



# Effect of pre-processing on radiomics features estimation from computed tomography imaging in patients with rectal cancer.

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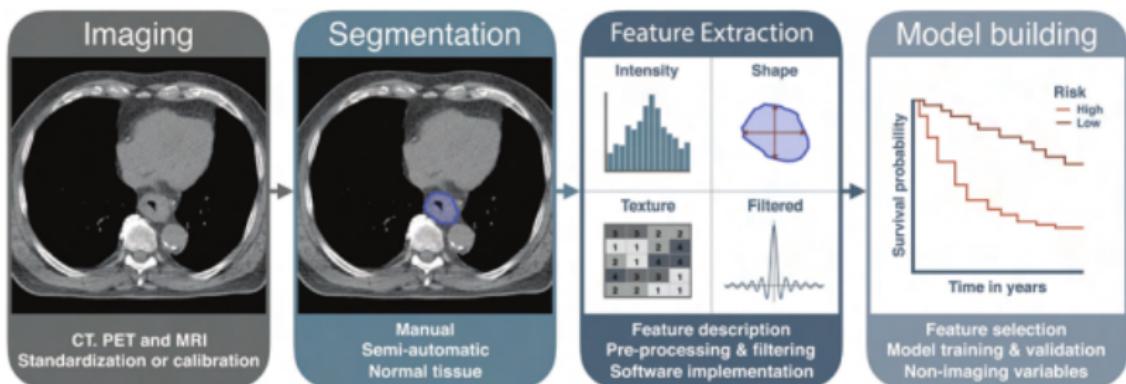
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# Radiomics: Images Are More than Pictures, They Are Data

"high-throughput extraction of **quantitative features** that result in the conversion of images into *mineable data* and the subsequent analysis of these data for decision support." <sup>a</sup>

<sup>a</sup> Gillies RJ et al. Radiomics: Images Are More than Pictures, They Are Data. doi:10.1148/radiol.2015151169

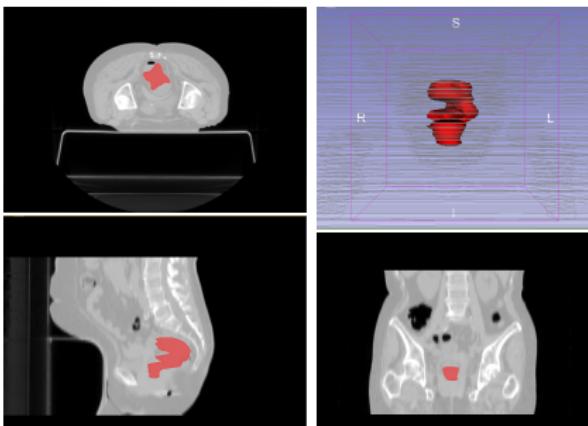


Deng, J. (Ed.), Xing, L. Big Data in Radiation Oncology. doi:10.1201/9781315207582

# Images:

Computed tomography images of 20 representative patients with **rectal cancer**, undergoing preoperative radiotherapy.

- Gross tumor volume manually **segmented** by a radiation oncologist



- CT acquisitions:

|                  |                    |
|------------------|--------------------|
| Scanner          | GE LightSpeed RT16 |
| Acquisition type | Helical            |
| Tube voltage     | 120 kVp            |
| Tube load        | 140-170 mAs        |
| Slice thickness  | 5 mm               |
| Pixel size       | 0.82-1.27 mm       |
| Matrix           | 512×512            |

- 105 **features** divided into 7 **classes**:

- 14 shape
- 18 first-order
- 22 Gray Level Co-occurrence Matrix (glcm)
- 14 Gray Level Dependence Matrix (gldm)
- 16 Gray Level Run Length Matrix (glrlm)
- 16 Gray Level Size Zone Matrix (glszm)
- 5 Neighbouring Gray Tone Difference Matrix (ngtdm)

# Features estimation with varying image pre-processing:

- Resampling:
  - 6 isotropic **voxel sizes**: 1, 1.3, 1.6, 1.9, 2.2, 2.5 mm
  - 10 **interpolation algorithms**: BSpline (BS), BlackmanWindowedSinc (BL), CosineWindowedSinc (CWS), Gaussian (G), HammingWindowedSinc (HWS), LabelGaussian (LG), LanczosWindowedSinc (LWS), Linear (L), NearestNeighbor (NN), WelchWindowedSinc (WWS)
- Quantization of gray-levels:
  - 6 **binwidths**: 3, 4, 5, 6, 7, 8 HU

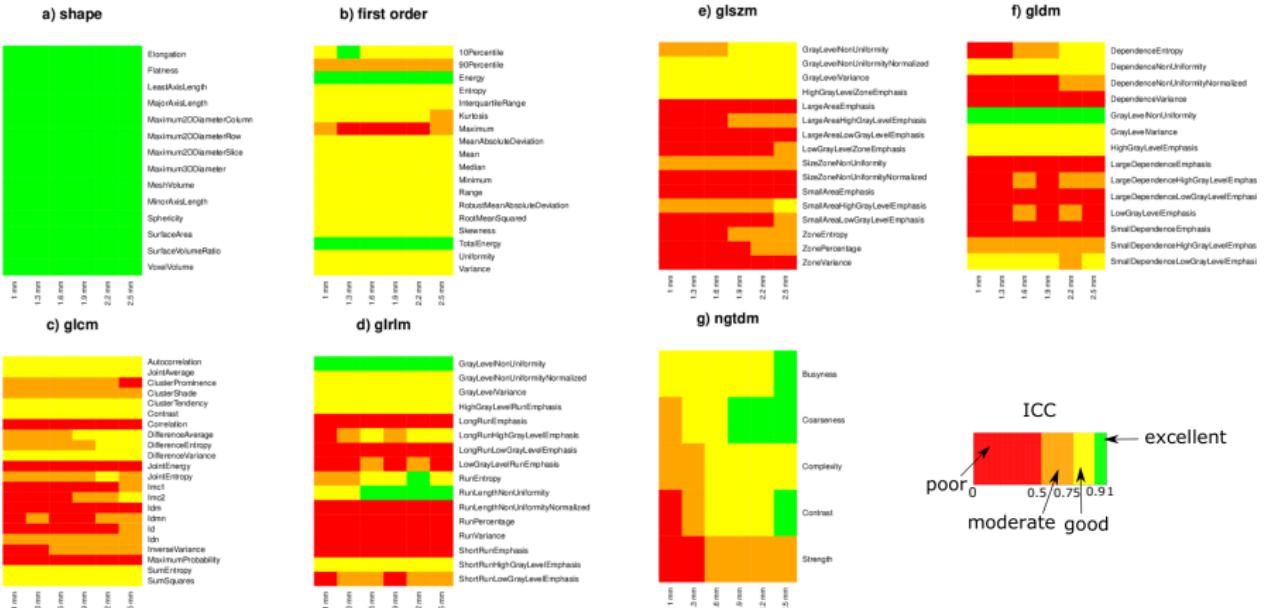
## 4 analyses:

- ① Effect of **interpolation algorithm** for different voxel sizes
- ② Effect of **binwidth** for different voxel sizes
- ③ Effect of **voxel size** for different binwidths
- ④ Effect of **voxel size** for different interpolation algorithms

### Statistical analysis

- **Intraclass Correlation Coefficient (ICC):** two-way mixed effects model, single rater, absolute agreement
- **Coefficient of Variation**
$$COV = \frac{\sigma}{|\mu|}$$

# 1) ICC values related to the effect of interpolation algorithm for different voxel sizes



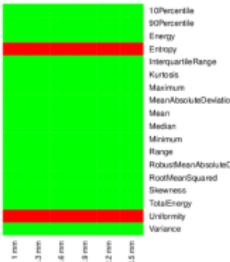
Features are divided into 4 reliability classes (poor, moderate, good, excellent) according to ICC values.

## 2) ICC values related to the effect of binwidth for different voxel sizes

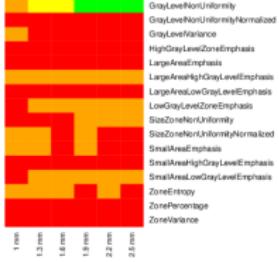
a) shape



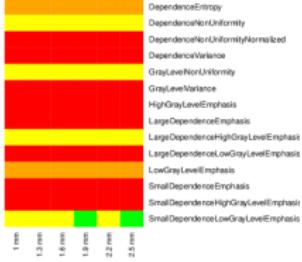
b) first order



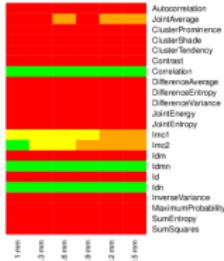
e) glszm



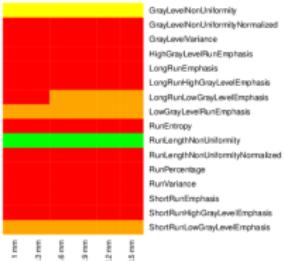
f) gldm



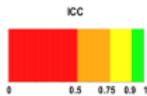
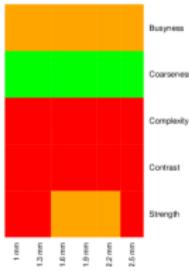
c) glcm



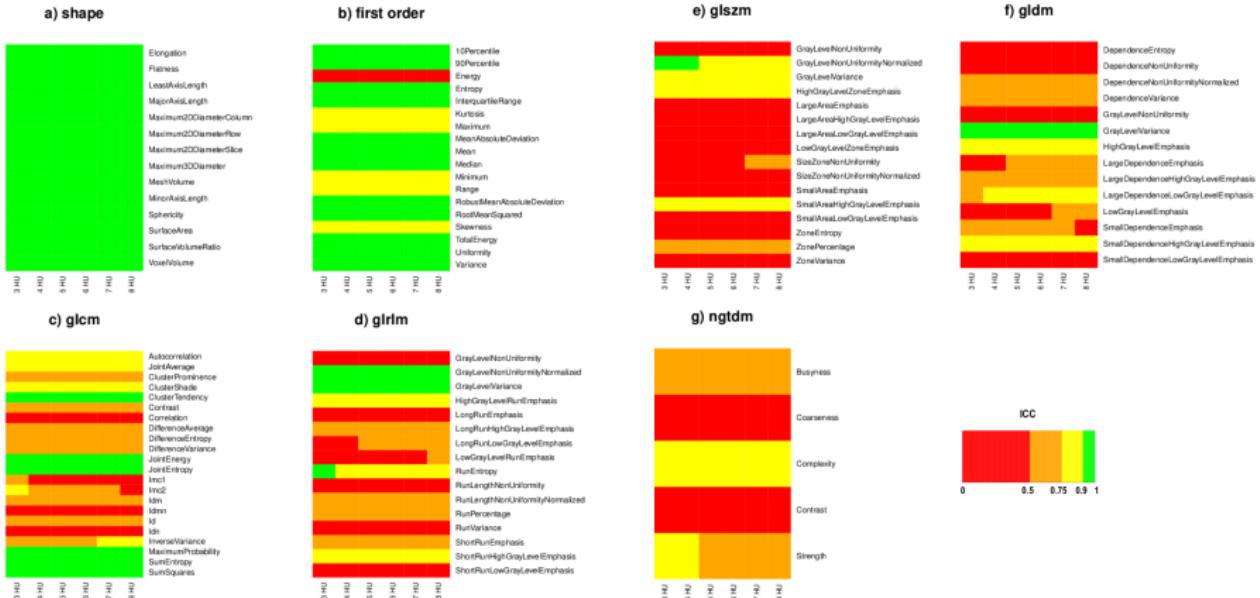
d) glrlm



g) ngtdm

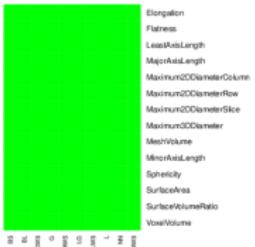


### 3) ICC values related to the effect of voxel size for different binwidths

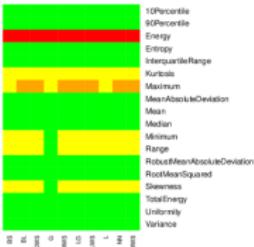


# 4) ICC values related to the effect of voxel size for different interpolation algorithms

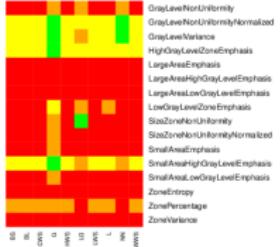
a) shape



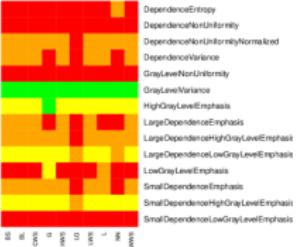
b) first order



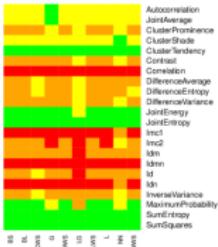
e) glsm



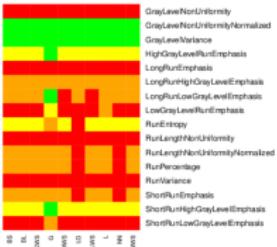
f) gldm



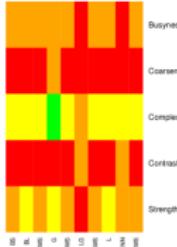
c) glcm



d) glrlm



g) ngldm



ICC



# Radiomic features median COV

| shape                   | 1   | 2   | 3   | 4   |
|-------------------------|-----|-----|-----|-----|
| Elongation              | 0,0 | 0,0 | 1,2 | 1,2 |
| Flatness                | 0,0 | 0,0 | 1,3 | 1,3 |
| LeastAxisLength         | 0,0 | 0,0 | 1,0 | 1,0 |
| MajorAxisLength         | 0,0 | 0,0 | 0,6 | 0,6 |
| Maximum2DDiameterColumn | 0,0 | 0,0 | 1,1 | 1,1 |
| Maximum2DDiameterRow    | 0,0 | 0,0 | 0,8 | 0,8 |
| Maximum2DDiameterSlice  | 0,0 | 0,0 | 0,9 | 0,9 |
| Maximum3DDiameter       | 0,0 | 0,0 | 0,8 | 0,8 |
| MeshVolume              | 0,0 | 0,0 | 1,7 | 1,7 |
| MinorAxisLength         | 0,0 | 0,0 | 0,6 | 0,6 |
| Sphericity              | 0,0 | 0,0 | 2,8 | 2,8 |
| SurfaceArea             | 0,0 | 0,0 | 3,1 | 3,1 |
| SurfaceVolumeRatio      | 0,0 | 0,0 | 2,8 | 2,8 |
| VoxelVolume             | 0,0 | 0,0 | 1,6 | 1,6 |

| first order                 | 1    | 2    | 3    | 4    |
|-----------------------------|------|------|------|------|
| 10Percentile                | 50,5 | 0,0  | 9,8  | 9,8  |
| 90Percentile                | 6,5  | 0,0  | 0,9  | 0,8  |
| Energy                      | 0,6  | 0,0  | 97,1 | 97,1 |
| Entropy                     | 3,1  | 11,8 | 0,6  | 0,6  |
| InterquartileRange          | 8,4  | 0,0  | 2,3  | 2,4  |
| Kurtosis                    | 23,2 | 0,0  | 21,8 | 18,8 |
| Maximum                     | 18,3 | 0,0  | 6,5  | 8,3  |
| MeanAbsoluteDeviation       | 6,4  | 0,0  | 2,1  | 1,9  |
| Mean                        | 11,0 | 0,0  | 1,4  | 1,4  |
| Median                      | 6,8  | 0,0  | 1,0  | 1,2  |
| Minimum                     | 20,1 | 0,0  | 20,0 | 18,0 |
| Range                       | 16,3 | 0,0  | 14,8 | 14,3 |
| RobustMeanAbsoluteDeviation | 7,7  | 0,0  | 1,7  | 1,7  |
| RootMeanSquared             | 0,3  | 0,0  | 0,0  | 0,0  |
| Skewness                    | 15,5 | 0,0  | 12,3 | 10,3 |
| TotalEnergy                 | 0,6  | 0,0  | 1,5  | 1,5  |
| Uniformity                  | 10,0 | 30,6 | 1,3  | 1,4  |
| Variance                    | 14,6 | 0,0  | 7,8  | 7,0  |

## Note:

- 1) Effect of interpolation algorithm
- 2) Effect of binwidth
- 3) Effect of voxel size
- 4) Effect of voxel size

| glm                | 1    | 2     | 3    | 4    |
|--------------------|------|-------|------|------|
| Autocorrelation    | 27,9 | 69,8  | 31,2 | 30,4 |
| JointAverage       | 16,0 | 34,4  | 16,5 | 15,8 |
| ClusterProminence  | 36,2 | 132,6 | 40,6 | 39,2 |
| ClusterShade       | 26,6 | 104,4 | 30,4 | 27,2 |
| ClusterTendency    | 12,6 | 70,8  | 13,2 | 11,9 |
| Contrast           | 22,0 | 70,5  | 25,6 | 24,9 |
| Correlation        | 20,7 | 0,4   | 29,8 | 28,1 |
| DifferenceAverage  | 14,5 | 35,0  | 12,7 | 12,6 |
| DifferenceEntropy  | 6,5  | 14,0  | 5,6  | 5,5  |
| DifferenceVariance | 22,1 | 70,1  | 27,1 | 26,1 |
| JointEnergy        | 39,8 | 55,0  | 7,7  | 8,3  |
| JointEntropy       | 4,8  | 12,3  | 1,3  | 1,2  |
| Imc1               | 36,1 | 7,3   | 24,7 | 23,8 |
| Imc2               | 8,3  | 6,0   | 7,4  | 7,0  |
| Idm                | 17,7 | 23,0  | 9,3  | 9,9  |
| Idmn               | 0,1  | 0,0   | 0,2  | 0,2  |
| Idn                | 12,2 | 16,2  | 6,4  | 6,8  |
| Indn               | 0,6  | 0,0   | 1,0  | 1,0  |
| InverseVariance    | 11,9 | 20,1  | 6,8  | 6,8  |
| MaximumProbability | 46,2 | 52,2  | 7,8  | 8,7  |
| SumEntropy         | 2,4  | 10,1  | 1,6  | 1,5  |
| SumSquares         | 14,1 | 70,7  | 7,2  | 6,7  |

| glm                                  | 1    | 2     | 3    | 4    |
|--------------------------------------|------|-------|------|------|
| DependenceEntropy                    | 2,7  | 2,5   | 3,6  | 3,7  |
| DependenceNonUniformity              | 20,6 | 26,8  | 84,8 | 83,8 |
| DependenceNonUniformityNormalized    | 20,6 | 26,8  | 12,7 | 13,6 |
| DependenceVariance                   | 62,8 | 40,9  | 21,7 | 24,2 |
| GrayLevelNonUniformity               | 10,0 | 30,8  | 97,1 | 97,0 |
| GrayLevelVariance                    | 14,5 | 70,8  | 7,8  | 7,1  |
| HighGrayLevelEmphasis                | 28,0 | 69,8  | 31,4 | 30,4 |
| LargeDependenceEmphasis              | 57,1 | 42,4  | 26,4 | 27,9 |
| LargeDependenceHighGrayLevelEmphasis | 55,1 | 19,6  | 47,5 | 50,2 |
| LargeDependenceLowGrayLevelEmphasis  | 98,0 | 84,3  | 25,2 | 25,8 |
| LowGrayLevelEmphasis                 | 45,8 | 40,2  | 51,0 | 49,1 |
| SmallDependenceEmphasis              | 31,8 | 33,3  | 22,9 | 23,1 |
| SmallDependenceHighGrayLevelEmphasis | 46,1 | 102,4 | 31,4 | 31,4 |
| SmallDependenceLowGrayLevelEmphasis  | 23,6 | 14,3  | 67,9 | 67,8 |

| ngldm      | 1    | 2    | 3    | 4    |
|------------|------|------|------|------|
| Busyness   | 46,2 | 50,6 | 51,7 | 51,8 |
| Courtness  | 22,8 | 0,5  | 68,1 | 68,5 |
| Complexity | 34,2 | 89,8 | 25,9 | 22,4 |
| Contrast   | 23,1 | 48,3 | 49,7 | 48,5 |
| Strength   | 37,2 | 51,8 | 31,8 | 32,0 |

# Conclusion



The estimation of radiomics features in rectal cancer from CT imaging appreciably **depends on pre-processing**, in terms of *resampling voxel size, interpolation algorithm and quantization binwidth*.

- **ICC analysis** shows that more than half of the radiomic features have *poor* or *moderate* reliability,
- A number of features with good ICC values ( $\text{ICC} > 0.75$ ) presents a **median COV**  $> 25\%$

Image pre-processing effect should be taken into account when planning a clinical study, comparing results from different studies and performing multicentric studies.