Physics-Driven Deep Learning Methods for Fast Quantitative Magnetic Resonance Imaging

### **CONTENTS**

- Introduction
- Physics and reconstruction of qMRI
- Training sample generation using physical models
- Physical model-based synthetic images via DL
- Physical model-integrated loss function design
- Physical model consistency network design

### Introduction

qMRI(quantitative MRI): quantitative measurements of tissue in physical units

- Acquire contrast-weighted images
- Obtain quantitative parameter



• **MR physics and physical models**

The bulk magnetization  $\mathbf{M} = M_x \mathbf{i} + M_y \mathbf{j} + M_z \mathbf{k}$ 

When put in a static main magnetic field  $B_0$ , **Bloch equation**:

$$
\frac{d\mathbf{M}(\tau)}{d\tau} = \mathbf{M}(\tau) \times \gamma \mathbf{B}(\tau) - \frac{M_x(\tau)\mathbf{i} + M_y(\tau)\mathbf{j}}{T_2} - \frac{(M_z(\tau) - M_0)\mathbf{k}}{T_1}
$$

 $\gamma$ : gyromagnetic ratio

 $M_0$ : equilibrium magnetization

 $\mathbf{B}(\tau)$ : spatially and time-varying total magnetic field

 $T_1$ ,  $T_2$  : longitudinal and transverse relaxation time

• **MR physics and physical models**

**Bloch simulations** describe the relationships between magnetization and biophysical parameters:

$$
\rho_{t_i} = \mathbf{A}(x,t_i)
$$

x : biophysical parameter

 $\rho_{t_i}$ : magnetization signal measured with the sequence parameter to

An MR sequence can be manipulated to "weight" the magnetization by changing a certain sequence parameter  $t_i$  to estimate the parameter  $x_i$ .





Table 1. The commonly used physical models and their dependencies on MRI sequences.



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### • **MR imaging model**

To form an MR image, the magnetization of the imaging region should be spatially encoded during data acquisition.

$$
b_{t_i}^{\ell}(\mathbf{k}) = \int c^{\ell}(\mathbf{r}) \cdot \rho_{t_i}(\mathbf{r}) \cdot e^{-j2\pi \mathbf{k}^{\mathrm{T}} \mathbf{r}} d\mathbf{r} + \varepsilon_{t_i}(\mathbf{k}), \ell = 1, ..., \mathcal{L}
$$

- $b_{t_i}^{\ell}(\mathbf{k})$  : measured k-space data from the *l* th receiver coil
- $\varepsilon_{t_i}(\mathbf{k})$  : associated measurement noise
- $\rho_{ti}(\mathbf{r})$  : image intensity that reflects magnetization distribution in the field of view
- $c^{\ell}(\mathbf{r})$  : *l* th coil sensitivity

 $\mathbf{k} \in [-0.5, 0.5)^d$  and  $\mathbf{r} \in \mathbb{R}^d$ , denote the k-space and image domain coordinates

$$
\mathbf{b}_{t_i}^{\ell} = \mathbf{F} \mathbf{C}^{\ell} \boldsymbol{\rho}_{t_i} + \boldsymbol{\varepsilon}_{t_i} \qquad \qquad \mathbf{b}_{t_i} = \mathbf{E} \boldsymbol{\rho}_{t_i} + \boldsymbol{\varepsilon}_{t_i}
$$

### • **Reconstruction**

Approaches of qMRI: (1) MR image reconstruction; (2) parameter fitting

Nonlinear inverse problems:

biophysical parameters(unknowns), contrast-weighted images(observations)

$$
\hat{\mathbf{x}} = \underset{\mathbf{x} \in \mathbb{C}^{N \times N_p}}{\operatorname{argmin}} \frac{1}{2} \sum_{i=1}^{L} \|\mathbf{A}(\mathbf{x}, t_i) - \boldsymbol{\rho}_{t_i}\|_2^2 \triangleq \frac{1}{2} \|\mathbf{A}(\mathbf{x}, t) - \boldsymbol{\rho}_{t}\|_F^2
$$

**x**: vectorized biophysical parameter map N<sub>p</sub>: the number of biophysical parameters L: the total number of images;  $\mathbf{A}(\mathbf{x},t) = [\mathbf{A}(\mathbf{x},t_1), \mathbf{A}(\mathbf{x},t_2), ..., \mathbf{A}(\mathbf{x},t_L)], \rho_t = [\rho_{t_1}, \rho_{t_2}, ..., \rho_{t_L}]$ 

- **Reconstruction**
- Heteroscedasticity of this noise process: weighted-least-squares
- Special applications with ill-posed models:  $\hat{\mathbf{x}} = \underset{\mathbf{x} \in \mathbb{C}^{N \times N_p}}{\text{argmin}} \frac{1}{2} ||\mathbf{A}(\mathbf{x}, t) \boldsymbol{\rho}_t||_F^2 + \mathcal{R}(\mathbf{x})$
- Fast qMRI, which undersamples the k-space:

$$
\hat{\mathbf{x}} = \underset{\mathbf{x} \in \mathbb{C}^{N \times N_p}}{\operatorname{argmin}} \frac{1}{2} \|\mathbf{EA}(\mathbf{x}, t) - \mathbf{B}_t\|_F^2 + \mathcal{R}(\mathbf{x}; \boldsymbol{\rho}_t)
$$

 $\mathbf{B}_t = [\mathbf{b}_{t_1}, \mathbf{b}_{t_2}, ..., \mathbf{b}_{t_L}]$  acquired k-space data of all contrast-weighted images **E** the forward imaging operator

### Categories of physics-driven DL-based fast qMRI methods

### Table 2. A summary of the strategies of DL-based fast qMRI methods.

#### Category

#### Type 1

Training sample generation via physical models

#### Type 2

Predicting missing/optimized contrast-weighted images via physical models and networks

#### Type 3

Loss function design using synthetic k-space data/images generated from physical models

#### Type 4

Network design using physics priors as a data consistency layer

#### **Advantages**

Easily implemented and low cost

More stable estimation from fewer contrast images

Adds an additional loss term, even allowing unsupervised learning

Explicitly incorporates physical models into a network and typically requires less training data

#### **Limitations**

The diversity of generated training data from real testing data may introduce uncertain errors

Synthesized images may be different from ideal images, and error will be propagated to quantitative maps.

It is difficult to choose weights for different loss terms.

Theoretical convergence is not guaranteed.

### **Applications**

 $T_{2}$  $T_1$  and  $T_2$ -MRF Magnetization transfer contrast (MTC)-MRF MT Diffusion tensor imaging **CEST** 

 $T_{2}$ **MTC-MRF** 

**QSM**  $T_{1}$ 

## Training sample generation using physical models

• Generating training samples using Bloch simulation with a predefined parameter range.



**FIGURE 1.** The type 1 category in Table 2. A deep network is trained for the parameter fitting of qMRI with training samples generated using a physical model.

# Physical model-based synthetic images via DL

- predict missing images by using a network for replenishment
- remove disturbances by using a network for correction and improving overall image quality



**FIGURE 2.** The type 2 category in Table 2. Physical models are implicitly involved when using the network to generate missing and improved images.

## Physical model-based synthetic images via DL —— qMTNet

• qMT(quantitative magnetic transfer): acquire images with multiple off-resonance frequencies for parameter fitting, typically 12 off-resonance images are acquired.

### • qMTNet:

produce 2D qMT parameter maps from 4 off-resonance images by generating missing images.



**FIGURE 3.** The gMTNet. Two subnetworks are involved. gMTNet-acq produces the missing eight offresonance images from the four acquired images. qMTNet-fit obtains qMT parameters from a total of 12 images.

### Physical model-integrated loss function design



 $\left\{\n\begin{array}{l}\n\mathcal{L} = \lambda_1 \mathcal{L}_1 + \lambda_2 \mathcal{L}_2 \\
\mathcal{L}_1 = \|\hat{\mathbf{x}} - \mathbf{x}\|_F^2 \\
\mathcal{L}_2 = \|\hat{\boldsymbol{\rho}}_t - \boldsymbol{\rho}_t\|_F^2 \text{ or } \|\hat{\mathbf{B}}_t - \mathbf{B}_t\|_F^2\n\end{array}\n\right.$ 

**FIGURE 4.** The physical model-integrated loss function design with synthetic data in the (a) image domain and (b) k- space domain. The network is used to generate parameter maps  $\hat{x}$  from input contrast images  $\rho$ . The conventional loss is the error between  $\hat{x}$  and the labels. The synthetic images  $\hat{\rho}_t$  and k-space  $\hat{\mathbf{B}}_t$  can be generated from  $\hat{\mathbf{x}}$  according to the physical model and image operator E. The error between the synthetic data and labeled data is used as an additional loss term.

#### Physical model-integrated loss function design  $\frac{1}{2}$  mapping **Varying Sequence Model Parameter** Parameter  $(t_i)$ **Sequence** Spin echo Echo time  $\rho_{\mu} = I_0 \cdot e^{-\frac{t_i}{T_2}}$ Relaxation  $T_2$

• MANTIS (model-augmented NN with incoherent k-space sampling): Directly estimate T<sup>2</sup> maps from undersampled k-space data, using a CNN with the loss function  $\mathcal{L} = \lambda_1 \| (\hat{\mathbf{I}}_0, \hat{\mathbf{T}}_2) - (\mathbf{I}_0, \mathbf{T}_2) \|_F^2 + \lambda_2 \| \hat{\mathbf{B}}_t - \mathbf{B}_t \|_F^2$ 

 $I_0$  and  $T_2$ : the proton density and  $T_2$  maps to be reconstructed  $\hat{\mathbf{B}}_t = \mathbf{EA}((\hat{\mathbf{I}}_0, \hat{\mathbf{T}}_2), \mathbf{TE})$ : the synthetic k-space data from the forward physical model : the echo times of  $T_2$  mapping **TIR** 

• RELAX (reference-free latent map extraction):

$$
\mathcal{L} = \|\hat{\mathbf{B}}_t - \mathbf{B}_t\|_F^2 + \lambda \mathcal{R}(\hat{\mathbf{I}}_0, \hat{\mathbf{T}}_2)
$$

## Physical model consistency network design

• With DL, iterations can be unrolled into a deep network to learn regularization and data fidelity terms.



**FIGURE 5.** The unrolling-based network design for qMRI. Each iteration in traditional iterative reconstruction is unrolled as a network module. The physical model is incorporated as a part of the measurement operator that transfers parameter maps into the raw data space to enforce DC.

#### Physical model consistency network design **Varying Sequence**  $\frac{1}{1}$  mapping **Model Parameter Sequence** Parameter  $(t_i)$ Relaxation  $T_1$  $\rho_n = I_0 \cdot \sin(t_i) \cdot \frac{1 - E_1}{1 - E_1 \cos(t_i)}$ Gradient echo Flip angle

DOPAMINE: A deep model-based MR parameter mapping network  $\bullet$  $\mathcal{R}(\mathbf{X}) = \|\mathbf{X} - \mathcal{D}_{\mathrm{R}}(\mathbf{X})\|_{2}^{2}$ 

where  $\mathcal{D}_R(\mathbf{X})$ :  $\mathbb{C}^{N\times 1}$   $\mapsto$   $\mathbb{C}^{N\times 1}$  denotes a CNN denoiser.

$$
\hat{\mathbf{X}} = \underset{\mathbf{X}}{\operatorname{argmin}} \frac{1}{2} ||\mathbf{A}(\mathbf{X}) - \mathbf{B}_t||_F^2 + \lambda ||\mathbf{X} - \mathcal{D}_R(\mathbf{X})||_2^2
$$

$$
\mathbf{X}_{k+1} = \mathbf{X}_k - 2\mu_k [\mathbf{J}_{\mathbf{A}}^{\mathbf{H}}(\mathbf{X}_k)(\mathbf{A}(\mathbf{X}_k) - \mathbf{B}_t) + \lambda_k (\mathbf{X}_k - \mathcal{D}_{\mathbf{R}}(\mathbf{X}_k))]
$$
  
=  $(1 - 2\lambda_k \mu_k) \mathbf{X}_k - 2\lambda_k \mu_k \mathcal{D}_{\mathbf{R}}(\mathbf{X}_k) - 2\mu_k \mathbf{J}_{\mathbf{A}}^{\mathbf{H}}(\mathbf{X}_k)(\mathbf{A}(\mathbf{X}_k) - \mathbf{B}_t)$ 

 $E_1 = e^{-\frac{TR}{T_1}}$ 

### Physical model consistency network design  $\frac{1}{1}$  mapping



**FIGURE 8.** The overall architecture of DOPAMINE. The initial  $X_i$  is generated by the mapping network from zero-filling  $T_i$ -weighted images. The network  $\mathcal{D}_R$  serves as a CNN-based denoiser, and the physical model is incorporated into the operation of  $J_A^H$  in the DC layer.