

1 **Supplementary Material:**

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3 **Supplementary Table S1: List of Radiomic Features**

Feature Category	Features
First Order Statistics	Energy Total Energy Entropy Minimum 10th Percentile 90th Percentile Maximum Mean Median Interquartile Range Range Mean Absolute Deviation Robust Mean Absolute Deviation Root Mean Squared Standard Deviation Skewness Kurtosis Variance Uniformity
3D Shape	Mesh Volume Voxel Volume Surface Area Surface Area to Volume Ratio Sphericity Compactness Spherical Disproportion Maximum 3D Diameter Maximum 2D Diameter (Axial) Maximum 2D Diameter (Coronal) Maximum 2D Diameter (Sagittal)

	Major Axis Length Minor Axis Length Least Axis Length Elongation Flatness
Gray level Co-occurrence Matrix	Autocorrelation Joint Average Cluster Prominence Cluster Shade Cluster Tendency Contrast Correlation Difference Average Difference Entropy Difference Variance Difference Average Joint Energy Joint Entropy Informational Correlation Inverse Difference Moment Inverse Difference Moment Normalized Inverse Difference Inverse Difference Normalized Inverse Variance Maximum Probability Sum Average Sum Entropy Sum of Squares
Gray Level Size Zone Matrix	Small Area Emphasis Large Area Emphasis Gray Level Non-Uniformity Gray Level Non-Uniformity Normalized Size-Zone Non-Uniformity Size-Zone Non-Uniformity Normalized

	Zone Percentage Gray Level Variance Zone Variance Zone Entropy Low Gray Level Zone Emphasis High Gray Level Zone Emphasis Small Area Low Gray Level Emphasis Small Area High Gray Level Emphasis Large Area Low Gray Level Emphasis Large Area High Gray Level Emphasis
Gray Level Run Length Matrix	Short Run Emphasis Long Run Emphasis Gray Level Non-Uniformity Gray Level Non-Uniformity Normalized Run Length Non-Uniformity Run Length Non-Uniformity Normalized Run Percentage Gray Level Variance Run Variance Run Entropy Low Gray Level Run Emphasis High Gray Level Run Emphasis Short Run Low Gray Level Emphasis Short Run High Gray Level Emphasis Long Run Low Gray Level Emphasis Long Run High Gray Level Emphasis
Gray Level Dependence Matrix	Small Dependence Emphasis Large Dependence Emphasis Gray Level Non-Uniformity Dependence Non-Uniformity Dependence Non-Uniformity Normalized Gray Level Variance Dependence Variance Dependence Entropy

	Low Gray Level Emphasis High Gray Level Emphasis Small Dependence Low Gray Level Emphasis Small Dependence High Gray Level Emphasis Large Dependence Low Gray Level Emphasis Large Dependence High Gray Level Emphasis
Neighboring Gray Tone Difference Matrix	Coarseness Contrast Busyness Complexity Strength

Supplementary Table S1: A list of radiomic features extracted from a liver volume. The features include computations related to the statistics, shape, and gray-level relationships of the image.

8 **Supplementary Table S2: Radiomic Deviations from IBSI Standards**

Computation	PyRadiomics Implementation	IBSI Guidelines
Binning	Discretizes gray values with fixed bins with edges equally spaced from 0.	Discretizes using fixed bin width equally spaced from minimum of resegmentation range
Resampling	Aligns to the corner of the original voxel	Aligns to the center of the image
Gray value rounding	Does not implement resampling of similar resolution to original intensity values, with the argument that differences are likely to be minor and may add complexity.	Resamples to similar resolution of original CT image i.e. rounding to integer resolution of Hounsfield Units from the original CT intensity data.
Mask resampling	Resamples to nearest neighbor	Allows selection of different interpolators for resampling

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10 Supplementary Table S2: A list of deviations from the feature extraction guidelines by the Image
11 Biomarker Standardisation Initiative (IBSI).

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Supplementary Equation S3: Random Survival Forest Algorithm

To build a survival tree that predicts survival from an input vector of radiomic features, the following steps are taken:

1. Select N samples from the dataset.
2. For each sample $i = 1, 2, \dots, N$, initialize a binary decision tree with max depth D .
3. At each node, iterate through set of features $X = \{x_1, x_2, \dots, x_N\}$ and its range of feature values $S = \{S_{min}, S_{max}\}$ to select feature x_i and a threshold split value s_i such that:

$$L(x_i, s_i) \geq L(x, s) \quad \forall x \in X, s \in S$$

Where $L(x, s)$ is the log rank test such that

$$L(x, c) = \frac{\sum_{i=1}^N E_{i,1} - Y_{i,1} \frac{E_i}{Y_i}}{\sqrt{\sum_{i=1}^N \left(\frac{Y_{i,1}}{Y_i} \right) \left(1 - \frac{Y_{i,1}}{Y_i} \right) \left(\frac{Y_i - E_i}{Y_i - 1} \right) E_i}}$$

Where at time t_i , E_i is the number of events at time t_i , $E_{i,j}$ is the number of events at a daughter node j , Y_i is the number of patients with an events or at risk at time t_i , and $Y_{i,j}$ is the number of patients with an event or at risk at a daughter node j

4. Continue to grow children nodes unless the children node has no more than M surviving samples, where M is a user-defined hyperparameter
5. Calculate the cumulative hazard function for the decision tree with the Nelson-Aalen estimator:

$$H_q(t) = \sum_{t_{p,q} \leq t} \frac{E_{p,q}}{Y_{p,q}}$$

Where p is a patient in the set of M patients in set $P = \{p_1, p_2, \dots, p_M\}$, q is a node in the set of N nodes in set $Q = \{q_1, q_2, \dots, q_N\}$, $E_{p,q}$ is the number of events at time $t_{p,q}$, and $Y_{p,q}$ is the number of patients with an event or at risk at time $t_{p,q}$.

6. Repeat steps 1-5 K times to create K separately initialized trees, where K is a user-defined hyperparameter.
7. Average the cumulative hazard function over all trees to compute the ensembled cumulative hazard.

40 **Supplementary Table S4: Baseline variable distributions by lesion**

Characteristics	All lesions (n=129)
Number of patients/lesions	97/129
Sex, n (%)	
Male	83 (64.3)
Female	46 (36.7)
Metastasis at time of diagnosis, n (%)	
M0	40 (30.0)
M1	89 (70.0)
Colorectal Histology, n (%)	
Colon	104 (80.6)
Rectal	20 (15.6)
Undetermined	5 (3.8)
Number of liver lesions at diagnosis, n (%)	
0	5 (3.9)
1	25 (19.4)
2	9 (7.0)
3-5	32 (24.8)
> 5	53 (41.0)
Undetermined	5 (3.9)
Other sites at diagnosis, n (%)	
None	101 (78.3)
Lung	12 (9.3)
Non-regional LN	3 (2.3)
Lung and non-regional LN	4 (3.1)
Other	7 (5.4)
Undetermined	2 (1.6)
RT to other sites, n (%)	
No	75 (58.1)
Before liver RT	28 (21.7)
After Liver RT	21 (16.3)
Before and after liver RT	2 (1.6)
Undetermined	3 (2.3)
RT fraction delivered, Median (IQR)	6 (5-15)
RT dose delivered, Median (IQR)	4500 (3000 - 6000)
Dose Painting - Yes, n (%)	55 (42.6)
Intended Dose Median (IQR)	6000 (4000 - 6750)
Mean RT length \pm SD (Days)	11.6 (8.5)
PTV volume (cm ³), Median (IQR)	94.4 (39.2 - 174.4)

Mean D95 \pm SD (% of intended dose)	97.7 (11.0)
Reirradiation - Yes, n (%)	8 (6.2)
Surgery before RT, n (%)	91 (70.5)
Systemic before RT, n (%)	126 (97.7)
Pump before RT, n (%)	81 (62.8)
Lines of Chemo, Median (IQR)	3 (2 - 4)
RFA before RT, n (%)	45 (34.9)
RFA to RT lesions - Yes, n (%)	13 (10.1)
Y90 before RT - Yes, n (%)	10 (7.8)
Embolization before RT, n (%)	12 (9.3)
CEA at diagnosis, Median (IQR)	15.7 (3.38 - 176.9)
CEA at RT, Median (IQR)	18.7 (4.8 - 127.2)
Number of liver lesions at RT, n (%)	
1	57 (44.2)
2	43 (33.3)
3	12 (9.4)
≥ 4	16 (12.4)
Undetermined	1 (0.7)
Other sites at RT, n (%)	
None	52 (40.3)
Lung	27 (21.0)
Non-regional LN	10 (7.8)
Lung and non-regional LN	25 (19.3)
Other	15 (11.6)
Mean lesion 1 dimension 2 \pm SD	35.2 (22.3)
Mean lesion 1 dimension 1 \pm SD	26.0 (17.9)
Freedom from local progression (FFLP), n (%)	
Progression	55 (42.6)
No progression	67 (52.0)
Undetermined	7 (5.4)
Mean FFLP (months) \pm SD	10.5 (0.4)
Any hepatic progression (AHP), n (%)	
Progression	99 (76.8)
No progression	25 (19.4)
Undetermined	5 (3.8)
Mean time to AHP (months) \pm SD	7.3 (7.1)

Abbreviations: LN = lymph node, RT = radiotherapy, PTV = planning target volume, CEA = carcinoembryonic antigen, HAIP = hepatic arterial infusion pump, TARE = transarterial radioembolization.

Supplementary Table S4: A table of baseline clinical variables recorded as part of standard of care, with averages computed from the set of variables per lesion. The clinical variables will be utilized alongside computational radiomic features from computed tomography scans as input data to a machine learning model to predict local progression.

Supplementary Table S5: Baseline variable distributions by patient

Characteristics	All Patients (n=97)
Sex, n (%)	
Male	63 (64.9)
Female	34 (35.1)
Metastasis at time of diagnosis, n (%)	
M0	32 (33)
M1	65 (67)
Other sites at diagnosis, n (%)	
None	74 (76.3)
Lung	9 (9.3)
Non-regional LN	3 (3.1)
Lung and non-regional LN	4 (4.1)
Other	5 (5.2)
Undetermined	2 (2.0)
RT to other sites, n (%)	
No	58 (59.8)
Before liver RT	23 (23.7)
After Liver RT	13 (13.4)
Before and after liver RT	2 (2.1)
Undetermined	1 (1)
Number of liver lesions at RT, n (%)	
1	56 (57.7)
2	25 (25.8)
3	6 (6.2)
≥ 4	9 (9.3)
Undetermined	1 (1)
Other sites at RT, n (%)	
None	37 (38.1)
Lung	21 (21.7)
Non-regional LN	8 (8.3)
Lung and non-regional LN	17 (17.5)
Other	14 (14.4)
Freedom from local progression (FFLP), n (%)	
Progression	50 (51.6)
No progression	40 (41.2)
Undetermined	7 (7.2)
Mean time to local progression (months) ± SD	10.5 (8.8)

Any hepatic progression (AHP), n (%)	
Progression	76 (78.4)
No progression	16 (16.4)
Undetermined	5 (5.2)
Mean time to AHP (months) \pm SD	7.4 (6.9)

Abbreviations: LN = lymph node, RT = radiotherapy

Supplementary Table S5: A table of baseline clinical variables recorded as part of standard of care, with averages computed from the set of variables per patient. Lesion-specific variables were excluded.

60 **Supplementary Table S6: Dosages and number of fractions to liver metastases**

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Total Dose (Gy)	Fractions	BED₁₀ (Gy)	Patient Count
24	1	82	9
24	3	43	1
27	3	51	1
30	3	60	1
30	5	48	8
30	10	39	1
35	3	60	2
36	6	68	1
38	15	48	1
40	5	72	4
45	3	113	5
45	5	86	1
50	5	100	12
50	10	75	1
60	3	180	2
60	5	132	4
60	6	120	2
60	10	96	5
60	15	84	1
67.5	15	98	15
70	10	119	9
75	3	263	1

75	5	188	2
75	15	113	1
75	25	98	5
75	50	86	1
80	10	144	1

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63 Supplementary Table S6: A list of doses to liver metastases, fractions, biologically effective dose (BED),
64 and number of patients treated with the combination.
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66 **Supplementary Material S7: Link to survival models**

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68 The survival models from the different feature sets are uploaded to an open repository at:

69 https://github.com/ricky-hu/local_control_radiomics_survival_model

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