Supplementary File for: MedFusionGAN: Multimodal Medical Image Fusion Using an Unsupervised Deep Generative Adversarial Network

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I. Quantitative metrics

The brief definition of the metrics are as follows:

1. Entropy (H): This metric measured in bits provides the amount of information required to code a given image. The higher the image entropy is, the better the image fusion result is. It is defined as follows:

$$H(X) = -\sum_{x} p(x) \log_2(x) \tag{1}$$

where p(x) is probability image intensity x for a give image X.

2. Standard deviation (SD): SD given in Equation 2 is a statistical measure that provides information about image contrast. For a noiseless image, an higher SD means a better image contrast.

$$SD = \sqrt{\mathbb{E}\{(X - \mathbb{E}\{X\})^2\}}$$
(2)

3. Mean gradient (MG): MG quantifies an image's gradient information, and higher MG means more edge information. MG is mathematically defined as follows:

$$SD = \frac{1}{\Omega} \sum_{x \in X} \sqrt{0.5 \times \left((\mathbf{D}_1 x)^2 + (\mathbf{D}_2 x)^2 \right)}$$
(3)

Where $\mathbf{D}_{\mathbf{i}}$ is a differentiation along a given dimension *i*, and Ω is the image size.

4. **spatial frequency (SF):** SF measures an image gradient (image edge) and texture. It is calculated using row frequency (RF) and column frequency (CF) as follows:

$$SD = \sqrt{RF^2 + CF^2} \tag{4}$$

where $RF = \sqrt{\frac{1}{\Omega} \sum_{x \in X} (\mathbf{D}_1 x)^2}$ and $CF = \sqrt{\frac{1}{\Omega} \sum_{x \in X} (\mathbf{D}_2 x)^2}$. An image with a higher SF provides more image edge information, and human eyes are more sensitive to images with higher SF.

5. Mutual information (MI): For a given two images X and F with marginal probability distributions $p_X(x)$ and $p_F(f)$ and joint probability $p_{X,F}(x, f)$. X and F are statistically independent if $p_{X,F}(x, f) = p_X(x).p_F(f)$. The mutual information I(X, F)of X and F quantify the distance between the joint distribution $p_{X,F}(x, f)$ and the distribution when two distributions are statistically independent $p_X(x).p_F(f)$ using the Kullback-Leibler measure, i.e.,

$$I(X,F) = \sum_{x,f} p_{X,F}(x,f) \log \frac{p_{X,F}(x,f)}{p_X(x).p_F(f)}$$
(5)

It is related to the entropy of the images as follows:

$$I(X, F) = H(X) + H(F) - H(X, F) = H(X) - H(X|F) = H(F) - H(F|X)$$
(6)

where $H(\bullet)$, $H(\bullet, \bullet)$, and $H(\bullet|\bullet)$ are the entropy (given in Equation 1), joint entropy (= $-\sum_{x,f} p_{X,F}(x, f) \log p_{X,F}(x, f)$), and conditional entropy (= $-\sum_{x,f} p_{X,F}(x, f) \log p_{X|F=f}(x)$), respectively.

6. Normalized cross correlation (NCC): NCC measures linear similarity between two images using the direct use of image signal intensities. It is calculated as follows:

$$NCC = \lambda_x \cdot r_{X,F} + \lambda_y \cdot r_{Y,F} \tag{7}$$

 $r_{Z,f} = \frac{\mathbb{E}\{(Z - \mathbb{E}\{Z\}) : (F - \mathbb{E}\{F\})\}}{\sqrt{\mathbb{E}\{(Z - \mathbb{E}\{Z\})^2\}} \cdot \sqrt{\mathbb{E}\{(F - \mathbb{E}\{F\})^2\}}}$ where $Z \in \{X, Y\}$. It ranges from 0 to 1 and greater value means a higher similarity between source images and fused image.

7. **Peak signal-to-noise ratio (PSNR):** PSNR calculated in decibel measure image representation fidelity. It is a ratio between maximum signal intensity power and the noise distortion power, i.e.,

$$PSNR = 10\log_{10}\frac{s^2}{MSE}\tag{8}$$

where s is the maximum signal intensity. MSE given in Equation 9 is the mean squared error that measure the image similarity.

$$MSE = \sum_{z \in \{X,Y\}} \lambda_z \cdot \frac{1}{M.N} \sum_{i}^{M} \sum_{j}^{N} (z_{i,j} - F_{i,j})^2$$
(9)

8. $\mathbf{Q}^{\mathbf{X}\mathbf{Y}/\mathbf{F}} : Q^{XY/F}$ quantifies the amount of the edge information of the fused image. Considering the fusing source images X and Y and fused image F, edge (g(i, j)) and orientation $(\alpha(i, j))$ for each pixel is calculated as follows:

$$g_Z = \sqrt{(\mathbf{D}_1 Z)^2 + (\mathbf{D}_2 Z)^2}$$

$$\alpha_Z = \tan^{-1} \left(\frac{(\mathbf{D}_2 Z)^2}{(\mathbf{D}_1 Z)^2} \right)$$
(10)

where $Z \in X, Y$ and $\mathbf{D}_{\mathbf{i}}Z$ is the differential of Z along dimension *i*.

The relative edge strength (G^{ZF}) and orientation (A^{ZF}) of an input image $Z \ (\in \{X, Y\})$ with respect to F are as follows:

$$G^{ZF}(i,j) = \begin{cases} \frac{g_F}{g_Z} & if \ g_Z > g_F \\ \frac{g_Z}{g_F} & otherwise \end{cases}, \text{ where } Z \in \{X,Y\} \\ A^{ZF}(i,j) = \frac{||\alpha_Z(i,j) - \alpha_F(i,j)| - \pi/2|}{\pi/2}, \text{ where } Z \in \{X,Y\} \end{cases}$$
(11)

Then, we can estimate the perceptual information loss in fused image for an input image $Z \ (\in \{X, Y\})$ as follows:

$$Q_g^{ZF} = \frac{\Gamma_g}{1 + e^{k_g} (G^{ZF} - \sigma_g)}$$

$$Q_\alpha^{ZF} = \frac{\Gamma_\alpha}{1 + e^{k_\alpha} (A^{ZF} - \sigma_\alpha)}$$
(12)

where constants Γ_g , k_g , σ_g , Γ_{α} , k_{α} , and σ_{α} determine the exact shape of the sigmoid functions that form the edge strength and orientation values. Equation 13 calculates the edge information preservation.

$$Q^{ZF} = Q_q^{ZF} \cdot Q_\alpha^{ZF} \tag{13}$$

where $Q^{ZF} = 1$ means lossless image fusion.

Finally, we obtain the fusion edge preservation from the weighted sum of the edge information, i.e.,

$$Q^{XY/F} = \frac{\lambda_X Q^{XF} + \lambda_Y Q^{YF}}{\lambda_X + \lambda_Y} \tag{14}$$

 $Q^{XY/F} \in [0,1]$ where the higher value means a better image fusion with lower information loss.

9. Structural similarity index (SSIM): SSIM assess the structural similarity of two images. It consists of three pixelwise comparisons including luminance, contrast, and structure. SSIM for two images Z and F is defined as follows:

$$SSIM(Z,F) = \frac{(2\mu_Z\mu_F + C_1)(2\sigma_{ZF} + C_2)}{(\mu_Z^2 + \mu_F^2 + C_1)(\sigma_Z^2 + \sigma_F^2 + C_2)}$$
(15)

where μ_z , σ_F , and σ_{ZF} are the local mean, local standard deviations, and local covariance between images Z and F, respectively. C_1 and C_2 are the constant parameters to stable the SSIM. The SSIM between source images and fused image is a weighted sum of the Equation 15 and is defined as:

$$SSIM = \lambda_X^{\text{SSIM}}.SSIM(X, F) + \lambda_Y^{\text{SSIM}}.SSIM(Y, F)$$
(16)