# **Assessment of Organ‑at‑risk Sparing in Esophageal Cancer: A Comparative Dosimetric Evaluation of Hybrid, Noncoplanar, and Coplanar RapidArc Plans**

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## **Abstract**

**Aim:** The purpose of this study is to improve the precision of radiation treatment and sparing of organ–at–risk (OAR) in patients with thoracic esophageal cancer (EC) affecting the heart, lung, and spinal cord. To improve and personalize cancer treatment plans, it assesses the dosimetric benefits of coplanar RapidArc (RA<sub>c</sub>), hybrid arc (RA<sub>Hyb</sub>), and noncoplanar RapidArc (RA<sub>nc</sub>). **Materials and Methods:** Fourteen patients with EC were chosen for our investigation from our hospital's database. RapidArc (RA) plan patients had already received treatment. Retrospectively, additional  $RA_{nc}$  and  $RA_{Hyb}$  plans were made with a prescription dose of 50.4 Gy in 28 fractions for the planning target volume (PTV). A prescription dose of 95% of PTV was used, so that three different treatment planning procedures could be compared. The cumulative dose‑volume histogram was used to analyze the plan quality indices homogeneity index (HI), conformity index (CI), conformation number (CN) as well as the OARs doses to the lung, heart, and spinal cord. **Results:** In comparison to RA<sub>c</sub> and RA<sub>nc</sub> techniques, the study indicated that RA<sub>Hyb</sub> plans significantly increased D95%, CI and HI; Dmax and CN did not differ substantially. In addition, compared to RA<sub>c</sub> (lung:  $16.15 \pm 0.03$  Gy and heart:  $23.91 \pm 4.67$  Gy) and RA<sub>nc</sub> (lung:  $15.24 \pm 0.03$  Gy and heart  $23.82 \pm 5.10$  Gy) plans, RA<sub>Hyb</sub> resulted in significantly lower mean lung doses (15.10  $\pm$  0.03 Gy) and heart doses (21.33  $\pm$  6.99 Gy). Moreover, the RA<sub>Hyb</sub> strategy showed a statistically significant  $(P < 0.05)$  lower average MU (452.7) than both the RA<sub>c</sub> (517.5) and RA<sub>c</sub> (566.2) plans. **Conclusion:** The D95%, conformity, and homogeneity indices were better for hybrid arc plans compared to RA<sub>c</sub> and RA<sub>nc</sub> plans. They also successfully managed to reduce the lung and heart doses as well as the mean MU per fraction.

**Keywords:** Esophageal cancer, hybrid, noncoplanar, radiotherapy, volumetric modulated arc therapy



# **Introduction**

Esophageal cancer (EC) poses a significant global health challenge, ranking seventh in cancer incidence and sixth in mortality worldwide. With over 600,000 new cases and  $500,000$  deaths annually,<sup>[1]</sup> addressing this aggressive disease requires innovative treatment approaches. Risk factors for EC include tobacco, alcohol, spicy foods, and betel nut chewing, prevalent in the Indian subcontinent.[2] Radical resection is only possible in 20% of patients, so in these situations, radiotherapy remains the primary treatment option for the majority.[3]

For several decades, conventional radiation therapy (2DRT), usually delivered in two stages, has been the norm. Phase 2 three fields (one anterior and two posterior oblique fields), with or without wedges are used, whereas Phase 1 frequently



uses parallel opposing, anterior, and posterior (AP/PA) fields. Nevertheless, the dose delivery restrictions of this 2DRT technique affect normal structures such as the heart, lung, and spinal cord, making it difficult to achieve dosage homogeneity in the target volume and comply with limitation for OAR. The challenge of increasing the local esophageal dose with 2DRT has contributed to the poor treatment success in EC, as evidenced by 5-year disease survival rate of approx 10.9%.<sup>[4]</sup>

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Recent developments in radiotherapy include the availability of multileaf collimators (MLCs) and improvements in treatment planning algorithms. The most important of these developments is intensity-modulated radiation therapy (IMRT), which stands out as a significant achievement in radiotherapy history, and as a major advance in radiation oncology and EC treatment.<sup>[5]</sup> According to numerous studies,<sup>[5-7]</sup> IMRT emerged as a revolutionary advancement in radiotherapy and offered superior dose distribution and prognosis compared to three‑dimensional conformal radiotherapy (3DCRT). Despite its benefits, IMRT comes with drawbacks such as prolonged treatment times and potentially increased risks of secondary cancer development.[8] RapidArc is a type of volumetric modulated arc therapy (VMAT) representing advanced dynamic volumetric IMRT technology developed by Varian Medical Systems. It provides a dynamic MLC, gantry speed adjustment, and variable dose-rate adjustment, even during gantry movement. Renowned for its efficiency, speed, and precision, RapidArc reduces treatment times while achieving comparable dose distributions to IMRT through single or multiple arc rotations of the gantry. Recent studies have showcased the superiority of RapidArc over traditional IMRT in terms of planning target volume (PTV) conformity and minimizing doses to OARs.[9,10]

In addition, hybrid arc plans combining static and dynamic beams have shown potential for optimizing dose distribution.<sup>[11-14]</sup> When comparing these methods with traditional IMRT and VMAT designs, better dosimetric results have been shown. To enhance OAR dose sparing in the treatment of EC, noncoplanar IMRT, noncoplanar VMAT, and VMAT schemes have been suggested.<sup>[15,16]</sup> Although patients with neurological or cardiovascular diseases should be treated with caution, these strategies seek to balance PTV coverage with dose reduction to adjacent organs.<sup>[15]</sup> The feasibility and effectiveness of noncoplanar VMAT plan to achieve ideal dose distributions with few side effects were shown by Martini

*et al*. [16] In this study, we conducted a dosimetric comparison of conventional RapidArc (RA<sub>c</sub>), noncoplanar RapidArc (RA<sub>nc</sub>), and hybrid arc  $(RA<sub>Hvb</sub>)$  plans. The purpose of this study was to establish whether the  $\text{RA}_{\text{nc}}$  and  $\text{RA}_{\text{Hvb}}$  plans have a dosimetric advantage over the  $RA_c$  plans and find out which is optimal for EC patients.

# **Materials and Methods**

This retrospective study was conducted in the Department of Radiation Oncology, State Cancer Institute, IGIMS, Patna, Bihar on 14 patients who were treated for thoracic EC between September 2021 and October 2023. All patients had previously received a RA<sub>c</sub> technique plan on a TrueBeam SVC linear accelerator machine with 6MV photon beam. This machine is equipped with a 120-millennium MLC from Varian Medical Systems, Palo Alto, CA, USA. The institutional ethics committee approved this study. Patient characteristics are summarized in Table 1.

#### **Target volume and organ‑at‑risk delineation**

Positron emission tomography‑computed tomography (PET-CT) simulation was utilized to exclude distant metastases. Patients were positioned supine with hands placed above the heads to create a thoracic mold using a uniform thermoplastic cast. CT simulation with a slice thickness of 2.5 mm was performed from C2 to L4 vertebrae. The gross tumor volume (GTV), clinical target volume (CTV), PTV, lymph nodes and OARs including the lungs, heart, and spinal cord, were delineated on the Eclipse Somavision contouring workstation version 16.0.14 according to the Radiation Therapy Oncology Group (RTOG) 0436 protocol.<sup>[17]</sup> The GTV was identified based on visible tumors or lymph nodes on CT scans, aided by fusion with PET‑CT simulation images, and confirmed by a radiologist. The CTV was delineated with 3–4.5 cm superior-inferior margins and a 1.5 cm radial margin relative to the GTV. Lymph nodes were contoured



PTV: Planning target volume, SD: Standard deviation

with 1 cm margins in all directions, considering anatomical boundaries such as lungs and bones. The PTV was defined with an additional 0.5 cm margin to the CTV, considering GTV delineation and nearby critical structures. The prescribed dose (PD) was 50.4 Gy delivered in 28 fractions with 1.8 Gy per fraction. OARs included the heart, lungs, spinal cord, liver, spleen, and trachea. Dose constraints for OARs are given in Table 2 and were used in optimization for treatment planning.

#### **Treatment planning**

Radiation therapy for RA is a novel treatment approach that involves a single 360° gantry rotation to deliver radiation to the patient. This method produces an intensity-modulated dose distribution by adjusting the gantry speed, dose rate, and MLC locations and velocities. To optimize the RA dose, an aperture‑based technique is used, which considers the monitor unit (MU) weights and MLC leaf positions as optimization parameters. One hundred and seventy‑seven control points are used to define the entire gantry rotation. The optimization procedure is separated into five distinct multiresolution levels(MRs) and is based on the photon optimizer (PO) algorithm (Version 16.0.1). To get to a convergent solution, the PO method uses fewer control points. Version 16.0.1 of the PO software was utilized to carry out inverse optimization for all planning strategies. The Eclipse treatment planning system (Version 16.0.14, Varian Medical System, Palo Alto, CA, USA) was utilized to build treatment plans for each patient for delivery with a Varian TrueBeam SVC linear accelerator, and the dose calculation was done using anisotropic analytical algorithm (AAA, 16.0.14).

# $\emph{Conventional RapidArc plan (RA<sub>c</sub>)}$

The two coplanar full-arc beams in this design are oriented 181° to 179° clockwise and 179° to 181° counterclockwise, with a collimator angle (CA) of 30° and 330°, respectively.

#### *Noncoplanar RapidArc plan*

This plan also contains two coplanar full arc and one noncoplanar partial arc beam with a range of 45° to 315° and a couch rotation





PTV: Planning target volume, OARs: Organs at risk, D<sub>mean</sub>: Average dose, Vxx Gy: Volume receiving xx Gy of dose,  $D_{\text{max}}$ : Maximum dose inside the organ

of  $315^{\circ} \pm 5^{\circ}$ , based on the spatial relationship between the PTV and OARs. The two coplanar full arcs have a CA of 3° or 357°, whereas the one noncoplanar partial arc has a CA of  $45^{\circ} \pm 10^{\circ}$ .

# *Hybrid RapidArc plan*

To generate hybrid arc plans for the treatment of EC, this study combines two different radiation techniques: 3D-CRT and RapidArc plans. Two steps are involved in preparing the hybrid arc plans.

# Step 1: (Three-dimensional conformal radiotherapy using a 10MV/15 MV photon photons)

The 3D–CRT approach makes use of a static anteroposterior beam (AP-0°) and a posteroanterior (PA-180°). To achieve a two-beam configuration that complies with the PTV with a margin of 0.5 cm, MLCs are added. 33% dose prescription is used to calculate the dose for the AP‑PA beam configuration, and the plan is normalized to the target mean.

#### Step 2: (RapidArc using 6 MV photon beam)

The RA method makes use of two complete arc beams. The two full arc beams are rotated from 179° to 181° and from 181° to 179°, respectively. For these arcs, the CAs are 30° and 330°, respectively. The full treatment prescription dose value is set. The procedure for optimization: hybrid arc plan optimization is performed using the 3D‑CRT (AP‑PA) plan from Step 1 as a base dose plan. After calculating the RA plan final dose, the beam arrangement is transferred to the two complete arc RapidArc plans from Step 2 after calculating the 3D-CRT dose distribution. The MUs of the AP-PA beam are matched to those from the initial 3D-CRT plan once the final dose calculation is completed. About one‑third of the treatment comes from the 3D‑CRT technique, and the remaining two-third uses the RapidArc approach. This means the Hybrid plan is composed of 33% and 67% of 3D‑CRT and RapidArc plans, respectively. All hybrid arc plans are normalized to deliver 100% dose coverage to 50% of the target volume (PTV). This normalization ensures that the PD is adequately delivered to the target area.

#### **Dosimetric analysis**

Cumulative dose‑volume histograms (c‑DVHs) based on the International Commission on Radiological Units and Measurements Report 83, 2010 were used to evaluate the dosimetric parameters of each plan. The planned objective was to provide 95% of the PD to 95% of the PTV volume, with a maximum dose (Dmax) of <110% of the PD and no more than 2% of the PTV volume receiving 107% of the PD. The quality of the plan was further assessed using the homogeneity index (HI), the conformity index (CI),<sup>[18]</sup> and the conformation number (CN);<sup>[18,19]</sup> uniformity index.<sup>[20]</sup>

CI is defined as:

 $CI = Volume of reference isodose (RI)/Volume of planning$ target volume (1)

CI is a measure of how well the PD conforms to the PTV, CI ideal value is one.

HI is defined as:

$$
HI = (D_{2\%} - D_{98\%})/D_{50\%}
$$
 (2)

This index represents the difference between the dose delivered to 2% of the target volume  $(D_{20})$  and that delivered to 98% of the target volume  $(D_{98\%})$  divided by 50% of the target volume  $(D_{50\%})$ . HI evaluates the dose homogeneity within the PTV. A value of zero is ideal and the closer the HI is to zero, the better the homogeneity.

CN is defined as:

$$
CN = (TV_{RI} / TV) \times (TV_{RI} / V_{RI})
$$
\n(3)

where, TV, TV $_{\text{RI}}$ , and V $_{\text{RI}}$  represents the treatment volume, the treatment volume at RI of the PD, and the total volume at RI of the PD. The RI was defined as 95% of PTV PD. The maximum value for CN is 1 which corresponds to perfect PTV coverage.

#### *Uniformity index*

It is defined as a ratio between minimum doses reached in 5% of the PTV volume (D 5%) and the minimum dose reached in 95% of the PTV volume (D 95%).

$$
Uniformity index (UI) = D5\%}/D95\%
$$
 (4)

#### *Integral dose*

It refers to the total amount of energy absorbed within an organ.<sup>[8]</sup> It is calculated using the mean organ dose  $(D_{mean})$ , mean organ volume ( $V_{\text{mean}}$ ), and mean organ density ( $\rho_{\text{mean}}$ ) with the formula:

$$
Integral dose (ID) = Dmean \times Vmean \times \rhomean (Gy-L)
$$
 (5)

In this study, all organs have the same density ( $\rho =1$ ), simplifying the calculation to:

$$
ID = D_{\text{mean} \times} V_{\text{mean}} \text{ (Gy-L)} \tag{6}
$$

#### **Statistical analysis**

The statistical difference between the dose–volume data for  $\text{RA}_{c}$ ,  $\text{RA}_{nc}$ , and  $\text{RA}_{Hyb}$  was determined using a one-way ANOVA test. The mean and standard deviation (SD) of the data were displayed. The three plans were compared using Wilcoxon signed-rank tests. A significance level of  $P < 0.05$ 

was considered statistically achievable. The Jamovi software 2.3.28 was used for all statistical analyses.

# **Results**

The patients' characteristics and demographics are shown in Table 1. Atotal of 14 patients were chosen for our study including 7 male and 7 female patients. The patients' ages ranged from 41 to 83 years. The PTVs long axis had an average length of  $19.99 \pm 3.57$  cm and a range of  $14.3 - 25.61$  cm. The volume of the PTVs varied from 260.2 and 898.1 cm<sup>3</sup>, averaging  $613.28 \pm 215$  cm<sup>3</sup>. Table 3 presents the target coverage analysis for three different treatment plans ( $\text{RA}_{\text{C}_1}\text{RA}_{\text{NC}}$ ,  $\text{RA}_{\text{Hyb}}$ : 3DCRT with RapidArc). Table 3 shows the mean and SDs for various DVH parameters related to the PTV. The *P*values were reported to indicate significant differences between delivery methods.

No statistically significant difference was found between the three delivery methods  $(RA_C, RA_{NC}$ , and  $RA_{Hvb}$ ) as the Dmax received by the PTV was 54.28 Gy, 54.79 Gy, and 54.42 Gy respectively (*P* > 0.05). D95% (dose received from 95% of PTV volume) was significantly higher in  $\text{RA}_{\text{Hyb}}$ , 96.85% compared to  $\text{RA}_{c}$ , 96.58% and  $\text{RA}_{nc}$ , 96.28%. There was no statistically significant difference in the dose received by 95% of the PTV volume among the three delivery methods  $(P > 0.05)$ . The CI of  $\text{RA}_{\text{Hvb}}$  showed a statistically significant improvement in conformity,  $0.983$  compared to  $RA_c$ ,  $0.977$  and  $RA_{nc}$ , 0.979 ( $P < 0.05$ ). The HI of RA<sub>Hyb</sub> demonstrated a statistically significant improvement in dose homogeneity, 0.079 within the PTV compared to  $\text{RA}_{c}$ , 0.082 and  $\text{RA}_{nc}$ , 0.088 ( $P < 0.05$ ). No statistically significant difference was observed in the UI between the three delivery methods  $(P > 0.05)$ . The UI values reported were  $1.064 \pm 0.013$  for RA<sub>c</sub>,  $1.068 \pm 0.011$  for  $\text{RA}_{\text{nc}}$ , and  $1.060 \pm 0.010$  for  $\text{RA}_{\text{Hvb}}$ . No statistically significant difference was observed in the CN between the three plan delivery methods RA<sub>c</sub> versus RA<sub>nc</sub> versus RA<sub>Hyb</sub>, 0.857 versus 0.921 versus  $0.846$  ( $P > 0.05$ ). However, the average of MUs per fraction was statistically lower in  $\text{RA}_{\text{Hvb}}$ , 452.7 compared to RA<sub>c</sub>, 517.5 and RA<sub>nc</sub>, 566.2 ( $P < 0.05$ ).

Table 4 presents the extracted parameters comparing OAR doses, specifically focusing on the lungs and heart, obtained from the

**Table 3: Average and standard deviation of the dosimetric and plan quality indices for the planning target volume and monitor unit for three different RapidArc planning schemes**

<b>PTV</b>	$RAn$ , mean $\pm$ SD	$RA_{n}$ , mean $\pm$ SD	$RA_{Hv}$ , mean $\pm SD$	$RA_{n}$ versus $RA_{n}$ (P)	$RA_{c}$ versus $RA_{Hv}$ (P)	$RA_{nc}$ versus $RA_{Hv}$ (P)
$D_{max}$	$54.28 \pm 1.163$	54.79±0.489	54.42±0.905	0.328	0.626	0.241
$V_{105\%}$ (cc)	$2.44 \pm 1.86$	$3.23 \pm 3.32$	$2.57 \pm 1.42$	0.246	0.411	0.815
$D_{95\%}$	$96.58 \pm 0.967$	$96.28 \pm 0.921$	$96.85 \pm 0.929$	0.296	0.173	0.017
HI	$0.082 \pm 0.017$	$0.088 \pm 0.014$	$0.079 \pm 0.014$	0.104	0.153	0.002
UI	$1.064 \pm 0.013$	$1.068 \pm 0.011$	$1.060 \pm 0.010$	0.622	0.678	0.190
<b>CI</b>	$0.979 \pm 0.015$	$0.977 \pm 0.018$	$0.983 \pm 0.016$	0.529	0.119	0.011
<b>CN</b>	$0.857 \pm 0.060$	$0.921 \pm 0.264$	$0.846 \pm 0.055$	0.576	0.119	0.241
MU	$517.5 \pm 84.43$	$566.2 \pm 131.7$	$452.7 \pm 63.15$	0.081	$0.002*$	< 0.001

RA<sub>c</sub>: Coplanar RapidArc, RA<sub>nc</sub>: Noncoplanar RapidArc, RA<sub>Hyb</sub>: Hybrid arc (three-dimensional conformal radiotherapy + Arc), D<sub>max</sub>: Maximum dose,  $D_{95\%}$ : Dose received by 95%, SD: Standard deviation, CI: Conformity index, HI: Homogeneity index, UI: Uniformity index, CN: Conformation, MU: Monitor unit, PTV: Planning target volume

c‑DVH as shown in Figure 1. Significantly V5 Gy of both lungs was 85.62% in RAHyb plan compared to RAc (89.15% )and RAnc(86.76%) , and no statistical difference was observed in RAnc versus RAHyb plan ( $P > 0.05$ ). V10 Gy, RA<sub>Hyb</sub> shows a statistically significant reduction of 60.85% compared to  $\text{RA}_c$ , 72% and RA<sub>ns</sub>, 64.14% ( $P < 0.01$ ). There was a significant difference between  $\text{RA}_{\text{c}}$  versus  $\text{RA}_{\text{nc}}$  (*P* value = 0.003) and RA<sub>nc</sub> vs. RA<sub>Hyb</sub> ( $P < 0.001$ ). No significant difference was observed between  $\text{RA}_{\text{nc}}$  and  $\text{RA}_{\text{Hvb}}$  ( $P = 0.217$ ). V20 Gy for each lung was statistically lower by 23.52% in the  $\text{RA}_{\text{Hub}}$ plan compared to the RA<sub>c</sub> plan, 25.51%; however, there was no statistically significant difference in V20 Gy when considering all three delivery methods together  $(P > 0.05)$ . V30 for both lung volumes showed a statistically significant increase of 11.53% in  $\text{RA}_{\text{Hyb}}$  plan compared to  $\text{RA}_{c}$ , 10.01% and RA<sub>nc</sub>, 9.94% ( $P < 0.001$ ). The mean of the RA<sub>Hyb</sub> plan resulted in a statistically significant reduction in the mean lung dose of 15.10 Gy compared to  $RA_c$ , 16.15 Gy and RA<sub>nc</sub>, 15.24 Gy with  $P < 0.01$ . The ID for the lungs showed



**Figure 1:** The dose–volume histogram comparison for planning target volume and various organs at risk between conventional arc (RAc), noncoplanar arc (RAnc), and hybrid (RA $_{\text{Hv}}$ ) Arc plans

a statistically insignificant increase in ascending order for  $\text{RA}_{\text{Hyb}}$  <  $\text{RA}_{\text{nc}}$  <  $\text{RA}_{\text{c}}$ .

Heart V30 Gy resulted in a statistically significant increase of 33.92 Gy in the  $\text{RA}_{\text{Hyb}}$  plan compared to  $\text{RA}_{c}$ , 27.82 Gy and  $\text{RA}_{nc}$ , 28.38 Gy (*P* < 0.001). The Heart V40 Gy showed a statistically significant increase of 18.85 Gy in the  $\text{RA}_{\text{Hvb}}$  plan compared to  $\text{RA}_{\text{c}}(17.31 \text{ Gy})$  and  $\text{RA}_{\text{nc}}(17.54 \text{ Gy})$  ( $P < 0.01$ ). No statistically significant difference in V40 Gy was observed between the  $\text{RA}_{\text{c}}$  and  $\text{RA}_{\text{nc}}$  plans ( $P > 0.05$ ). The mean heart dose in the  $RA<sub>Hyb</sub>$  plan demonstrated a statistically significant increase to 26.25 Gy compared to  $\text{RA}_{\text{c}}$  (23.91 Gy) and  $\text{RA}_{\text{nc}}$  (23.82 Gy)  $(P < 0.001)$ . There was no statistically significant difference in the mean heart dose between the  $RA<sub>c</sub>$  and  $RA<sub>nc</sub>$  plans  $(P > 0.05)$ . The ID for hearts was  $1.128 \pm 0.419 \times 10^4$  (Gy-L) for RA<sub>c</sub>,  $1.132 \pm 0.456 \times 10^4$  (Gy-L) for RAn<sub>c</sub>, and  $1.254 \pm 0.468 \times 10^4$  (Gy-L) for RA<sub>Hyb</sub>, respectively, with no statistically significant difference.

There was no statistically significant difference between the three delivery methods  $(RA_c, RA_{nc}, and RA_{Hyb})$  for the Dmax (44 Gy, 45.35 Gy, and 44.28 Gy, respectively) received by the spinal cord ( $P > 0.05$ ).

# **Discussion**

Dose escalation was found to improve survival in the RTOG94‑05 trial; however, the study was constrained by the investigator's use of standard 2DRT techniques, which resulted in greater doses being delivered to nearby OARs.[21] In EC, local failures and residual disease present persistent therapeutic problems that this study was intended to address. This study compared  $\text{RA}_{\text{Hyb}}$  plans with  $\text{RA}_{\text{c}}$  and  $\text{RA}_{\text{nc}}$  plans to examine the dosimetric characteristics of these treatment options for esophageal malignancies. Compared to RA<sub>c</sub> and  $RA_{nc}$  plans, the results showed that  $RA_{Hv}$  plans significantly

**Table 4: The comparison of different dose levels between conventional and noncoplanar and hybrid RapidArc plan for various organs at risk**

<b>OARs</b> <b>parameters</b>	RA, $mean \pm SD$	$RA_{nc}$ $mean \pm SD$	$RA_{Hv}$ $mean \pm SD$	RA, versus $RA_{nc}(P)$	RA, versus $RA_{Hyb} (P)$	RA <sub>nc</sub> versus $RA_{Hyb} (P)$
Both lung						
$V5$ Gy $(\% )$	$89.15 \pm 12.1$	$86.76 \pm 13.19$	$85.62 \pm 14.85$	0.032	0.014	0.625
V10(%)	$72 \pm 16.97$	$64.14 \pm 18.98$	$60.85 \pm 16.64$	0.003	< 0.001	0.217
$V20$ Gy $(\% )$	$25.51 \pm 9.66$	23.75±9.57	$23.52 \pm 10.01$	0.011	0.217	0.761
$V30$ Gy $(\% )$	$10.01 \pm 5.87$	$9.947 \pm 6.15$	$11.53\pm 6.06$	0.442	< 0.001	< 0.001
$D_{mean}$ (Gy)	$16.15 \pm 0.03$	$15.24 \pm 0.035$	$15.10 \pm 0.037$	0.004	0.011	
ID (Gy-L) $\times$ 10 <sup>4</sup>	$3.95 \pm 1.19$	$3.66 \pm 1.11$	$3.63 \pm 1.18$	0.822	0.789	0.997
Heart						
V30 Gy $(\% )$	$27.82 \pm 8.75$	$28.38 \pm 8.78$	$33.92 \pm 9.77$	0.442	< 0.001	< 0.001
$V40$ Gy $(\% )$	$17.31 \pm 6.39$	$17.54 \pm 6.62$	$18.85 \pm 6.68$	0.484	0.002	0.003
$D_{mean}$ (Gy)	$23.91 \pm 4.67$	$23.82 \pm 5.10$	$26.25 \pm 4.89$	0.625	< 0.001	< 0.001
ID (Gy-L) $\times$ 10 <sup>4</sup>	$1.128 \pm 0.419$	$1.132 \pm 0.456$	$1.254 \pm 0.468$	0.991	0.772	0.783
Spinal cord						
$D_{max}$ (Gy)	$44.0 \pm 5.47$	$45.4 \pm 3.12$	$44.28 \pm 4.15$	0.174	0.114	0.671

RA<sub>c</sub>: Coplanar RapidArc, RA<sub>nc</sub>: Noncoplanar RapidArc, RA<sub>Hyb</sub>: Hybrid arc (three-dimensional conformal radiotherapy + Arc), D<sub>mean</sub>: Mean dose, SD: Standard deviation, Vxx: XX Gy dose received by % of the volume, Gy-L: Gray-liter, ID: Integral dose

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**Figure** 2: The maximum dose to 20% color dose wash comparison between (a) conventional arc (RAc), (b) noncoplanar arc (RA<sub>nc</sub>), (c) hybrid (RAHyb) treatment plans for one of the patients

improved dose conformance, homogeneity, and PTV coverage as shown in Figure 2. However, as shown in Figure 3a,  $RA_{n}$ plans had a higher CN value than both  $\text{RA}_c$  and  $\text{RA}_{\text{Hyb}}$  plans, whereas the difference was not statistically significant.  $RA<sub>Hub</sub>$ shows the better uniformity of doses compared to  $\text{RA}_c$  and  $\text{RA}_{nc}$ treatment plans. In addition, from Figure 3b, average MUs of the Hybrid plan  $(RA<sub>Hvb</sub>)$  were significantly lower than those of the  $\text{RA}_{c}$  and  $\text{RA}_{nc}$  plans. Scatter dose is directly related to MU, resulting in higher MU in a treatment plan with more scattered dose to the surrounding healthy tissue. In our study, the  $RA_{Hv}$  plan shows the 25% and 14% less MU compared to  $\text{RA}_{\text{nc}}$  and  $\text{RA}_{\text{c}}$  treatment plans, respectively.

When evaluating treatment regimens for EC, pulmonary toxicity represents a significant problem that shows the need for dose restrictions for both lungs. Comparing the results of many studies is difficult since different, contradictory measures have been used for predicting pulmonary toxicity.  $[22,23]$  In this investigation, measures including V10 Gy, V20 Gy, and V30 Gy were used to assess pulmonary toxicity. According to M. V. Graham *et al*.,[24] a total lung V20 Gy of 25% indicates a minimal risk of pneumonitis. On the other hand, Asakura *et al*. [25] found that radiation pneumonitis during radiotherapy for EC is significantly more likely when the V20 Gy is higher than 37%. Furthermore, a correlation between Dmean values in the lungs and pneumonitis was found,[26] with suggested values ranging from 20 to 23 Gy. Tonison et al reported that their institutional data for esophageal cancer showed no patient with a V5 Gy greater than 71% developed grade 2 or higher pulmonary toxicity.<sup>[27]</sup> Milano data analysis revealed that the suggested ranges for Dmean and V30 Gy were  $10-20$  Gy and  $10\%$ -15%, respectively. Moreover, keeping V20 Gy below 25%–30% was associated with a lower risk of late Grade 3 toxicity, to  $\leq 5\% - 10\%$ .<sup>[28]</sup> As illustrated in Figure 4a, the lung dosimetric parameters in our investigation were within the limits indicated by the Milano data for toxicity evaluation.



**Figure 3:** (a) Shows the comparison of various plan quality indices and (b) shows the monitor unit for conventional arc (RAc), noncoplanar arc (RAnc), and hybrid ( $RA<sub>sub</sub>$ ) arc plans

Pneumonitis on RT for lung lesions is associated with a higher incidence of V5 Gy of  $>65\%$ , according to I.Y. Jo *et al.*<sup>[29]</sup> The maximum V5 Gy was achieved by the RA plan (89.15%), followed by the  $RA_{nc}$  (81.8%) and hybrid plan (85.62%). While  $\text{RA}_{\text{c}}$  and  $\text{RA}_{\text{nc}}$  planning techniques provide improved target dose painting capabilities, they also expose a large lung volume to a large number of low doses of radiation. Figure 5a shows the comparison of ID between  $\text{RA}_{c}$ ,  $\text{RA}_{\text{nc}}$ , and  $\text{RA}_{\text{Hyb}}$  for lungs.  $\text{RA}_{\text{Hyb}}$  plans exhibited the lowest ID values for the lungs. A 9.30% increase in ID was

observed for the lungs in  $RA_c$  plans compared to  $RA_{Hyb}$ treatment plans.

The significant incidence of radiation-induced cardiac damage at doses higher than 40 Gy has been reported in a number of studies, highlighting the significance of lowering V40 Gy to minimize cardiac toxicities. Strong correlations between V30 Gy exceeding 33% and Dmean exceeding 20 Gy and different grades of myocardial effusion in IMRT for EC were shown by Pao *et al.*[26] In addition, the results of their analysis showed that cardiac effusion Grade 3 or higher could be significantly predicted when V30 Gy exceeds 65% and V40 Gy exceeds



**Figure 4:** (a) Compares various dose–volume parameters for the lungs, and (b) compares the heart and spinal cord between conventional arc (RA<sub>c</sub>), noncoplanar arc (RA<sub>nc</sub>), and hybrid arc (RA<sub>Hyb</sub>) plans

55%. Wei *et al*. [30] reported that the retention of cardiac effusion following chemotherapy-radiation therapy for esophageal cancer was associated with V30 Gy of 46% and Dmean of less than 26 Gy. According to our analysis, the  $RA<sub>Hub</sub>$  exhibits higher mean doses to the heart compared to both the  $\text{RA}_\text{c}$  and  $\text{RA}_\text{nc}$ plans. In contrast to  $\text{RA}_{\text{c}}$  and  $\text{RA}_{\text{nc}}$  plans,  $\text{RA}_{\text{Hyb}}$  also results in a statistically significant increased V30 Gy and V40 Gy for the heart. This increase in heart doses is due to the inclusion of AP‑PA 3DCRT beam into the hybrid RA planning. In spite of this, in all plans, we have achieved a heart mean dose of less than 26 Gy, V30 Gy, and V40 Gy of <30%. In contrast to the  $RA<sub>c</sub>$  plans, the  $RA<sub>nc</sub>$  plans in our study resulted in somewhat better dose distribution, except for the heart and spinal cord. This is mainly due to the constraints imposed by the noncoplanar arc beams. For the spinal cord, the National Comprehensive Cancer Network Guidelines propose a dose limit of no more than 45 Gy. At total spinal cord doses of 50 Gy, 60 Gy, and 69 Gy, respectively, Kirkpatrick *et al*. [31] showed that myelopathy rates were 0.2%, 6%, and 50% with standard fraction doses of 2 Gy per day. As illustrated in Figure 4b, our investigation sustained a maximum spinal cord exposure of 45 Gy and found no statistically significant variations. Figure 5b shows the comparison of ID between  $RA_c$ ,  $RA_{nc}$ , and  $RA_{Hyb}$  for the heart. RA<sub>Hyb</sub> plans exhibited the highest ID values for the hearts among  $\overline{RA}_c$  and  $\overline{RA}_{nc}$ . A 12.51% increase in ID was observed for the hearts in  $\text{RA}_{\text{Hyb}}$  plans compared to  $\text{RA}_{\text{c}}$  treatment plans.

# **Conclusion**

Compared to  $\mathsf{RA}_{\mathrm{c}}$  and  $\mathsf{RA}_{\mathrm{nc}}$  plans, hybrid arc plans demonstrated better dose coverage, homogeneity, lower mean lung doses, and lung volume reduction at low doses such as V5 Gy, V10 Gy, and V20 Gy. In addition, each fraction uses fewer MUs. Based on these findings, hybrid arc planning can potentially reduce radiation toxicity and improve treatment outcomes for EC. Complete integration of the hybrid arc plan into clinical practice requires additional study and clinical validation.

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**Figure 5:** (a) Compares the integral dose distribution for the whole lung, whereas (b) compares the integral dose distribution for the heart between conventional arc (RAc), noncoplanar arc (RAnc), and hybrid arc (RAHyb) plans

#### **Conflicts of interest**

There are no conflicts of interest.

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