

The role of delta radiomics in gastric cancer

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Submitted Jul 20, 2018. Accepted for publication Jul 27, 2018.

doi: 10.21037/qims.2018.07.08

View this article at: <http://dx.doi.org/10.21037/qims.2018.07.08>

We have read with great interest and appreciated the paper by Hou *et al.* entitled “Radiomic analysis using contrast-enhanced computed tomography (CT): predict treatment response to pulsed low dose rate radiotherapy in gastric carcinoma with abdominal cavity metastasis” (1).

The authors successfully tried to correlate radiomic features, calculated on three-dimensional 3D VOI'S drawn slightly within the border of the tumoral lesions, with therapeutic response assessed according to the response evaluation criteria in solid tumors (RECIST v1.1). In particular, they found that 6 features (1 first order-based, 1 texture-based, 1 log-based, and 3 wavelet-based) were significantly different between responders (patients with complete or partial response) and non-responders (patients with stable or progressive disease). These findings suggest that radiomic analysis of contrast-enhanced CT images may provide additional information regarding therapeutic response, which could be important for the clinical decision-making process.

In fact, texture analysis (TA) of CT images seems to be able to detect subtle differences in CT values which cannot be recognized by human eyes, providing quantitative data on tumour phenotype and microenvironment by analyzing the relation, distribution and relationship of pixel intensities (2).

Another therapeutic option for metastatic gastric cancer (GC), which has spread during the last years, is conversion surgery, defined as a surgical treatment aiming at R0 resection after chemotherapy in patients with originally unresectable or only marginally resectable advanced gastric cancer (AGC) (3). For this reason, neoadjuvant

chemotherapy (NAC) still remains a standard of care both for metastatic and locally AGC and an accurate evaluation of overall tumour burden and discrimination between responders and non-responders, prior to surgery, has than become increasingly important to customize patient treatment and reduce healthcare costs, delivering aggressive multimodal treatments only to patients who potentially might benefit from these (4,5).

In this sense it is rising interest to identify surrogate imaging markers for predicting tumour histopathological regression prior to surgery and CT studies have shown promising results for improving patient stratification, by using different approach, such as linear or volumetric measurement systems or radiomic analysis (6-8).

In our experience, delta radiomics, defined as the calculation of the modification of texture parameters before and after NAC, has shown the potentiality to predict histopathological response and patient outcome, overcoming the limit of a model based only on pre or post-treatment radiomic features (9). We have preliminary analyzed a homogeneous cohort of 23 patients with biopsy proven resectable AGC ($\geq T3$ or N+), treated with NAC and radical surgery. Gastric lesion, defined as gross tumour volume (GTV) was contoured on every slice of pre and post NAC contrast enhanced CT scan, obtaining a 3D region of interest (3D-ROI) with the contouring software RayStation, using a method already validated in different settings (10). Finally, a correlation between the texture parameters [histogram, shape and grey-level co-occurrence matrix (GLCM)] extracted from the 3D-ROI and the complete

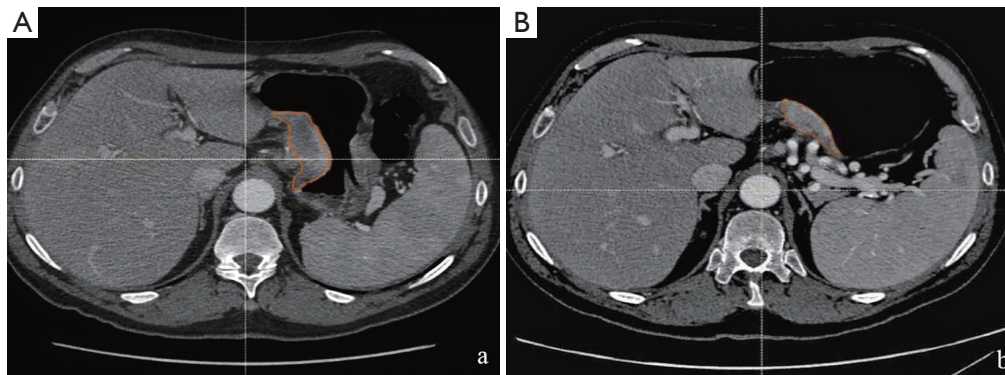


Figure 1 A 67-year-old man with advanced gastric cancer of the lesser curvature who underwent contrast enhanced CT pre (A) and post (B) NAC. Although the post-NAC CT clearly shows a reduction in the size of tumoral mass, delta parameters (entropy, GLCM-contrast, GLCM-entropy, GLCM-dissimilarity) classify this patient as non-responder. The finding is confirmed on final histopathology after surgery (adenocarcinoma ypT3, tumor regression grade 3 according to Becker *et al.*). NAC, neoadjuvant chemotherapy; CT, computed tomography; GLCM, grey-level co-occurrence matrix.

pathological response (ypCR: tumour regression grade 1 according to Becker *et al.*) was searched. In our preliminary case population only three patients showed a ypCR, and only delta parameters entropy ($P=0.003$), GLCM-contrast ($P<0.001$), GLCM-entropy ($P=0.004$), GLCM-dissimilarity ($P=0.001$) were correlated with this outcome at univariate analysis whereas at multivariate analysis, the only parameter that showed a significant correlation was the delta GLCM-contrast ($R^2: 0.539$, sensibility 66%, specificity 100%, AUC 0.733) (Figure 1).

We are going to test these results with external validation, from a wider cohort of about 100 patients coming from different centres of the Italian Research Group for Gastric Cancer (GIRCG).

Acknowledgements

None.

Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

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Cite this article as: Mazzei MA, Nardone V, Di Giacomo L, Bagnacci G, Gentili F, Tini P, Marrelli D, Volterrani L. The role of delta radiomics in gastric cancer. *Quant Imaging Med Surg* 2018;8(7):719-721. doi: 10.21037/qims.2018.07.08