 2020 Nov;165:109334. doi: 10.1016/j.apradiso.2020.109334. Epub 2020 Jul 23. PMID: 32739796.
- 23: Miyatake SI, Wanibuchi M, Hu N, Ono K. Boron neutron capture therapy for malignant brain tumors. *J Neurooncol.* 2020 Aug;149(1):1-11. doi: 10.1007/s11060-020-03586-6. Epub 2020 Jul 16. PMID: 32676954.
- 24: Takeuchi K, Hattori Y, Kawabata S, Futamura G, Hiramatsu R, Wanibuchi M, Tanaka H, Masunaga SI, Ono K, Miyatake SI, Kirihata M. Synthesis and Evaluation of Dodecaboranethiol Containing Kojic Acid (KA-BSH) as a Novel Agent for Boron Neutron Capture Therapy. *Cells.* 2020 Jun 25;9(6):1551. doi: 10.3390/cells9061551. PMID: 32630612; PMCID: PMC7349888.
- 25: Yoshida F, Kurita T, Endo K, Nakai K, Shirakawa M, Zaboronok A, Tsurubuchi T, Ishikawa E, Matsumura A. Difference in BPA uptake between glioma stem-like cells and their cancerous cells. *Appl Radiat Isot.* 2020 Oct;164:109234. doi: 10.1016/j.apradiso.2020.109234. Epub 2020 Jun 3. PMID: 32554123.
- 26: Nakahara Y, Ito H, Masuoka J, Abe T. Boron Neutron Capture Therapy and Photodynamic Therapy for High-Grade Meningiomas. *Cancers (Basel).* 2020 May 23;12(5):1334. doi: 10.3390/cancers12051334. PMID: 32456178; PMCID: PMC7281755.
- 27: Tsurubuchi T, Shirakawa M, Kurosawa W, Matsumoto K, Ubagai R, Umishio H, Suga Y, Yamazaki J, Arakawa A, Maruyama Y, Seki T, Shibui Y, Yoshida F, Zaboronok A, Suzuki M, Sakurai Y, Tanaka H, Nakai K, Ishikawa E, Matsumura A. Evaluation of a Novel Boron-Containing  $\alpha$ -D-Mannopyranoside for BNCT. *Cells.* 2020 May 21;9(5):1277. doi: 10.3390/cells9051277. PMID: 32455737; PMCID: PMC7290312.
- 28: Chio CM, Huang YC, Chou FC, Hsu FC, Lai YB, Yu CS. Boron Accumulation in Brain Tumor Cells through Boc-Protected Tryptophan as a Carrier for Boron Neutron Capture Therapy. *ACS Med Chem Lett.* 2020 Mar 16;11(4):589-596. doi: 10.1021/acsmmedchemlett.0c00064. PMID: 32292568; PMCID: PMC7153283.
- 29: Altinoz MA, Topcu G, Elmaci İ. Boron's neurophysiological effects and tumoricidal activity on glioblastoma cells with implications for clinical treatment. *Int J Neurosci.* 2019 Oct;129(10):963-977. doi: 10.1080/00207454.2019.1595618. Epub 2019 Apr 2. PMID: 30885023.
- 30: Kanemitsu T, Kawabata S, Fukumura M, Futamura G, Hiramatsu R, Nonoguchi N, Nakagawa F, Takata T, Tanaka H, Suzuki M, Masunaga SI, Ono K, Miyatake SI, Nakamura H, Kuroiwa T. Folate receptor-targeted novel boron compound for boron neutron capture therapy on F98 glioma-bearing rats. *Radiat Environ Biophys.* 2019 Mar;58(1):59-67. doi: 10.1007/s00411-018-0765-2. Epub 2018 Nov 24. PMID: 30474719.
- 31: Shiba H, Takeuchi K, Hiramatsu R, Furuse M, Nonoguchi N, Kawabata S, Kuroiwa T, Kondo N, Sakurai Y, Suzuki M, Ono K, Oue S, Ishikawa E, Michiue H, Miyatake SI. Boron Neutron Capture Therapy Combined with Early Successive Bevacizumab Treatments for Recurrent Malignant Gliomas - A Pilot Study. *Neurol Med Chir (Tokyo).* 2018 Dec 15;58(12):487-494. doi: 10.2176/nmc.oa.2018-0111. Epub 2018 Nov 21. PMID: 30464150; PMCID: PMC6300692.

- 32: Takeuchi K, Kawabata S, Hiramatsu R, Matsushita Y, Tanaka H, Sakurai Y, Suzuki M, Ono K, Miyatake SI, Kuroiwa T. Boron Neutron Capture Therapy for High- Grade Skull-Base Meningioma. *J Neurol Surg B Skull Base.* 2018 Oct;79(Suppl 4):S322-S327. doi: 10.1055/s-0038-1666837. Epub 2018 Jul 3. PMID: 30210985; PMCID: PMC6133692.
- 33: Barth RF, Mi P, Yang W. Boron delivery agents for neutron capture therapy of cancer. *Cancer Commun (Lond).* 2018 Jun 19;38(1):35. doi: 10.1186/s40880-018-0299-7. PMID: 29914561; PMCID: PMC6006782.
- 34: Zelenetskii AN, Uspenskii S, Zaboronok A, Cherkaev G, Shchegolihin A, Mathis BJ, Selyanin M, Yamamoto T, Matsumura A. Polycomplexes of Hyaluronic Acid and Borates in a Solid State and Solution: Synthesis, Characterization and Perspectives of Application in Boron Neutron Capture Therapy. *Polymers (Basel).* 2018 Feb 13;10(2):181. doi: 10.3390/polym10020181. PMID: 30966217; PMCID: PMC6415177.
- 35: Sato E, Zaboronok A, Yamamoto T, Nakai K, Taskaev S, Volkova O, Mechetina L, Taranin A, Kanygin V, Isobe T, Mathis BJ, Matsumura A. Radiobiological response of U251MG, CHO-K1 and V79 cell lines to accelerator-based boron neutron capture therapy. *J Radiat Res.* 2018 Mar 1;59(2):101-107. doi: 10.1093/jrr/rrx071. PMID: 29281044; PMCID: PMC5950924.
- 36: Yeh CN, Chang CW, Chung YH, Tien SW, Chen YR, Chen TW, Huang YC, Wang HE, Chou YC, Chen MH, Chiang KC, Huang WS, Yu CS. Dataset on the synthesis and characterization of boron fenbufen and its F-18 labeled homolog. *Data Brief.* 2017 Sep 5;15:174-202. doi: 10.1016/j.dib.2017.08.048. PMID: 29021997; PMCID: PMC5633353.
- 37: Yeh CN, Chang CW, Chung YH, Tien SW, Chen YR, Chen TW, Huang YC, Wang HE, Chou YC, Chen MH, Chiang KC, Huang WS, Yu CS. Synthesis and characterization of boron fenbufen and its F-18 labeled homolog for boron neutron capture therapy of COX-2 overexpressed cholangiocarcinoma. *Eur J Pharm Sci.* 2017 Sep 30;107:217-229. doi: 10.1016/j.ejps.2017.07.019. Epub 2017 Jul 17. PMID: 28728977.
- 38: Kondo N, Barth RF, Miyatake SI, Kawabata S, Suzuki M, Ono K, Lehman NL. Cerebrospinal fluid dissemination of high-grade gliomas following boron neutron capture therapy occurs more frequently in the small cell subtype of IDH1<sup>R132H</sup> mutation-negative glioblastoma. *J Neurooncol.* 2017 May;133(1):107-118. doi: 10.1007/s11060-017-2408-x. Epub 2017 May 22. PMID: 28534152; PMCID: PMC5786264.
- 39: Futamura G, Kawabata S, Nonoguchi N, Hiramatsu R, Toho T, Tanaka H, Masunaga SI, Hattori Y, Kirihata M, Ono K, Kuroiwa T, Miyatake SI. Evaluation of a novel sodium borocaptate-containing unnatural amino acid as a boron delivery agent for neutron capture therapy of the F98 rat glioma. *Radiat Oncol.* 2017 Jan 23;12(1):26. doi: 10.1186/s13014-017-0765-4. PMID: 28114947; PMCID: PMC5260095.
- 40: Ishikawa M, Yamamoto T, Matsumura A, Hiratsuka J, Miyatake S, Kato I, Sakurai Y, Kumada H, Shrestha SJ, Ono K. Early clinical experience utilizing scintillator with optical fiber (SOF) detector in clinical boron neutron capture therapy: its issues and solutions. *Radiat Oncol.* 2016 Aug 9;11(1):105. doi: 10.1186/s13014-016-0680-0. PMID: 27506665; PMCID: PMC4977859.
- 41: Gao Z, Horiguchi Y, Nakai K, Matsumura A, Suzuki M, Ono K, Nagasaki Y. Use of boron cluster-containing redox nanoparticles with ROS scavenging ability in boron neutron capture therapy to achieve high therapeutic efficiency and low adverse effects. *Biomaterials.* 2016 Oct;104:201-12. doi: 10.1016/j.biomaterials.2016.06.046. Epub 2016 Jul 11. PMID: 27467416.

- 42: Sun T, Li Y, Huang Y, Zhang Z, Yang W, Du Z, Zhou Y. Targeting glioma stem cells enhances anti-tumor effect of boron neutron capture therapy. *Oncotarget.* 2016 Jul 12;7(28):43095-43108. doi: 10.18632/oncotarget.9355. PMID: 27191269; PMCID: PMC5190011.
- 43: Kondo N, Sakurai Y, Hirota Y, Tanaka H, Watanabe T, Nakagawa Y, Narabayashi M, Kinashi Y, Miyatake S, Hasegawa M, Suzuki M, Masunaga S, Ohnishi T, Ono K. DNA damage induced by boron neutron capture therapy is partially repaired by DNA ligase IV. *Radiat Environ Biophys.* 2016 Mar;55(1):89-94. doi: 10.1007/s00411-015-0625-2. Epub 2015 Nov 16. PMID: 26573366.
- 44: Takahara K, Inamoto T, Minami K, Yoshikawa Y, Takai T, Ibuki N, Hirano H, Nomi H, Kawabata S, Kiyama S, Miyatake S, Kuroiwa T, Suzuki M, Kirihata M, Azuma H. The Anti-Proliferative Effect of Boron Neutron Capture Therapy in a Prostate Cancer Xenograft Model. *PLoS One.* 2015 Sep 1;10(9):e0136981. doi: 10.1371/journal.pone.0136981. PMID: 26325195; PMCID: PMC4556531.
- 45: Zaboronok A, Yamamoto T, Nakai K, Yoshida F, Uspenskii S, Selyanin M, Zelenetskii A, Matsumura A. Hyaluronic acid as a potential boron carrier for BNCT: Preliminary evaluation. *Appl Radiat Isot.* 2015 Dec;106:181-4. doi: 10.1016/j.apradiso.2015.08.020. Epub 2015 Aug 18. PMID: 26302663.
- 46: Okamoto E, Yamamoto T, Nakai K, Fumiyo Yoshida, Matsumura A. Detection of DNA double-strand breaks in boron neutron capture reaction. *Appl Radiat Isot.* 2015 Dec;106:185-8. doi: 10.1016/j.apradiso.2015.08.019. Epub 2015 Aug 17. PMID: 26302660.
- 47: Kulvik M, Kallio M, Laakso J, Vähätilo J, Hermans R, Järvinen E, Paetau A, Rasilainen M, Ruokonen I, Seppälä M, Jääskeläinen J. Biodistribution of boron after intravenous 4-dihydroxyborylphenylalanine-fructose (BPA-F) infusion in meningioma and schwannoma patients: A feasibility study for boron neutron capture therapy. *Appl Radiat Isot.* 2015 Dec;106:207-12. doi: 10.1016/j.apradiso.2015.08.006. Epub 2015 Aug 10. PMID: 26298436.
- 48: Nakai K, Yamamoto Y, Okamoto E, Yamamoto T, Yoshida F, Matsumura A, Yamada N, Kitamura A, Koka M, Satoh T. Boron analysis for neutron capture therapy using particle-induced gamma-ray emission. *Appl Radiat Isot.* 2015 Dec;106:166-70. doi: 10.1016/j.apradiso.2015.07.035. Epub 2015 Jul 29. PMID: 26242558.
- 49: Kageji T, Sogabe S, Mizobuchi Y, Nakajima K, Shinji N, Nakagawa Y. Radiation-induced meningiomas after BNCT in patients with malignant glioma. *Appl Radiat Isot.* 2015 Dec;106:256-9. doi: 10.1016/j.apradiso.2015.06.004. Epub 2015 Jun 20. PMID: 26122975.
- 50: Hiramatsu R, Kawabata S, Tanaka H, Sakurai Y, Suzuki M, Ono K, Miyatake SI, Kuroiwa T, Hao E, Vicente MGH. Tetrakis(p-Carboranylthio- Tetrafluorophenyl)Chlorin (TPFC): Application for Photodynamic Therapy and Boron Neutron Capture Therapy. *J Pharm Sci.* 2015 Mar;104(3):962-970. doi: 10.1002/jps.24317. Epub 2016 Jan 8. PMID: 28756849.
- 51: Hiramatsu R, Kawabata S, Tanaka H, Sakurai Y, Suzuki M, Ono K, Miyatake S, Kuroiwa T, Hao E, Vicente MG. Tetrakis(p-carboranylthio- tetrafluorophenyl)chlorin (TPFC): application for photodynamic therapy and boron neutron capture therapy. *J Pharm Sci.* 2015 Mar;104(3):962-70. doi: 10.1002/jps.24317. Epub 2014 Dec 26. PMID: 25546823; PMCID: PMC4415589.
- 52: Futamura G, Kawabata S, Siba H, Kuroiwa T, Suzuki M, Kondo N, Ono K, Sakurai Y, Tanaka M, Todo T, Miyatake S. A case of radiation-induced osteosarcoma treated effectively by boron neutron capture therapy. *Radiat Oncol.* 2014 Nov 4;9:237. doi: 10.1186/s13014-014-0237-z. PMID: 25366059; PMCID: PMC4228084.

- 53: Kageji T, Nagahiro S, Mizobuchi Y, Matsuzaki K, Nakagawa Y, Kumada H. Boron neutron capture therapy (BNCT) for newly-diagnosed glioblastoma: comparison of clinical results obtained with BNCT and conventional treatment. *J Med Invest.* 2014;61(3-4):254-63. doi: 10.2152/jmi.61.254. PMID: 25264042.
- 54: Sun T, Li Y, Wu T, Xie X, Chen G, Wei Y, Li B, Zhou Y, Du Z. Comparative analysis of pathology and boronophenylalanine uptake in experimental orthotopic and heterotopic amelanotic melanoma. *Melanoma Res.* 2014 Aug;24(4):315-21. doi: 10.1097/CMR.0000000000000086. PMID: 24915302.
- 55: Kawaji H, Miyatake S, Shinmura K, Kawabata S, Tokuyama T, Namba H. Effect of boron neutron capture therapy for recurrent anaplastic meningioma: an autopsy case report. *Brain Tumor Pathol.* 2015 Jan;32(1):61-5. doi: 10.1007/s10014-014-0189-x. Epub 2014 May 8. PMID: 24807102.
- 56: Yoshida F, Yamamoto T, Nakai K, Zaboronok A, Matsuda M, Akutsu H, Ishikawa E, Shirakawa M, Matsumura A. Pretreatment with buthionine sulfoximine enhanced uptake and retention of BSH in brain tumor. *Appl Radiat Isot.* 2014 Jun;88:86-8. doi: 10.1016/j.apradiso.2014.02.025. Epub 2014 Mar 28. PMID: 24731546.
- 57: Kumada H, Matsumura A, Sakurai H, Sakae T, Yoshioka M, Kobayashi H, Matsumoto H, Kiyanagi Y, Shibata T, Nakashima H. Project for the development of the linac based NCT facility in University of Tsukuba. *Appl Radiat Isot.* 2014 Jun;88:211-5. doi: 10.1016/j.apradiso.2014.02.018. Epub 2014 Feb 22. PMID: 24637084.
- 58: Kageji T, Mizobuchi Y, Nagahiro S, Nakagawa Y, Kumada H. Correlation between radiation dose and histopathological findings in patients with glioblastoma treated with boron neutron capture therapy (BNCT). *Appl Radiat Isot.* 2014 Jun;88:20-2. doi: 10.1016/j.apradiso.2013.12.014. Epub 2013 Dec 25. PMID: 24480727.
- 59: Michiue H, Sakurai Y, Kondo N, Kitamatsu M, Bin F, Nakajima K, Hirota Y, Kawabata S, Nishiki T, Ohmori I, Tomizawa K, Miyatake S, Ono K, Matsui H. The acceleration of boron neutron capture therapy using multi-linked mercaptoundecahydrododecaborate (BSH) fused cell-penetrating peptide. *Biomaterials.* 2014 Mar;35(10):3396-405. doi: 10.1016/j.biomaterials.2013.12.055. Epub 2014 Jan 20. PMID: 24452095.
- 60: Okazaki T, Kageji T, Mizobuchi Y, Miyamoto T, Nagahiro S. Nine-year interval recurrence after treatment of boron neutron capture therapy in a patient with glioblastoma: a case report. *Appl Radiat Isot.* 2014 Jun;88:28-31. doi: 10.1016/j.apradiso.2013.12.029. Epub 2014 Jan 1. PMID: 24440540.
- 61: Miyatake S, Kawabata S, Hiramatsu R, Furuse M, Kuroiwa T, Suzuki M. Boron neutron capture therapy with bevacizumab may prolong the survival of recurrent malignant glioma patients: four cases. *Radiat Oncol.* 2014 Jan 6;9:6. doi: 10.1186/1748-717X-9-6. PMID: 24387301; PMCID: PMC3923505.
- 62: Aleynik V, Bashkirtsev A, Kanygin V, Kasatov D, Kuznetsov A, Makarov A, Schudlo I, Sorokin I, Taskaev S, Tiunov M. Current progress and future prospects of the VITA based neutron source. *Appl Radiat Isot.* 2014 Jun;88:177-9. doi: 10.1016/j.apradiso.2013.11.132. Epub 2013 Dec 11. PMID: 24369890.
- 63: Sun T, Zhang Z, Li B, Chen G, Xie X, Wei Y, Wu J, Zhou Y, Du Z. Boron neutron capture therapy induces cell cycle arrest and cell apoptosis of glioma stem/progenitor cells in vitro. *Radiat Oncol.* 2013 Aug 6;8(1):195. doi: 10.1186/1748-717X-8-195. PMID: 23915425; PMCID: PMC3751121.
- 64: Hiramatsu R, Kawabata S, Furuse M, Miyatake S, Kuroiwa T. Identification of early and distinct

glioblastoma response patterns treated by boron neutron capture therapy not predicted by standard radiographic assessment using functional diffusion map. *Radiat Oncol.* 2013 Aug 1;8(1):192. doi: 10.1186/1748-717X-8-192. PMID: 23915330; PMCID: PMC3751226.

65: Kawabata S, Hiramatsu R, Kuroiwa T, Ono K, Miyatake S. Boron neutron capture therapy for recurrent high-grade meningiomas. *J Neurosurg.* 2013 Oct;119(4):837-44. doi: 10.3171/2013.5.JNS122204. Epub 2013 Jun 28. PMID: 23808536.

66: Miyatake S, Furuse M, Kawabata S, Maruyama T, Kumabe T, Kuroiwa T, Ono K. Bevacizumab treatment of symptomatic pseudoprogression after boron neutron capture therapy for recurrent malignant gliomas. Report of 2 cases. *Neuro Oncol.* 2013 Jun;15(6):650-5. doi: 10.1093/neuonc/not020. Epub 2013 Mar 3. PMID: 23460324; PMCID: PMC3661101.

67: Dai C, Cai F, Hwang KC, Zhou Y, Zhang Z, Liu X, Ma S, Yang Y, Yao Y, Feng M, Bao X, Li G, Wei J, Jiao Y, Wei Z, Ma W, Wang R. Folate receptor-mediated boron-10 containing carbon nanoparticles as potential delivery vehicles for boron neutron capture therapy of nonfunctional pituitary neuroendocrine tumors. *Sci China Life Sci.* 2013 Feb;56(2):163-73. doi: 10.1007/s11427-012-4433-5. Epub 2013 Jan 18. PMID: 23334699.

68: Sun T, Zhou Y, Xie X, Chen G, Li B, Wei Y, Chen J, Huang Q, Du Z. Selective uptake of boronophenylalanine by glioma stem/progenitor cells. *Appl Radiat Isot.* 2012 Aug;70(8):1512-8. doi: 10.1016/j.apradiso.2012.04.005. Epub 2012 Apr 12. PMID: 22728842.

69: Aiyama H, Nakai K, Yamamoto T, Nariai T, Kumada H, Ishikawa E, Isobe T, Endo K, Takada T, Yoshida F, Shibata Y, Matsumura A. A clinical trial protocol for second line treatment of malignant brain tumors with BNCT at University of Tsukuba. *Appl Radiat Isot.* 2011 Dec;69(12):1819-22. doi: 10.1016/j.apradiso.2011.04.031. Epub 2011 Jul 20. PMID: 21778066.

70: Miyatake S. [Boron neutron capture therapy for brain glioma]. *Gan To Kagaku Ryoho.* 2011 Jun;38(6):927-32. Japanese. PMID: 21761624.

71: Kageji T, Mizobuchi Y, Nagahiro S, Nakagawa Y, Kumada H. Clinical results of boron neutron capture therapy (BNCT) for glioblastoma. *Appl Radiat Isot.* 2011 Dec;69(12):1823-5. doi: 10.1016/j.apradiso.2011.05.029. Epub 2011 Jun 12. PMID: 21684170.

72: Nakai K, Yamamoto T, Aiyama H, Takada T, Yoshida F, Kageji T, Kumada H, Isobe T, Endo K, Matsuda M, Tsurubuchi T, Shibata Y, Takano S, Mizumoto M, Tsuboi K, Matsumura A. Boron neutron capture therapy combined with fractionated photon irradiation for glioblastoma: a recursive partitioning analysis of BNCT patients. *Appl Radiat Isot.* 2011 Dec;69(12):1790-2. doi: 10.1016/j.apradiso.2011.03.049. Epub 2011 May 5. PMID: 21565517.

73: Kageji T, Mizobuchi Y, Nagahiro S, Nakagawa Y, Kumada H. Long-survivors of glioblastoma treated with boron neutron capture therapy (BNCT). *Appl Radiat Isot.* 2011 Dec;69(12):1800-2. doi: 10.1016/j.apradiso.2011.03.021. Epub 2011 Mar 21. PMID: 21463946.

74: Kawabata S, Miyatake S, Hiramatsu R, Hirota Y, Miyata S, Takekita Y, Kuroiwa T, Kirihata M, Sakurai Y, Maruhashi A, Ono K. Phase II clinical study of boron neutron capture therapy combined with X-ray radiotherapy/temozolomide in patients with newly diagnosed glioblastoma multiforme—study design and current status report. *Appl Radiat Isot.* 2011 Dec;69(12):1796-9. doi: 10.1016/j.apradiso.2011.03.014. Epub 2011 Mar 21. PMID: 21459588.

75: Matsuda M, Yamamoto T, Ishikawa E, Nakai K, Zaboronok A, Takano S, Matsumura A. Prognostic

factors in glioblastoma multiforme patients receiving high-dose particle radiotherapy or conventional radiotherapy. *Br J Radiol.* 2011 Dec;84 Spec No 1(Spec Iss 1):S54-60. doi: 10.1259/bjr/29022270. Epub 2011 Mar 22. PMID: 21427185; PMCID: PMC3473893.

76: Yamamoto T, Nakai K, Nariai T, Kumada H, Okumura T, Mizumoto M, Tsuboi K, Zaboronok A, Ishikawa E, Aiyama H, Endo K, Takada T, Yoshida F, Shibata Y, Matsumura A. The status of Tsukuba BNCT trial: BPA-based boron neutron capture therapy combined with X-ray irradiation. *Appl Radiat Isot.* 2011 Dec;69(12):1817-8. doi: 10.1016/j.apradiso.2011.02.013. Epub 2011 Feb 15. PMID: 21393005.

77: Miyata S, Kawabata S, Hiramatsu R, Doi A, Ikeda N, Yamashita T, Kuroiwa T, Kasaoka S, Maruyama K, Miyatake S. Computed tomography imaging of transferrin targeting liposomes encapsulating both boron and iodine contrast agents by convection-enhanced delivery to F98 rat glioma for boron neutron capture therapy. *Neurosurgery.* 2011 May;68(5):1380-7; discussion 1387. doi: 10.1227/NEU.0b013e31820b52aa. PMID: 21273928.

78: Hiramatsu R, Kawabata S, Miyatake S, Kuroiwa T, Easson MW, Vicente MG. Application of a novel boronated porphyrin ( $H_2OCP$ ) as a dual sensitizer for both PDT and BNCT. *Lasers Surg Med.* 2011 Jan;43(1):52-8. doi: 10.1002/lsm.21026. PMID: 21254143; PMCID: PMC3164306.

79: Wang P, Zhen H, Jiang X, Zhang W, Cheng X, Guo G, Mao X, Zhang X. Boron neutron capture therapy induces apoptosis of glioma cells through Bcl-2/Bax. *BMC Cancer.* 2010 Dec 2;10:661. doi: 10.1186/1471-2407-10-661. PMID: 21122152; PMCID: PMC3003659.

80: Kawabata S, Yang W, Barth RF, Wu G, Huo T, Binns PJ, Riley KJ, Ongayi O, Gottumukkala V, Vicente MG. Convection enhanced delivery of carboranylporphyrins for neutron capture therapy of brain tumors. *J Neurooncol.* 2011 Jun;103(2):175-85. doi: 10.1007/s11060-010-0376-5. Epub 2010 Sep 17. PMID: 20848301; PMCID: PMC3005081.

81: Furuse M, Kawabata S, Kuroiwa T, Miyatake S. Repeated treatments with bevacizumab for recurrent radiation necrosis in patients with malignant brain tumors: a report of 2 cases. *J Neurooncol.* 2011 May;102(3):471-5. doi: 10.1007/s11060-010-0333-3. Epub 2010 Aug 7. PMID: 20694573.

82: Abdalla K, Naqvi AA, Maalej N, Elshahat B. Dose calculation from a D-D- reaction-based BSA for boron neutron capture synovectomy. *Appl Radiat Isot.* 2010 Apr-May;68(4-5):751-4. doi: 10.1016/j.apradiso.2009.09.041. Epub 2009 Sep 20. PMID: 19828325.

83: Yang W, Barth RF, Wu G, Huo T, Tjarks W, Ciesielski M, Fenstermaker RA, Ross BD, Wikstrand CJ, Riley KJ, Binns PJ. Convection enhanced delivery of boronated EGF as a molecular targeting agent for neutron capture therapy of brain tumors. *J Neurooncol.* 2009 Dec;95(3):355-365. doi: 10.1007/s11060-009-9945-x. Epub 2009 Jul 9. PMID: 19588228; PMCID: PMC2830857.

84: Tsurubuchi T, Yamamoto T, Nakai K, Zaboronok A, Yoshida F, Miyakawa M, Shirakawa M, Yamamoto Y, Matsuda M, Matsumura A. Intracellular uptake of a new boronated porphyrin EC032. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S94-6. doi: 10.1016/j.apradiso.2009.03.098. Epub 2009 Mar 31. PMID: 19410469.

85: Nakagawa Y, Kageji T, Mizobuchi Y, Kumada H, Nakagawa Y. Clinical results of BNCT for malignant brain tumors in children. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S27-30. doi: 10.1016/j.apradiso.2009.03.090. Epub 2009 Mar 28. PMID: 19406652.

86: Yamamoto Y, Isobe T, Yamamoto T, Shibata Y, Anno I, Nakai K, Shirakawa M, Matsushita A, Sato E, Matsumura A. T2 corrected quantification method of L-p- boronophenylalanine using proton magnetic

resonance spectroscopy for boron neutron capture therapy. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S345-7. doi: 10.1016/j.apradiso.2009.03.060. Epub 2009 Mar 27. PMID: 19406648.

87: Kawabata S, Miyatake S, Nonoguchi N, Hiramatsu R, Iida K, Miyata S, Yokoyama K, Doi A, Kuroda Y, Kuroiwa T, Michiue H, Kumada H, Kirihata M, Imahori Y, Maruhashi A, Sakurai Y, Suzuki M, Masunaga S, Ono K. Survival benefit from boron neutron capture therapy for the newly diagnosed glioblastoma patients. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S15-8. doi: 10.1016/j.apradiso.2009.03.015. Epub 2009 Mar 25. PMID: 19398348.

88: Miyatake S, Kawabata S, Yokoyama K, Kuroiwa T, Michiue H, Sakurai Y, Kumada H, Suzuki M, Maruhashi A, Kirihata M, Ono K. Survival benefit of boron neutron capture therapy for recurrent malignant gliomas. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S22-4. doi: 10.1016/j.apradiso.2009.03.032. Epub 2009 Mar 27. PMID: 19394240.

89: Nakai K, Kumada H, Yamamoto T, Tsurubuchi T, Zaboronok A, Matsumura A. Feasibility of boron neutron capture therapy for malignant spinal tumors. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S43-6. doi: 10.1016/j.apradiso.2009.03.089. Epub 2009 Mar 28. PMID: 19376723.

90: Matsuda M, Yamamoto T, Kumada H, Nakai K, Shirakawa M, Tsurubuchi T, Matsumura A. Dose distribution and clinical response of glioblastoma treated with boron neutron capture therapy. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S19-21. doi: 10.1016/j.apradiso.2009.03.054. Epub 2009 Mar 27. PMID: 19375933.

91: Nariai T, Ishiwata K, Kimura Y, Inaji M, Momose T, Yamamoto T, Matsumura A, Ishii K, Ohno K. PET pharmacokinetic analysis to estimate boron concentration in tumor and brain as a guide to plan BNCT for malignant cerebral glioma. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S348-50. doi: 10.1016/j.apradiso.2009.03.061. Epub 2009 Mar 27. PMID: 19375930.

92: Yamamoto T, Nakai K, Tsurubuchi T, Matsuda M, Shirakawa M, Zaboronok A, Endo K, Matsumura A. Boron neutron capture therapy for newly diagnosed glioblastoma: a pilot study in Tsukuba. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S25-6. doi: 10.1016/j.apradiso.2009.03.011. Epub 2009 Mar 24. PMID: 19375927.

93: Matsumura A, Yamamoto T, Tsurubuchi T, Matsuda M, Shirakawa M, Nakai K, Endo K, Tokuee K, Tsuboi K. Current practices and future directions of therapeutic strategy in glioblastoma: survival benefit and indication of BNCT. *Appl Radiat Isot.* 2009 Jul;67(7-8 Suppl):S12-4. doi: 10.1016/j.apradiso.2009.03.010. Epub 2009 Mar 24. PMID: 19375923.

94: Miyatake S, Kawabata S, Nonoguchi N, Yokoyama K, Kuroiwa T, Matsui H, Ono K. Pseudoprogression in boron neutron capture therapy for malignant gliomas and meningiomas. *Neuro Oncol.* 2009 Aug;11(4):430-6. doi: 10.1215/15228517-2008-107. Epub 2009 Mar 16. PMID: 19289492; PMCID: PMC2743223.

95: Yamamoto T, Nakai K, Kageji T, Kumada H, Endo K, Matsuda M, Shibata Y, Matsumura A. Boron neutron capture therapy for newly diagnosed glioblastoma. *Radiother Oncol.* 2009 Apr;91(1):80-4. doi: 10.1016/j.radonc.2009.02.009. Epub 2009 Mar 11. PMID: 19285355.

96: Detta A, Cruickshank GS. L-amino acid transporter-1 and boronophenylalanine- based boron neutron capture therapy of human brain tumors. *Cancer Res.* 2009 Mar 1;69(5):2126-32. doi: 10.1158/0008-5472.CAN-08-2345. Epub 2009 Feb 24. PMID: 19244126.

97: Kawabata S, Miyatake S, Kuroiwa T, Yokoyama K, Doi A, Iida K, Miyata S, Nonoguchi N, Michiue H,

- Takahashi M, Inomata T, Imahori Y, Kirihata M, Sakurai Y, Maruhashi A, Kumada H, Ono K. Boron neutron capture therapy for newly diagnosed glioblastoma. *J Radiat Res.* 2009 Jan;50(1):51-60. doi: 10.1269/jrr.08043. Epub 2008 Oct 29. PMID: 18957828.
- 98: Miyatake S, Kawabata S, Yokoyama K, Kuroiwa T, Michiue H, Sakurai Y, Kumada H, Suzuki M, Maruhashi A, Kirihata M, Ono K. Survival benefit of Boron neutron capture therapy for recurrent malignant gliomas. *J Neurooncol.* 2009 Jan;91(2):199-206. doi: 10.1007/s11060-008-9699-x. Epub 2008 Sep 24. PMID: 18813875.
- 99: Yamamoto T, Nakai K, Matsumura A. Boron neutron capture therapy for glioblastoma. *Cancer Lett.* 2008 Apr 18;262(2):143-52. doi: 10.1016/j.canlet.2008.01.021. Epub 2008 Mar 4. PMID: 18313207.
- 100: Doi A, Kawabata S, Iida K, Yokoyama K, Kajimoto Y, Kuroiwa T, Shirakawa T, Kirihata M, Kasaoka S, Maruyama K, Kumada H, Sakurai Y, Masunaga S, Ono K, Miyatake S. Tumor-specific targeting of sodium borocaptate (BSH) to malignant glioma by transferrin-PEG liposomes: a modality for boron neutron capture therapy. *J Neurooncol.* 2008 May;87(3):287-94. doi: 10.1007/s11060-008-9522-8. Epub 2008 Jan 25. PMID: 18219552.
- 101: Nakagawa N, Akai F, Fukawa N, Fujita Y, Suzuki M, Ono K, Taneda M. Early effects of boron neutron capture therapy on rat glioma models. *Brain Tumor Pathol.* 2007;24(1):7-13. doi: 10.1007/s10014-007-0214-4. Epub 2007 May 25. PMID: 18095138.
- 102: Shibata Y. Boron distribution in the normal rat brain after intravenous injection of boronophenylalanine-fructose. *J Neurooncol.* 2008 Mar;87(1):35-41. doi: 10.1007/s11060-007-9494-0. Epub 2007 Nov 20. PMID: 18026910.
- 103: Miyatake S, Tamura Y, Kawabata S, Iida K, Kuroiwa T, Ono K. Boron neutron capture therapy for malignant tumors related to meningiomas. *Neurosurgery.* 2007 Jul;61(1):82-90; discussion 90-1. doi: 10.1227/01.neu.0000279727.90650.24. PMID: 17621022.
- 104: Hoh DJ, Liu CY, Pagnini PG, Yu C, Wang MY, Apuzzo ML. Chained lightning, part I: Exploitation of energy and radiobiological principles for therapeutic purposes. *Neurosurgery.* 2007 Jul;61(1):14-27; discussion 27-8. doi: 10.1227/01.neu.0000279720.83026.49. PMID: 17621015.
- 105: Tamura Y, Miyatake S, Nonoguchi N, Miyata S, Yokoyama K, Doi A, Kuroiwa T, Asada M, Tanabe H, Ono K. Boron neutron capture therapy for recurrent malignant meningioma. Case report. *J Neurosurg.* 2006 Dec;105(6):898-903. doi: 10.3171/jns.2006.105.6.898. PMID: 17405262.
- 106: Yokoyama K, Miyatake S, Kajimoto Y, Kawabata S, Doi A, Yoshida T, Okabe M, Kirihata M, Ono K, Kuroiwa T. Analysis of boron distribution in vivo for boron neutron capture therapy using two different boron compounds by secondary ion mass spectrometry. *Radiat Res.* 2007 Jan;167(1):102-9. doi: 10.1667/RR0501.1. PMID: 17214510.
- 107: Kawabata S, Miyatake S. [Boron neutron capture therapy for malignant glioma]. *No To Shinkei.* 2006 Dec;58(12):1051-9. Japanese. PMID: 17193955.
- 108: Endo K, Yamamoto T, Shibata Y, Tsuboi K, Matsumura A, Kumada H, Yamamoto K, Sakai T, Sato T, Oikawa M, Ohara Y, Ishii K. Demonstration of inter- and intracellular distribution of boron and gadolinium using micro-proton-induced X-ray emission (Micro-PIXE). *Oncol Res.* 2006;16(2):57-65. doi: 10.3727/000000006783981198. PMID: 16898266.
- 109: Kageji T, Nagahiro S, Matsuzaki K, Mizobuchi Y, Toi H, Nakagawa Y, Kumada H. Boron neutron capture therapy using mixed epithermal and thermal neutron beams in patients with malignant

glioma-correlation between radiation dose and radiation injury and clinical outcome. *Int J Radiat Oncol Biol Phys.* 2006 Aug 1;65(5):1446-55. doi: 10.1016/j.ijrobp.2006.03.016. Epub 2006 Jun 5. PMID: 16750328.

110: Yokoyama K, Miyatake S, Kajimoto Y, Kawabata S, Doi A, Yoshida T, Asano T, Kirihata M, Ono K, Kuroiwa T. Pharmacokinetic study of BSH and BPA in simultaneous use for BNCT. *J Neurooncol.* 2006 Jul;78(3):227-32. doi: 10.1007/s11060-005-9099-4. Epub 2006 Mar 24. PMID: 16557351.

111: Sakurai Y, Ono K, Miyatake S, Maruhashi A. Improvement effect on the depth- dose distribution by CSF drainage and air infusion of a tumour-removed cavity in boron neutron capture therapy for malignant brain tumours. *Phys Med Biol.* 2006 Mar 7;51(5):1173-83. doi: 10.1088/0031-9155/51/5/009. Epub 2006 Feb 8. PMID: 16481686.

112: Miyatake S, Kawabata S, Kajimoto Y, Aoki A, Yokoyama K, Yamada M, Kuroiwa T, Tsuji M, Imahori Y, Kirihata M, Sakurai Y, Masunaga S, Nagata K, Maruhashi A, Ono K. Modified boron neutron capture therapy for malignant gliomas performed using epithermal neutron and two boron compounds with different accumulation mechanisms: an efficacy study based on findings on neuroimages. *J Neurosurg.* 2005 Dec;103(6):1000-9. doi: 10.3171/jns.2005.103.6.1000. PMID: 16381186.

113: Miyatake S, Kuwabara H, Kajimoto Y, Kawabata S, Yokoyama K, Doi A, Tsuji M, Mori H, Ono K, Kuroiwa T. Preferential recurrence of a sarcomatous component of a gliosarcoma after boron neutron capture therapy: case report. *J Neurooncol.* 2006 Jan;76(2):143-7. doi: 10.1007/s11060-005-4174-4. PMID: 16234987.

114: Miyatake S, Kawabata S, Kajimoto Y, Kuroiwa T, Ono K. [Boron neutron capture therapy without craniotomy for malignant gliomas]. *Nihon Rinsho.* 2005 Sep;63 Suppl 9:447-51. Japanese. PMID: 16201562.

115: Kageji T, Nagahiro S, Toi H, Mizobuchi Y, Nakagawa Y. [Boron neutron capture therapy (BNCT) for malignant glioma-present status and the points at issue]. *Nihon Rinsho.* 2005 Sep;63 Suppl 9:442-6. Japanese. PMID: 16201561.

116: Imahori Y, Sasajima H, Mineura K, Imahori Y. [Molecular imaging for brain tumor using positron emission tomography]. *Nihon Rinsho.* 2005 Sep;63 Suppl 9:250-7. Japanese. PMID: 16201531.

117: Matsumura A, Zhang T, Nakai K, Endo K, Kumada H, Yamamoto T, Yoshida F, Sakurai Y, Yamamoto K, Nose T. Combination of boron and gadolinium compounds for neutron capture therapy. An in vitro study. *J Exp Clin Cancer Res.* 2005 Mar;24(1):93-8. PMID: 15943038.

118: Bergenheim AT, Capala J, Roslin M, Henriksson R. Distribution of BPA and metabolic assessment in glioblastoma patients during BNCT treatment: a microdialysis study. *J Neurooncol.* 2005 Feb;71(3):287-93. doi: 10.1007/s11060-004-1724-0. PMID: 15735919.

119: Yamamoto T, Matsumura A, Nakai K, Shibata Y, Endo K, Sakurai F, Kishi T, Kumada H, Yamamoto K, Torii Y. Current clinical results of the Tsukuba BNCT trial. *Appl Radiat Isot.* 2004 Nov;61(5):1089-93. doi: 10.1016/j.apradiso.2004.05.010. PMID: 15308197.

120: Kageji T, Nagahiro S, Mizobuchi Y, Toi H, Nakagawa Y, Kumada H. Radiation injury of boron neutron capture therapy using mixed epithermal- and thermal neutron beams in patients with malignant glioma. *Appl Radiat Isot.* 2004 Nov;61(5):1063-7. doi: 10.1016/j.apradiso.2004.05.058. PMID: 15308193.

- 121: Kageji T, Nagahiro S, Uyama S, Mizobuchi Y, Toi H, Nakamura M, Nakagawa Y. Histopathological findings in autopsied glioblastoma patients treated by mixed neutron beam BNCT. *J Neurooncol.* 2004 May;68(1):25-32. doi: 10.1023/b:neon.0000024725.31515.22. PMID: 15174518.
- 122: Honová H, Safanda M, Petruzelka L, Burian J, Marek M, Rejchrt J, Sus F, Tovarys F, Dbalý V, Honzátko J, Tomandl I, Mares V. Neutronová záchytová terapie v léčbě glioblastoma multiforme. První zkušenosti v České republice [Neutron capture therapy in the treatment of glioblastoma multiforme. Initial experience in the Czech Republic]. *Cas Lek Cesk.* 2004;143(1):44-7. Czech. PMID: 15061119.
- 123: Shibata Y, Matsumura A, Yamamoto T, Akutsu H, Yasuda S, Nakai K, Nose T, Yamamoto K, Kumada H, Hori N, Otake S. Prediction of boron concentrations in blood from patients on boron neutron capture therapy. *Anticancer Res.* 2003 Nov- Dec;23(6D):5231-5. PMID: 14981995.
- 124: Kawabata S, Miyatake S, Kajimoto Y, Kuroda Y, Kuroiwa T, Imahori Y, Kirihata M, Sakurai Y, Kobayashi T, Ono K. The early successful treatment of glioblastoma patients with modified boron neutron capture therapy. Report of two cases. *J Neurooncol.* 2003 Nov;65(2):159-65. doi: 10.1023/b:neon.000003751.67562.8e. PMID: 14686736.
- 125: Takahashi Y, Imahori Y, Mineura K. Prognostic and therapeutic indicator of fluoroboronophenylalanine positron emission tomography in patients with gliomas. *Clin Cancer Res.* 2003 Dec 1;9(16 Pt 1):5888-95. PMID: 14676111.
- 126: Yamamoto T, Matsumura A, Yamamoto K, Kumada H, Hori N, Torii Y, Shibata Y, Nose T. Characterization of neutron beams for boron neutron capture therapy: in- air radiobiological dosimetry. *Radiat Res.* 2003 Jul;160(1):70-6. doi: 10.1667/rr3012. PMID: 12816525.
- 127: Nakagawa Y, Pooh K, Kobayashi T, Kageji T, Uyama S, Matsumura A, Kumada H. Clinical review of the Japanese experience with boron neutron capture therapy and a proposed strategy using epithermal neutron beams. *J Neurooncol.* 2003 Mar- Apr;62(1-2):87-99. doi: 10.1007/BF02699936. PMID: 12749705.
- 128: Yamamoto T, Matsumura A, Yamamoto K, Kumada H, Shibata Y, Nose T. In- phantom two-dimensional thermal neutron distribution for intraoperative boron neutron capture therapy of brain tumours. *Phys Med Biol.* 2002 Jul 21;47(14):2387-96. doi: 10.1088/0031-9155/47/14/302. PMID: 12171329.
- 129: Iwakura M, Kondoh H, Hiratsuka J, Ehara K, Tamaki N, Mishima Y. Effect of boron neutron capture therapy for melanotic and amelanotic melanoma transplanted into mouse brain. *Pigment Cell Res.* 2002 Feb;15(1):67-75. doi: 10.1034/j.1600-0749.2002.00059.x. PMID: 11837459.
- 130: Shinomura T, Furutani H, Osawa M, Ono K, Fukuda K. Anaesthetic management of 27 cases of boron neutron capture therapy for glioblastoma. *Anaesthesia.* 2001 Jul;56(7):665-7. doi: 10.1046/j.1365-2044.2001.01912.x. PMID: 11437767.
- 131: Goodman JH, Yang W, Barth RF, Gao Z, Boesel CP, Staubus AE, Gupta N, Gahbauer RA, Adams DM, Gibson CR, Ferketich AK, Moeschberger ML, Soloway AH, Carpenter DE, Albertson BJ, Bauer WF, Zhang MZ, Wang CC. Boron neutron capture therapy of brain tumors: biodistribution, pharmacokinetics, and radiation dosimetry sodium borocaptate in patients with gliomas. *Neurosurgery.* 2000 Sep;47(3):608-21; discussion 621-2. doi: 10.1097/00006123-200009000-00016. PMID: 10981748.
- 132: Yang W, Barth RF, Bartus RT, Rotaru JH, Moeschberger ML, Ferketich AK, Nawrocky MM, Coderre JA, Goodman JH. Improved survival after boron neutron capture therapy of brain tumors by Cereport-

mediated blood-brain barrier modulation to enhance delivery of boronophenylalanine. *Neurosurgery*. 2000 Jul;47(1):189-97; discussion 197-8. doi: 10.1097/00006123-200007000-00039. PMID: 10917362.

133: Joensuu H, Tenhunen M. Physical and biological targeting of radiotherapy. *Acta Oncol*. 1999;38 Suppl 13:75-83. doi: 10.1080/028418699432806. PMID: 10612500.

134: Brada M. Current approaches to radiation therapy for malignant gliomas. *Front Radiat Ther Oncol*. 1999;33:139-49. doi: 10.1159/000061230. PMID: 10549484.

135: Matsumura A, Shibata Y, Yamamoto T, Yoshida F, Isobe T, Nakai K, Hayakawa Y, Kiriya M, Shimojo N, Ono K, Sakata I, Nakajima S, Okumura M, Nose T. A new boronated porphyrin (STA-BX909) for neutron capture therapy: an in vitro survival assay and in vivo tissue uptake study. *Cancer Lett*. 1999 Jul 1;141(1-2):203-9. doi: 10.1016/s0304-3835(99)00105-6. Erratum in: *Cancer Lett* 2000 Jul 31;155(2):209. PMID: 10454263.

136: Chanana AD, Capala J, Chadha M, Coderre JA, Diaz AZ, Elowitz EH, Iwai J, Joel DD, Liu HB, Ma R, Pendzick N, Peress NS, Shady MS, Slatkin DN, Tyson GW, Wielopolski L. Boron neutron capture therapy for glioblastoma multiforme: interim results from the phase I/II dose-escalation studies. *Neurosurgery*. 1999 Jun;44(6):1182-92; discussion 1192-3. doi: 10.1097/00006123-199906000-00013. PMID: 10371617.

137: Barth RF, Soloway AH, Goodman JH, Gahbauer RA, Gupta N, Blue TE, Yang W, Tjarks W. Boron neutron capture therapy of brain tumors: an emerging therapeutic modality. *Neurosurgery*. 1999 Mar;44(3):433-50; discussion 450-1. doi: 10.1097/00006123-199903000-00001. PMID: 10069580.

138: Barth RF, Yang W, Bartus RT, Moeschberger ML, Goodman JH. Enhanced delivery of boronophenylalanine for neutron capture therapy of brain tumors using the bradykinin analog Cereport (Receptor-Mediated Permeabilizer-7). *Neurosurgery*. 1999 Feb;44(2):351-9; discussion 359-60. doi: 10.1097/00006123-199902000-00062. PMID: 9932889.

139: Shibata Y, Matsumura A, Yamamoto T, Nakagawa K, Yoshii Y, Nose T, Sakata I, Nakajima S, Hayakawa Y, Ono K. Neutron capture therapy with a new boron- porphyrin compound in the rat 9L glioma model. *J Exp Clin Cancer Res*. 1998 Sep;17(3):285-9. PMID: 9894763.

140: Haselsberger K, Radner H, Pendl G. Boron neutron capture therapy for glioblastoma: improvement of boron biodistribution by hyaluronidase. *Cancer Lett*. 1998 Sep 11;131(1):109-11. doi: 10.1016/s0304-3835(98)00206-7. PMID: 9839625.

141: Imahori Y, Ueda S, Ohmori Y, Sakae K, Kusuki T, Kobayashi T, Takagaki M, Ono K, Ido T, Fujii R. Positron emission tomography-based boron neutron capture therapy using boronophenylalanine for high-grade gliomas: part I. *Clin Cancer Res*. 1998 Aug;4(8):1825-32. PMID: 9717808.

142: Elowitz EH, Bergland RM, Coderre JA, Joel DD, Chadha M, Chanana AD. Biodistribution of p-boronophenylalanine in patients with glioblastoma multiforme for use in boron neutron capture therapy. *Neurosurgery*. 1998 Mar;42(3):463-8; discussion 468-9. doi: 10.1097/00006123-199803000-00004. PMID: 9526978.

143: Imahori Y, Ueda S, Ohmori Y, Kusuki T, Ono K, Fujii R, Ido T. Fluorine-18-labeled fluoroboronophenylalanine PET in patients with glioma. *J Nucl Med*. 1998 Feb;39(2):325-33. PMID: 9476945.

144: Nakagawa Y, Hatanaka H. Boron neutron capture therapy. Clinical brain tumor studies. *J*

- Neurooncol. 1997 May;33(1-2):105-15. doi: 10.1023/a:1005781517624. PMID: 9151228.
- 145: Haselsberger K, Radner H, Pendl G. Na<sub>2</sub>B<sub>12</sub>H<sub>11</sub>SH (BSH) in combination with systemic hyaluronidase: a promising concept for boron neutron capture therapy for glioblastoma. Neurosurgery. 1996 Aug;39(2):321-5; discussion 325-6. doi: 10.1097/00006123-199608000-00016. PMID: 8832669.
- 146: Delattre JY, Uchuya M. Radiotherapy and chemotherapy for gliomas. Curr Opin Oncol. 1996 May;8(3):196-203. doi: 10.1097/00001622-199605000-00005. PMID: 8804817.
- 147: Yang W, Barth RF, Carpenter DE, Moeschberger ML, Goodman JH. Enhanced delivery of boronophenylalanine for neutron capture therapy by means of intracarotid injection and blood-brain barrier disruption. Neurosurgery. 1996 May;38(5):985-92. doi: 10.1097/00006123-199605000-00027. PMID: 8727825.
- 148: Leppälä J, Kallio M, Nikula T, Nikkinen P, Liewendahl K, Jääskeläinen J, Savolainen S, Gylling H, Hiltunen J, Callaway J, et al. Accumulation of 99mTc- low-density lipoprotein in human malignant glioma. Br J Cancer. 1995 Feb;71(2):383-7. doi: 10.1038/bjc.1995.78. PMID: 7841057; PMCID: PMC2033577.
- 149: Stragliotto G, Fankhauser H. Biodistribution of boron sulfhydryl for boron neutron capture therapy in patients with intracranial tumors. Neurosurgery. 1995 Feb;36(2):285-92; discussion 292-3. doi: 10.1227/00006123-199502000-00007. PMID: 7731508.
- 150: Haselsberger K, Radner H, Pendl G. Boron neutron capture therapy: boron biodistribution and pharmacokinetics of Na<sub>2</sub>B<sub>12</sub>H<sub>11</sub>SH in patients with glioblastoma. Cancer Res. 1994 Dec 15;54(24):6318-20. PMID: 7987820.
- 151: Haselsberger K, Radner H, Gössler W, Schlagenhaufen C, Pendl G. Subcellular boron-10 localization in glioblastoma for boron neutron capture therapy with Na<sub>2</sub>B<sub>12</sub>H<sub>11</sub>SH. J Neurosurg. 1994 Nov;81(5):741-4. doi: 10.3171/jns.1994.81.5.0741. PMID: 7931621.
- 152: Haritz D, Gabel D, Huiskamp R. Clinical phase-I study of Na<sub>2</sub>B<sub>12</sub>H<sub>11</sub>SH (BSH) in patients with malignant glioma as precondition for boron neutron capture therapy (BNCT). Int J Radiat Oncol Biol Phys. 1994 Mar 30;28(5):1175-81. doi: 10.1016/0360-3016(94)90492-8. PMID: 8175403.
- 153: Hatanaka H, Nakagawa Y. Clinical results of long-surviving brain tumor patients who underwent boron neutron capture therapy. Int J Radiat Oncol Biol Phys. 1994 Mar 30;28(5):1061-6. doi: 10.1016/0360-3016(94)90479-0. PMID: 8175390.
- 154: Nakagawa Y, Hatanaka H, Moritani M, Kitamura K, Matsumoto K, Kobayashi M. Partial deuteration and blood-brain barrier (BBB) permeability. Acta Neurochir Suppl (Wien). 1994;60:410-2. doi: 10.1007/978-3-7091-9334-1\_111. PMID: 7976604.
- 155: Fankhauser H, Gavin PR, Stragliotto G. Proposal of a treatment protocol for boron neutron capture therapy of supratentorial malignant gliomas. Strahlenther Onkol. 1993 Jan;169(1):71-5. PMID: 8434344.
- 156: Saris SC, Solares GR, Wazer DE, Cano G, Kerley SE, Joyce MA, Adelman LS, Harling OK, Madoc-Jones H, Zamenhof RG. Boron neutron capture therapy for murine malignant gliomas. Cancer Res. 1992 Sep 1;52(17):4672-7. PMID: 1511433.
- 157: Hatanaka H, Moritani M, Camillo M. Possible alteration of the blood-brain barrier by boron-

neutron capture therapy. *Acta Oncol.* 1991;30(3):375-8. doi: 10.3109/02841869109092389. PMID: 2036249.

158: Goodman JH, McGregor JM, Clendenon NR, Gahbauer RA, Barth RF, Soloway AH, Fairchild RG. Inhibition of tumor growth in a glioma model treated with boron neutron capture therapy. *Neurosurgery.* 1990 Sep;27(3):383-8. doi: 10.1097/00006123-199009000-00007. PMID: 2234330.

159: Clendenon NR, Barth RF, Gordon WA, Goodman JH, Alam F, Staubus AE, Boesel CP, Yates AJ, Moeschberger ML, Fairchild RG, et al. Boron neutron capture therapy of a rat glioma. *Neurosurgery.* 1990 Jan;26(1):47-55. doi: 10.1097/00006123-199001000-00007. PMID: 2294479.

160: Goodman JH, McGregor JM, Clendenon NR, Gahbauer RA, Barth RF, Soloway AH, Fairchild RG. Ultrastructural microvascular response to boron neutron capture therapy in an experimental model. *Neurosurgery.* 1989 May;24(5):701-8. doi: 10.1227/00006123-198905000-00007. PMID: 2716978.

161: Montagno Ede A, Moreira Filho L, Hatanaka H. Three cases of terminal stage malignant gliomas in which the diffusely disseminated intraventricular tumors became the target of boron neutron capture therapy. *Strahlenther Onkol.* 1989 Feb-Mar;165(2-3):238-40. PMID: 2494740.

162: Finkel GC, Poletti CE, Fairchild RG, Slatkin DN, Sweet WH. Distribution of 10B after infusion of Na210B12H11SH into a patient with malignant astrocytoma: implications for boron neutron capture therapy. *Neurosurgery.* 1989 Jan;24(1):6-11. doi: 10.1227/00006123-198901000-00002. PMID: 2927599.

163: Perks CA, Mill AJ, Constantine G, Harrison KG, Gibson JA. A review of boron neutron capture therapy (BNCT) and the design and dosimetry of a high-intensity, 24 keV, neutron beam for BNCT research. *Br J Radiol.* 1988 Dec;61(732):1115-26. doi: 10.1259/0007-1285-61-732-1115. PMID: 3064858.

164: Hatanaka H, Sano K, Yasukochi H. [Boron-neutron capture therapy in brain tumors and other cancers—a radiosurgery]. *Gan To Kagaku Ryoho.* 1988 Apr;15(4 Pt 2-2):1115-23. Japanese. PMID: 3289496.

165: Abe M, Amano K, Kitamura K, Ohta M, Tateishi J, Hatanaka H. Capillary permeability and boron distribution in ethylnitrosourea-induced rat glioma. *Neurosurgery.* 1988 Jan;22(1 Pt 1):23-31. doi: 10.1227/00006123-198801010-00004. PMID: 3344084.

<sup>1)</sup>

Takahara K, Miyatake SI, Azuma H, Shiroki R. Boron neutron capture therapy for urological cancers. *Int J Urol.* 2022 Mar 3. doi: 10.1111/iju.14855. Epub ahead of print. PMID: 35240726.

<sup>2)</sup>

Gao Z, Horiguchi Y, Nakai K, Matsumura A, Suzuki M, Ono K, Nagasaki Y. Use of boron cluster-containing redox nanoparticles with ROS scavenging ability in boron neutron capture therapy to achieve high therapeutic efficiency and low adverse effects. *Biomaterials.* 2016 Jul 11;104:201-212. doi: 10.1016/j.biomaterials.2016.06.046. [Epub ahead of print] PubMed PMID: 27467416.

<sup>3)</sup>

Kageji T, Nagahiro S, Mizobuchi Y, Matsuzaki K, Nakagawa Y, Kumada H. Boron neutron capture therapy (BNCT) for newly-diagnosed glioblastoma: comparison of clinical results obtained with BNCT and conventional treatment. *J Med Invest.* 2014;61(3-4):254-63. PubMed PMID: 25264042.

<sup>4)</sup>

Kageji T, Sogabe S, Mizobuchi Y, Nakajima K, Shinji N, Nakagawa Y. Radiation-induced meningiomas after BNCT in patients with malignant glioma. *Appl Radiat Isot.* 2015 Jun 20. pii: S0969-8043(15)30049-X. doi: 10.1016/j.apradiso.2015.06.004. [Epub ahead of print] PubMed PMID:

26122975.

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