

## Research Article

# Effectiveness of Newly Developed Water-Equivalent Mouthpiece during External Beam Radiotherapy for Oral Cancer

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## Abstract

The objective of this study was to research the effectiveness of newly developed water-equivalent mouthpiece during external beam radiotherapy for oral cancer. In external beam radiotherapy for cancer of the tongue, floor of the mouth, and lower gingiva, it is possible to prescribe a low dose to the upper gingiva and hard palate at an open mouth position using a mouthpiece. However, the inhomogeneity correction resulting from the air cavity and the mobility of the tongue produced by an open mouth position should be considered. Therefore, a new mouthpiece was designed to be fixed by the dental arch, and the air cavity of the mouth can be filled with water-equivalent material. In 30 patients with previously treated oral cancer, the simulated homogeneity index of the calculated water-equivalent mouthpiece by a treatment-planning system was significantly better than that of a conventional mouthpiece ( $p = 0.004$ ). This new mouthpiece facilitates excellent dose distribution while attaining immobilization of the tongue in patients with oral cancer.

**Keywords:** External beam radiation therapy; Oral cancer; Head and neck cancer; Mouthpiece; Water-equivalent; Immobilization

## Abbreviations

EBRT: External Beam Radiotherapy; 3D-CRT: Three-Dimensional Conformal Radiotherapy; CT: Computed Tomography; GTV: Gross Tumor Volume; CTV: Clinical Target Volume; PTV: Planning Target Volume; HI: Homogeneity Index; D max: maximum dose of Planning Target Volume (PTV); D min: minimum dose of PTV; IMRT: Intensity-Modulated Radiotherapy; N: Numbers; WE MP: Water-Equivalent Mouthpiece; C MP: Cotton piece; AL: Antero-Lateral beams; LO: Lateral Opposed beams; WE MP HI: Homogeneity Index of Water-Equivalent Mouthpiece; C MP HI: Homogeneity Index of rolled Cottonpiece; SD: Standard Deviation of mean; SEM: Standard Error of Mean

## Introduction

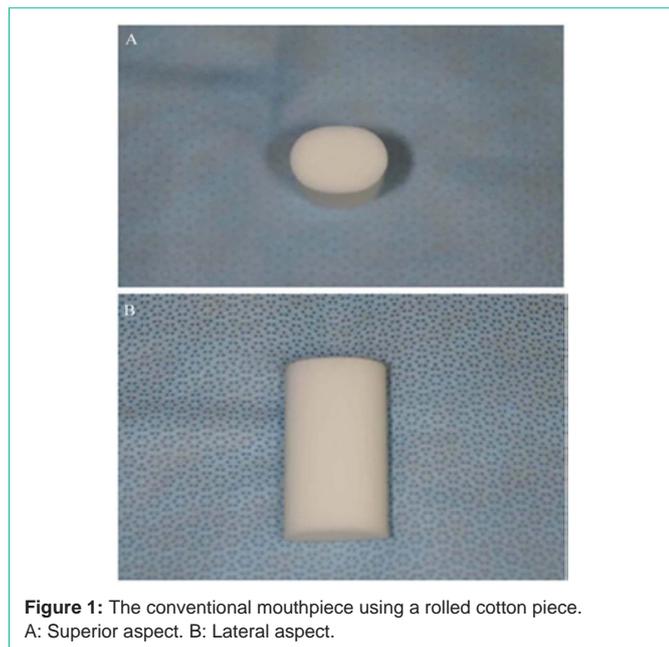
Radiotherapy is widely utilized for oral cancer treatment in an effort to preserve chewing, speech, and swallowing functions and the shape of the oral structures. A mouthpiece is used to depress the tongue away from the hard palate in External Beam Radiotherapy (EBRT) for patients with cancer of the floor of the mouth, lower gingiva, and tongue [1]. It is possible to administer adequate EBRT doses to tumors in patients with cancer of the tongue and floor of the mouth while sparing the upper gingiva and hard palate. Mouthpieces have the benefits of reducing the dose administered to tongue of patients with cancer of the upper gingiva, hard palate, and maxillary sinus and reducing the dose administered to the upper gingiva, hard palate, and maxillary sinus of patients with cancer of the tongue.

According to Willner, et al. an individualized mouthpiece can facilitate accurate and reproducible positioning in patients with head and neck cancer using Three-Dimensional Conformal Radiotherapy

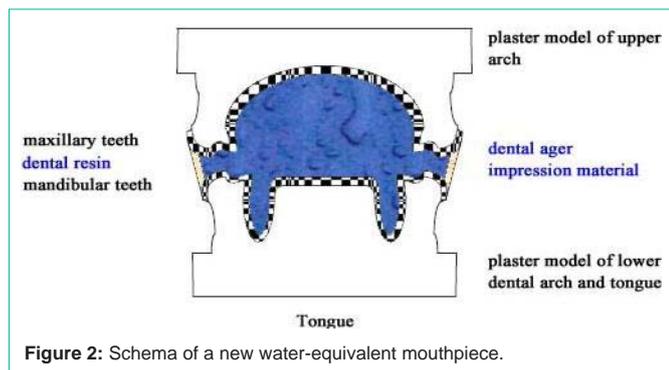
(3D-CRT) [2]. They reported that random and systematic deviations in each of the three directions (craniocaudal, anteroposterior, and mediolateral axes) are within the range of  $\pm 4$  mm and within the range or even less than the deviations described for most thermoplastic masks.

A mouthpiece is used to both depress and fix the tongue during simulation and treatment of EBRT, especially in patients with cancer of the tongue or floor of the mouth [3,4]. However, the position of the mouthpiece may be uncontrollable during treatment [5]. Additionally, interfractional organ motion of the tongue results in uncertainty because of involuntary tongue movement. The silicone mouthpiece, which is fixed by the dental arch, can be steadily immobilized while depressing the tongue. Stereotactic radiotherapy to metastatic brain tumors has a biological advantage over stereotactic radiosurgery [6-8]. The custom-made mouthpiece worn on the palate increases the setup accuracy in stereotactic radiotherapy to the metastatic brain tumor [9-11]. The silicone mouthpiece also allows the user to easily attain a stable tongue position. However, scattered radiation can be produced because the silicone must be within the treatment field of patients with oral cancer, and severe stomatitis as the reason of an interrupting radiotherapy occurs.

A conventional method using a rolled cotton piece has been used to depress the tongue and immobilize the mandible (Figure 1). This cotton piece is influenced by scattered radiation less than silicone mouthpiece, and is easy to wear, low-cost, and hygienic. However, it is thought to have the disadvantage of positioning accuracy compared with the silicone mouthpiece with respect to fixation of the tongue. Therefore, we developed a new water-equivalent mouthpiece that can improve the inhomogeneity correction while maintaining accurate



**Figure 1:** The conventional mouthpiece using a rolled cotton piece. A: Superior aspect. B: Lateral aspect.



**Figure 2:** Schema of a new water-equivalent mouthpiece.

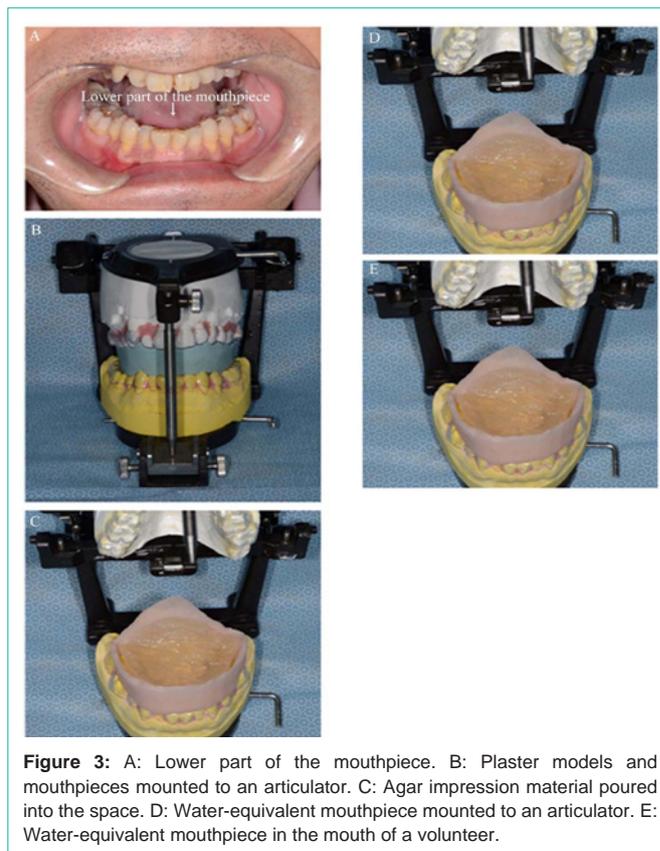
immobilization. It can be fixed by the dental arch, and the air cavity of the mouth can be filled with water-equivalent material.

The purpose of this study was to design a mouthpiece that can fill the air cavity of the mouth with a water-equivalent material while immobilizing the tongue without a delay in the commencement of EBRT, and to verify the usefulness of this new mouthpiece by simulation using a treatment-planning workstation.

**Materials and Methods**

**Preparation of water-equivalent mouthpiece to immobilize the tongue**

An impression of the upper dental arch was routinely taken. An impression of both the lower dental arch and the tongue together was taken using an upper impression tray to immobilize the tongue. Thermoplastic disks of 1-mm thickness (Erkodur; Erkodent, Pfalzgratenweiler, Germany) were then pressed by a thermoplastic former (Erkopress; Erkodent) to the upper and lower plaster models (Figure 2). The lower part of the mouthpiece was set onto the lower dental arch to immobilize the tongue (Figure 3a). A silicone bite was taken in the open mouth position and cut along the buccal edge of each dental arch for enclosure by the tray resin. The posterior limit was extended as long as possible to fill the air cavity of the mouth.



**Figure 3:** A: Lower part of the mouthpiece. B: Plaster models and mouthpieces mounted to an articulator. C: Agar impression material poured into the space. D: Water-equivalent mouthpiece mounted to an articulator. E: Water-equivalent mouthpiece in the mouth of a volunteer.

The plaster models and mouthpieces were mounted to an articulator (Figure 3b). The tray resin was enclosed around the silicone bite as thin as possible from the upper to lower teeth. After the silicone bite was removed, an agar impression material (Dupligel; Shofu, Kyoto, Japan) was poured into the space (Figure 3c). The final mouthpiece was almost equivalent to water, in contrast to silicone material, because thinner resin and water-equivalent agar impression material were used (Figure 3d). Figure 3e shows this mouthpiece in the mouth of a volunteer.

**Analyses of the effect of the new water-equivalent mouthpiece on dose distribution**

In total, 111 patients with oral cancer were treated by EBRT from September 2002 to August 2011 in Tokushima University Hospital. Thirty dentulous patients who underwent irradiation of cancer of the tongue, lower gingiva, and floor of the mouth by the conventional method using a rolled cotton piece in the open mouth position were included in this institutional review board-approved clinical investigation. The numbers of primary cancer sites involving the tongue, lower gingiva, and floor of the mouth were 13, 9, and 8, respectively (Table 1). Planning Computed Tomography (CT) images were reconstructed to a 3-mm thickness with a 1-mm slice thickness using an Asteion (Toshiba Medical Systems, Tochigi, Japan). The CT data were exported to a treatment planning workstation (XiO; Elekta,

**Table 1:** The numbers of primary cancer sites.

Primary site	Tongue	Lower gingiva	Floor of the mouth
N	13	9	8

N: Number

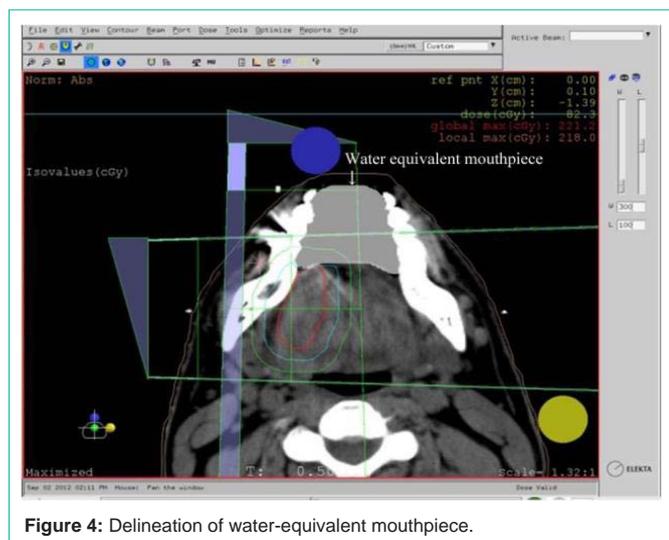


Figure 4: Delineation of water-equivalent mouthpiece.

Inc., MO, USA). The calculated water-equivalent electron density mouthpiece was delineated on the CT images using commercial treatment planning software possessing the ability to delineate an outline with any electron density. Figure 4 shows the delineation of the calculated water-equivalent mouthpiece. According to our policy, the Gross Tumor Volume (GTV) – the Clinical Target Volume (CTV) margin, Planning Target Volume (PTV) – CTV margin, and leaf margin were set to 5, 5, and 3 mm, respectively. A 4-MV photon-penetrating machine was used (MEVATRON KD2/7450; Siemens, Tokyo, Japan). The field arrangements were as follows: a 15° opposed lateral wedge pair technique if the GTV was across the midline, and a 30° anterolateral wedge pair technique if the GTV was not across the midline. A bolus was used to improve the dose distribution if the PTV was close to the skin surface. Dose homogeneity was assessed with the Homogeneity Index (HI), which is defined as the maximum PTV dose (Dmax) / minimum PTV dose (Dmin) to evaluate the usefulness of the water-equivalent mouthpiece and conventional method using a rolled cotton piece.

**Statistical analysis**

All statistical analyses were performed using PASW Statistics 18 (SPSS Japan Inc., Tokyo, Japan). The HI calculated from XiO of both the conventional method using a rolled cotton piece and the new mouthpiece was tested using Student’s t-test. The difference in the HI between with and without the bolus and in each beam direction was tested using Levene’s test. Intergroup differences were considered statistically significant when the p value was <0.05.

**Results**

According to the degree of invasion of the primary lesion, 21 patients were planned to undergo the 15° opposed lateral wedge pair technique and 9 patients were planned to undergo the 30° anterolateral wedge pair technique. Because the Dmin was <65% of the prescribed doses without a bolus, there were 13 patients with a bolus (data not shown).

**Inhomogeneity correction: analysis of calculated water-equivalent mouthpiece method**

The Dmax, Dmin, and HI of the water-equivalent mouthpiece method and conventional method using a rolled cotton piece

Table 2: Dmax, Dmin and HI of the water-equivalent and cotton piece by two different beam directions and with/without of bolus.

WE MP			C MP			Beam directions	Bolus
Dmax	Dmin	HI	Dmax	Dmin	HI		
212	167	1.27	212	167	1.27	AL	With
223	160	1.39	224	160	1.40	AL	With
227	161	1.41	228	159	1.43	AL	With
210	154	1.36	208	151	1.38	AL	With
213	160	1.33	212	159	1.33	AL	Without
218	156	1.40	218	156	1.40	AL	Without
218	168	1.30	218	168	1.30	AL	Without
215	156	1.38	217	156	1.39	AL	Without
220	156	1.41	224	154	1.45	AL	Without
219	153	1.43	219	153	1.43	LO	With
231	154	1.50	231	155	1.49	LO	With
233	158	1.47	235	161	1.46	LO	With
228	153	1.49	228	153	1.49	LO	With
213	155	1.37	214	145	1.48	LO	With
216	154	1.40	216	156	1.38	LO	With
219	153	1.43	219	153	1.43	LO	With
219	154	1.42	219	150	1.46	LO	With
211	145	1.46	218	136	1.60	LO	With
206	165	1.25	207	165	1.25	LO	Without
207	166	1.25	207	147	1.41	LO	Without
214	167	1.28	214	152	1.41	LO	Without
210	162	1.30	224	165	1.36	LO	Without
214	159	1.35	215	154	1.40	LO	Without
214	156	1.37	214	157	1.36	LO	Without
213	153	1.39	213	152	1.40	LO	Without
230	157	1.46	230	157	1.46	LO	Without
224	158	1.42	228	158	1.44	LO	Without
222	148	1.50	223	150	1.49	LO	Without
225	157	1.43	226	148	1.53	LO	Without
223	154	1.45	223	154	1.45	LO	Without

WE MP: Water-Equivalent Mouthpiece; C MP: Cotton Piece; Dmax (Gy): maximum dose of Planning Target Volume (PTV); Dmin (Gy): minimum dose of PTV; HI: Homogeneity Index; AL: Antero-Lateral beams; LO: Lateral Opposed beams.

in different beam directions and with and without a bolus were calculated (Table 2). Table 3 shows the HI of each method.

The mean HI using the calculated water-equivalent mouthpiece was 1.39, and that of the conventional method using a rolled cotton piece was 1.42. The HI of the calculated water-equivalent mouthpiece was significantly better than that of the conventional method using a rolled cotton piece (Student’s t-test, p: 0.004). The HIs of the calculated water-equivalent mouthpiece in two different beam directions and with and without the bolus are shown in (Tables 4 & 5). There were no significant differences with the two different beam directions (Levene’s test, p: 0.819) or with or without the bolus (Levene’s test, p: 0.079).

**Table 3:** HI of the ideal water-equivalent mouthpiece and rolled cotton piece.

	Mean	SD	SEM
WE MP HI	1.389	0.072	0.013
C MP HI	1.418	0.074	0.013

WE MP HI: Homogeneity Index of Water-Equivalent Mouthpiece; C MP HI: Homogeneity Index of rolled Cotton piece; SD: Standard Deviation of mean; SEM: Standard Error of Mean.

**Table 4:** HI of ideal water-equivalent mouthpiece by the different beam directions.

	N	Mean	SD	SEM
AL	9	1.361	0.051	0.017
LO	21	1.401	0.077	0.017

AL: Antero-Lateral beams; LO: Lateral Opposed beams; N: Number; SD: Standard Deviation of mean; SEM: Standard Error of Mean.

**Table 5:** HI of ideal water-equivalent mouthpiece with/without bolus.

	N	Mean	SD	SEM
Without bolus	17	1.369	0.075	0.018
With bolus	13	1.415	0.061	0.017

N: Numbers; SD: Standard Deviation of mean; SEM: Standard Error of Mean.

## Discussion

Positioning accuracy is required to minimize the CTV-PTV margin during EBRT. Because this new mouthpiece immobilizes the tongue, the inaccuracy caused by involuntary movement and flexibility of the tongue is none. The method used to prepare the mouthpiece can be standardized although it is custom-made. Additionally, the delay in the commencement of radiotherapy is shortened because the mouthpiece can be fabricated using a conventional upper impression tray and conventional dental materials. Finally, this new mouthpiece fills the air cavity of the mouth with water-equivalent material to improve the inhomogeneity correction. Therefore, the dose homogeneity using this new mouthpiece is expected to be better than that using the conventional method with a rolled cotton piece.

A two-dimensional treatment plan was used until 2002 in Tokushima University Hospital. Thereafter, 30 eligible edentulous patients were selected to undergo the conventional method using a rolled cotton piece to simulate the dose volume histogram using this new mouthpiece. Only two treatment field arrangements were used. The PTV of the primary lesion (not involving the neck area) was evaluated because the influence of the inhomogeneity correction mainly occurred around the mouthpiece. The mean HI of the calculated water-equivalent mouthpiece was better than that of the conventional method using rolled cotton piece although this study was small sample size. And there were no significant differences with the two different beam directions. On the other hand, there was a trend towards the significance with or without the bolus. For all 13 patients with a bolus, the thickness of the bolus was set to be minimum so that Dmin was to be more than 65% of the prescribed dose. Therefore, it was thought that the patients with a bolus had a trend toward high HI in spite of compensation by bolus. A previously described mouthpiece was used to depress the tongue and immobilize the mandible in patients with head and neck cancer [2,4,5]. The mouthpiece used during EBRT in patients with head and neck cancer is fabricated from high-density material more than soft tissues (i.e., silicone, or similar materials). The delineation of the tissue near the mouthpiece is influenced by the scattered radiation, even if the image

is fused with magnetic resonance imaging to delineate the contour.

There are two precautions associated with this new water-equivalent mouthpiece. First, this new water-equivalent mouthpiece cannot be stabilized during each treatment session in edentulous patients because it can be fixed with the teeth. Therefore, it is not applicable to edentulous patients. Second, the posterior border should be limited to the anterior border of the soft palate to reduce the gag reflex.

EBRT shift from 3D-CRT to Intensity-Modulated Radiotherapy (IMRT). Patients undergoing IMRT exhibit comparable acute and significantly less severe late toxicity than do those undergoing 3D-CRT [12]. Therefore, IMRT for oral cancer has usually been performed to spare the parotid gland [13-15]. Wagner, et al. refrained from the use of a mouthpiece during treatments such as IMRT to reduce the dose and increase the accuracy of the treatment in patients with head and neck cancer [5]. At Tokushima University Hospital, IMRT for head and neck cancer is currently performed with a closed mouth in patients with an unstable mandible. However, there are patients in whom the tongue contacts the palate on treatment-planning CT images in the closed mouth position. The dose for normal tissues within the PTV margins is high in these patients. The normal adjacent tissues near the tumor can be positioned for separation from the opposite side of the jaw using this new mouthpiece. Thus, the inhomogeneity correction may be improved because of the use of the water-equivalent material. The positioning accuracy of this new mouthpiece is considered to be equivalent to that of a silicone mouthpiece by fixation using the dental arch.

Postoperative radiation alone or combined with concurrent chemotherapy is an established adjuvant treatment in patients at high risk for disease recurrence. Such risk factors include positive margins, extracapsular nodal extension, lymphovascular invasion, and perineural invasion [16]. The optimal management of oral cancer typically involves surgical resection followed by adjuvant radiotherapy or chemoradiotherapy in the setting of such risk factors, and the outcomes of patients who were treated with a radiation-based approach were consistently inferior to those treated with surgery followed by adjuvant radiotherapy or chemoradiotherapy [17]. Definitive IMRT using this new mouthpiece would allow for better locoregional control in patients with oral cancer because a better dose distribution can be attained. Furthermore, this new mouthpiece is thought to be effective against oral cancer and other head and neck cancers, such as cancer of the maxillary sinus, soft palate, and base of the tongue.

In conclusion, the calculated HI for 30 previously treated patients with oral cancer significantly improved with the use of the herein-described water-equivalent mouthpiece. The mouthpiece is custom-made for immobilization of the tongue, but the preparation method can be standardized. In addition, the mouthpiece can be rapidly fabricated by a general practitioner without a delay in the commencement of EBRT. This new mouthpiece is expected to be used in not only 3D-CRT but also IMRT for head and neck cancer.

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