

Integrative and Multiscale Image Registration: Application to Fetal and **Neonatal Brain Development Analysis**

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Abstract

The quantification of abnormal brain development is important for the study of diseases such as Intrauterine Growth Restriction (IUGR) or Ventriculomegaly, where already in utero changes in brain structure and function can be noticed and where the neurodevelopmental outcome is unknown. To be able to characterize early the severness of the disease, quantification of the abnormal fetal brain development and comparison to normality are necessary. To achieve this, image registration is an important preprocessing step for spatial normalization of subjects and/or populations. To obtain good registration results, which is crucial for the quantification, we adress two different issues of image registration, the integration of additional sources of information and the multiscale nature of medical image data. For the former, we propose a spatially weighted multichannel similarity measure to distiguish between useful and redundant additional information. For the latter, we present an adaptive multiscale similarity measure which combines image statistics at multiple scales for a multiscale representation of regional image similarities. In addition, we present the idea of simultaneously optimization of transformation parameters to capture better the different scales in the datasets. Both approaches show improvements over common registration methods. Preliminary results are presented on synthetic and clinical data.

Application

Development of an integrative and multiscale registration framework to model and quantify fetal and neonatal abnormal brain development and to estimate their variability within and across populations.

Intrauterine Growth Restriction:

- occurs 5 10 % of pregnancies
- due to placental insufficiency
- preterm birth
- associated with neurostructural and neurodevelopmental anomalies



Gray and white matter volume changes between IUGR and term infants [1].

Ventriculomegaly:

• occurs in 0.15 - 0.7 % of pregnancies • ventricular enlargement



US of fetus with ventriculomegaly.

 Association with schizophrenia, autism and ADHD



MRI of normal control fetus (A) and fetus with ventriculomegaly (B) [2].

 $\lambda_l(p) = \mathcal{S}^{l,p} - 1 \in [0,1]$

 $+ \alpha_3 S_3(D_3; T_3(T_2(T_1)))$

 $\mathcal{S}^{0,1} = \mathcal{M}^{0,1}$

 $\text{ if }p'\in P$

otherwise

	Methods		
Image Registration:	Integration of information	Multiscale Image Registration:	
Images:	Multichannel similarity measure [3]:	Adaptive Multiscale Similarity Measure [4]: $S^{2,p}$ $S^{1,p}$	
$F, M : \mathbb{R}^{n} \to \mathbb{R}$ Transformation:	$\mathcal{S}(\mathbf{F}, \mathbf{M}; T) = \sum_{c=1} \lambda_c \cdot \mathcal{S}_c(F_c, M_c; T)$	• Measuring image similarity on multiple $l=2$ levels at spatially varying locations	
$T: \mathrm{I\!R}^a \to \mathrm{I\!R}^a$	$\mathbf{F} = [F_1, \dots, F_C] \qquad \mathbf{M} = [M_1, \dots, M_C]$	 Combination of regional similarities through adaptive weights: low regional 	
Find a transformation T such		similarity → higher weight Patchwise matching measure: 	
image M is most similar to a	FA FA FA FA	$\mathcal{M}^{l,p} = (1 - \lambda_l)\mathcal{S}^{l,p} + \lambda_l \sum^{2^{d(l-1)}} \alpha_{p'} \mathcal{M}^{l-1,p'} \qquad \lambda_l(p) = \mathcal{S}$	
$\mathcal{I}(T) = \mathcal{S}(F \ M \circ T_{*})$	T1 T2 T1 T1 T2 T1 T1 T2 T1 T2 T1 T2 T1 T2 T1 T1 T1 T2 T1	• Final measure: $\mathcal{M}^{(L)} = \frac{1}{2^{dL}} \sum_{j=1}^{2^{dL}} \mathcal{M}^{L,p} \qquad \qquad$	
$+ \alpha \mathcal{R}(T)$	$S_c(F_c, M_c; T) =$	$\frac{2^{nL} p=1}{p=1}$	
$T^* = \min(\mathcal{J}(T))$	$\sum_{x \in \Omega_F} w_c(x) (F_c(x) - M_c(T(x)))^2$	Fixed and I_{-1} $\mathcal{S}(D_L; T_L) = \alpha_1 \mathcal{S}_1(D_1; T_1) + \alpha_2 \mathcal{S}_2(D_2; T_2(D_1; T_1))$	
		moving $L=2$ + $\alpha_3 S_3(L)$	

 \mathcal{R} : Regularization term, S : Similarity measure



Gradient Magnitude Local Entropy

NMI

L=3 pyramids $D_L = [F_L, M_L]$



Multiscale transform reperesentation

Multichannel Image Registration with spatial weighting:							
X / C	no weighting	GM	LE				
T2	-2.170		Sec. 19				
T2-FA	-2.196	-2.023	-2.241				
T2-ADC	-2.248	-2.023	-2.303				
T2-FA-ADC	-2.295	-2.300	-2.319				

- Data: neonatal MRI and DTI (T1,T2,FA,ADC)
- Registration results assessed with MI
- Spatial weighting with Gradient Magnitude and Local Entropy
- \rightarrow Including DTI info through spatially weighted/
 - multichannel can improve results.

Results and Future Work

image

Multisca



le Image Registration:							
	Mod.	INU	NMI	AM-NMI			
	T1-T1 T1-T2 T1-PD	0%	$\begin{array}{c} 0.459 \pm 0.012 \\ 0.965 \pm 0.012 \\ 0.821 \pm 0.009 \end{array}$	$\begin{array}{c} 0.419 \pm 0.013 \\ \textbf{0.929} \pm \textbf{0.012} \\ 0.806 \pm 0.010 \end{array}$			
	T1-T1 T1-T2 T1-PD	20%	$\begin{array}{c} 0.465 \pm 0.011 \\ 1.164 \pm 0.011 \\ 1.237 \pm 0.031 \end{array}$	$\begin{array}{c} 0.420 \pm 0.020 \\ \textbf{0.940} \pm \textbf{0.026} \\ \textbf{0.913} \pm \textbf{0.025} \end{array}$			
-10 0 10 20 translation	T1-T1 T1-T2 T1-PD	40%	$\begin{array}{c} 0.468 \pm 0.010 \\ 1.329 \pm 0.016 \\ 1.485 \pm 0.050 \end{array}$	$\begin{array}{c} 0.421 \pm 0.007 \\ 1.007 \pm 0.046 \\ 0.972 \pm 0.025 \end{array}$			

- synthetic ex: AM-NMI is, in contrast to NMI, able to detect the minimum for translation
- BrainWeb data with random deformation: AM-NMI is robust to increasing INU fields

Future Work:

- Investigate the power of simultaneously multiscale transform optimization:
- Adaptation of transformation model to anatomy, e.g. including growth models
- Population analysis with respect to abnormal brain development for fetuses affected by IUGR and/or by Ventriculomegaly.

References: [1] N. Padilla et al: Differential effects of intrauterine growth restriction on brain structure and development in preterm infants: A magnetic resonance imaging study. Brain Research 1382, pp. 98–108, 2011. [2] V. Kyriakopoulou et al: Cortical Overgrowth in Fetuses With Isolated Ventriculomegaly. Cerebral Cortex, March 2013. [3] H.-J. Park et al: Spatial normalization of diffusion tensor mri using multiple channels. NeuroImage 20, pp. 1995–2009, 2003. [4] V.A. Zimmer and G. Piella: An adaptive multiscale similarity measure for non-rigid registration. WBIR 2014.

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