Empirical field mapping for gradient nonlinearity correction of multi-site diffusion weighted MRI

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ABSTRACT

Background: Achieving inter-site / inter-scanner reproducibility of diffusion weighted magnetic
resonance imaging (DW-MRI) metrics has been challenging given differences in acquisition
protocols, analysis models, and hardware factors.

5 Purpose: Magnetic field gradients impart scanner-dependent spatial variations in the applied 6 diffusion weighting that can be corrected if the gradient nonlinearities are known. However, 7 retrieving manufacturer nonlinearity specifications is not well supported and may introduce errors 8 in interpretation of units or coordinate systems. We propose an empirical approach to mapping the 9 gradient nonlinearities with sequences that are supported across the major scanner vendors.

10 Study Type: Prospective observational study

11 Subjects: A spherical isotropic diffusion phantom, and a single human control volunteer

Field Strength/Sequence: 3T (two scanners). Stejskal-Tanner spin echo sequence with b-values
of 1000, 2000 s/mm² with 12, 32, and 384 diffusion gradient directions per shell.

Assessment: We compare the proposed correction with the prior approach using manufacturer specifications against typical diffusion pre-processing pipelines (i.e., ignoring spatial gradient nonlinearities). In phantom data, we evaluate metrics against the ground truth. In human and phantom data, we evaluate reproducibility across scans, sessions, and hardware.

18 **Statistical Tests:** Wilcoxon rank-sum test between uncorrected and corrected data.

19 **Results:** In phantom data, our correction method reduces variation in mean diffusivity across 20 sessions over uncorrected data (p<0.05). In human data, we show that this method can also reduce 21 variation in mean diffusivity across scanners (p<0.05).

22 Conclusion: Our method is relatively simple, fast, and can be applied retroactively. We advocate

23 incorporating voxel-specific b-value and b-vector maps should be incorporated in DW-MRI

24 harmonization preprocessing pipelines to improve quantitative accuracy of measured diffusion

25 parameters.

26 Keywords: Gradient Nonlinearity, Field Estimation, Pre-processing, DW-MRI

28

INTRODUCTION

29 Physics underlying magnetic resonance imaging (MRI) gradient coil designs result in nonuniform 30 magnetic field gradients during acquisition. This leads to spatial image warping [1-4] in magnetic 31 resonance images and gradient distortion in diffusion weighted magnetic resonance imaging (DW-32 MRI) [5-9]. The introduced spatial variation can impact estimated diffusion tensor information 33 [10] or high-angular resolution diffusion measurements [11]. Bammer et al. show in extreme cases 34 the gradient nonuniformity can lead to an overestimation in the diffusion coefficient up to 30% 35 and an underestimation up to 15% [12]. The severity of the effect increases with distance from the magnet's isocenter [12] and with higher gradient amplitudes [12, 13]. The artifact becomes 36 37 especially troubling for multi-site studies that have varying scanner models and manufacturers [14] 38 and for studies utilizing very large gradient amplitudes such as in the human connectome project 39 (HCP) which utilized amplitudes up to 300 mT/m [13, 15, 16]. Recent work has shown the effect 40 of gradient nonlinearities in the HCP cohort results in considerable bias in tractography results and 41 potentially incorrect interpretations in group-wise studies [17].

42 Various estimates of the coil magnetic field nonlinearities have been applied to improve accuracy 43 within and across sites [18-21]. An adaptive correction of diffusion information proposed by 44 Bammer et al. relies on calculating the spatially varying gradient coil L. This approach is achieved 45 by relating the actual gradients with the desired gradients [12], and has become standard practice 46 [22, 23]. However, this approach assumes that the gradient calibration specified by the manufacturer is readily available. Spherical harmonics (SH) based techniques are already 47 48 implemented by manufacturers in the scanning systems to account for the spatial image warping 49 effects of gradient nonlinearities [1, 24-27]. Yet, the spherical harmonic coefficients are not

usually provided to regular users and may be subject to non-disclosure criteria. Additionally
gradient nonlinearity correction has been approached using noncartesian MR image reconstruction
[28].

53 To remove the need for the manufacturer supplied specifications, we demonstrate an empirical field-mapping procedure which can be universally applied across platforms as defined by Rogers 54 55 et al. [29, 30]. At two scanners (scanner A and scanner B), a large oil-filled phantom is used to 56 measure the magnetic field produced by each gradient coil. To estimate the achieved diffusion 57 gradient directions and b-values on a voxel-wise basis, solid harmonic basis functions are fit to the 58 measured magnetic field. The measured diffusivity (MD) and fractional anisotropy (FA) are 59 compared without nonlinearity correction, with nonlinearity correction using estimated fields, and 60 with nonlinearity correction using fields specified by the manufacturer for an ice-water diffusion phantom. The reproducibility is compared between without nonlinearity correction and with 61 62 nonlinearity correction with the estimated fields for a subject scanned at two positions within the 63 scanner at scanner A. We show that our method removes the need for manufacturer specified 64 spherical harmonic coefficients and that the method reduces MD reproducibility error in-vivo 65 when the effect of gradient nonlinearities is present.

66

METHODS

67 Measurement of gradient coil-generated magnetic fields

Data were acquired across two 3T scanners: Scanner A and scanner B. Both of these are 94 cm
bore Philips Intera Achieva MR whole-body systems and have a gradient strength of 80 mT/m, a
200 T/m/s slew-rate. A phantom is used to estimate the gradient coil fields. The phantom is 24

71 liters of a synthetic white oil (SpectraSyn 4 polyalphaolefin, ExxonMobil) in a polypropylene 72 carboy with an approximate diameter of 290mm and height of 500mm [30]. This oil is used by the 73 manufacturer for some of their calibration phantoms which made it a reasonable choice. The 74 phantom was placed approximately at scanner isocenter and imaged with a dual echo GRE-based 75 field mapping sequence. Images are acquired at two echo times 1ms apart, and the fieldmap is 76 computed from the phase difference of the two images. This follows the manufacturer's field 77 mapping and provides a field map with minimal phase wrapping or distortion. Four field maps 78 were acquired, one with shim field set to 0.05 mT/m on each axis X, Y, Z plus a final image with gradient coil shim fields set to zero. Each used a 384 mm field of view with 4 mm isotropic voxel 79 80 size. Total scan time was approximately 5 minutes. Gradient coil fields were estimated by 81 subtracting the zero-shim field map from each coil's respective 0.05 mT/m field map. It should be 82 noted that the proposed method requires that the field maps are made using the same coils used to produce the diffusion gradients, and systems that utilize gradient coil inserts may not be able to 83 84 directly utilize the technique. Field maps were acquired on 40 dates over the course of a year at 85 scanner B while scanner A only one session was acquired with the fieldmapping phantom.

For each coil, we modeled the magnetic field spatial variation as a sum of solid harmonics [12, 31, 32] to 7th order, excluding even order terms due to the coils' physical symmetry. These basis functions were fit to the field measurements with robust least squares, using all voxels within a 270 mm diameter sphere at isocenter. For comparison, the general shape of the human head is an ellipsoid with an average height of 180 to 200mm [33]. The result was an analytically differentiable estimate of the true magnetic field produced by each gradient coil (Figure 1). This fitting procedure was performed on an average field map derived from a series of scans to ensure stability. On 93 Scanner B, the fitting procedure is also performed on the scanner manufacturer's estimate of the 94 coil fields as measured during manufacturing and installation. These are provided as a set of solid 95 harmonic functions and corresponding coefficients. The series of scans which are averaged are 96 defined for each subject session according to the closest 10 field map sessions in terms of date for 97 scanner B whereas 10 acquisitions were acquired within a single session at scanner A which are 98 averaged.

99 Estimating achieved b-values and gradient directions

100 A spatially varying tensor *L* relates the achieved magnetic field gradient to the intended one [12]:

101
$$L = \begin{bmatrix} \frac{\partial B_z^{(x)}}{\partial x} & \frac{\partial B_z^{(y)}}{\partial x} & \frac{\partial B_z^{(z)}}{\partial x} \\ \frac{\partial B_z^{(x)}}{\partial y} & \frac{\partial B_z^{(y)}}{\partial y} & \frac{\partial B_z^{(z)}}{\partial y} \\ \frac{\partial B_z^{(x)}}{\partial z} & \frac{\partial B_z^{(y)}}{\partial z} & \frac{\partial B_z^{(z)}}{\partial z} \end{bmatrix}$$
(1)

102 where $B_z^{(x)}$ is the z component of the magnetic field produced by unit amplitude of a nominal x-103 gradient coil current, and similarly for (y) and (z). This tensor may be computed analytically from 104 the solid harmonic approximation to the measured field, then evaluated at spatial locations of 105 interest. We can use *L* to relate the assumed gradient vector to the achieved gradient field and as 106 well as the assumed b-value to the achieved one. If we assume |g| = 1 then the adjusted gradient 107 vector and b-value become:

$$g' = Lg \tag{2}$$

$$g'' = \frac{g'}{|g'|} \tag{3}$$

110
$$b' = b|g'|^2$$
 (4)

111 where b' is the adjusted b-value and g'' is the adjusted and normalized gradient vector. In the 112 common situation where the scanner reports the intended gradient direction and amplitude but the 113 full b-matrix [34-36] is not known, an approximate correction to adjust the signal S_i for the i^{th} 114 diffusion acquisition relative to the reference signal S_0 is [18]:

115
$$\ln\left(\frac{s_i}{s_0}\right) = -bg_i^T Dg_i^T = -bg_i^T L^T DLg_i$$
(5)

where *b* is the scalar b-value, *g* is the intended gradient vector, g' is the actual gradient vector, and *D* is the diffusion tensor. If we substitute with b' and g'' equation 5 can be re-written as:

118
$$\ln\left(\frac{S_i}{S_0}\right) = -b'g_i^{"T}Dg_i^{"} = -b|g_i'|^2 \frac{g_i'^{T}}{|g_i'|} D\frac{g_i'}{|g_i'|} = -bg_i^{T}L^{T}DLg_i$$
(6)

Importantly, this is spatially varying and processing occurs voxelwise, but this may be used in any
desired way for further processing of the diffusion images. Figure 2 shows *L* for each voxel
estimated using our empirical fieldmapping acquired on scanner B.

122

EXPERIMENTS

123 This section describes the set of analyses which aim to show the accuracy of the estimated fields 124 as well as their impact on resulting DW-MRI metrics in phantom and human data. All DW-MRI 125 are corrected for susceptibility distortion [37] and eddy current distortion [15] using FSL.

126 Empirically Estimated Fieldmaps

127 Gradient nonlinearity correction is only viable if we can depend on the estimation to match the 128 true fields. To investigate if the magnitude estimated fieldmaps closely approximate the true fields, 129 we compare them to the field maps specified by the manufacturer on scanner B. This was not done 130 for scanner A as the manufacturer specifications for scanner A were not provided. For comparison, 131 we take the average fieldmap from the latest 10 oil phantom scans on scanner B and calculate the 132 voxel-wise difference between this and the manufacturer specified fields. To evaluate the stability 133 of the empirical estimations, we report the variance across fields estimated from 40 individual oil 134 phantom scans acquired over time on scanner B. These additional acquisitions are unnecessary for 135 practical use and are strictly for evaluation purposes. Only a single acquisition would be needed 136 for this method to be deployed on a scanner to be applied to all previous and future acquisitions. 137 All evaluations on the empirical fields use a spherical mask with a radius of 135mm from isocenter.

138 Polyvinylpyrrolidone (PVP) phantom

139 To evaluate the intra-scanner performance of the gradient field nonlinearity correction with the 140 empirical fieldmaps in a controlled environment, we use a 43% Polyvinylpyrrolidone (PVP) 141 aqueous solution in a sealed spherical container that is 160mm in diameter (PVP phantom) [38]. 142 The PVP phantom is a large homogeneous material, and estimated metrics are expected to be the 143 same across the entire volume. Additionally, toxicology has shown PVP to be safe for use, and 144 PVP is stable and uniform. At scanner B, the phantom was scanned at three positions along the 145 magnet axis: superior (4cm above isocenter), isocenter, and inferior (8cm below isocenter). At 146 each position DWI data was acquired with diffusion weighting applied in twelve directions at a bvalue of 1000 s/mm² and twelve more were acquired at 2000 s/mm² with a TR of 7775, a voxel 147

resolution of 2.5mm by 2.5mm by 2.5mm, and a FOV of 240mm by 240mm by 170mm. 148 149 Susceptibility distortion correction and eddy current distortion correction are applied without 150 movement correction. Signal to noise ratio (SNR) was calculated by fitting the signal to a tensor 151 in the phantom and taking the residuals after the fit. Using all diffusion volumes at each position, 152 MD is calculated without and with gradient nonlinearity correction using the empirically derived 153 fields and using the manufacturer specified fields. When calculating MD with the correction, the 154 estimated achieved b-values and gradient directions for each voxel are used. We report error in 155 terms of absolute percent error (APE) between each scan out of isocenter and the scan at isocenter. 156 All non-diffusion volumes to a structural T1 image using a rigid body transform restricted to only 157 use translations, and this registration is applied to the calculated MD before analysis.

158 Human subject

159 To evaluate the intra-scanner and inter-scanner performance of the gradient field nonlinearity 160 correction with the empirical fieldmaps in-vivo, we scanned a single subject at scanner A and 161 scanner B. At scanner B, two sessions were acquired of the subject with one session acquired with 162 the bridge of the subject's nose positioned at isocenter within the magnet and one session acquired 163 with the subject positioned 6cm superior from isocenter. At scanner A, only one session is acquired 164 at isocenter. Each session consisted of twelve gradient directions at a b-value of 1000 s/mm², twelve at a b-value of 2000 s/mm², a TR of 3700ms, a voxel resolution of 2.5mm by 2.5mm by 165 166 2.5mm, and a FOV of 240mm by 240mm by 170mm. Susceptibility distortion correction and eddy 167 current distortion correction are applied with movement correction for each session. Using all 168 diffusion volumes from each session, MD is calculated without and with gradient nonlinearity 169 correction using the empirically derived fields. At scanner B, MD is also calculated after correction

170 with the manufacturer specifications. For analysis the scans are registered to a T1 acquired at 171 isocenter using FSL Flirt [39]. We report MD error as the absolute percent error between the two 172 scans acquired at scanner B and between the scan acquired at scanner A and the out of isocenter 173 scan acquired at scanner B.

174 We also evaluate the performance of the empirical correction with higher quality acquisitions on 175 scanner A. Again, two sessions are acquired of the subject: one with the bridge of the subject's 176 nose positioned at isocenter and one where the subject is shifted 4cm inferior from isocenter. Each 177 session consisted of 384 gradient directions at a b-value of 1000 s/mm², a voxel resolution of 178 2.5mm by 2.5mm by 2.5mm, and a FOV of 240mm by 240mm by 170mm. Susceptibility distortion 179 correction and eddy current distortion correction are applied with movement correction for each 180 session. Using all diffusion volumes from each session, MD is calculated without and with gradient 181 nonlinearity correction using the empirically derived fields. For analysis the scans are registered 182 to a T1 acquired at isocenter using FSL Flirt [39]. We report MD error as the absolute percent error 183 between the two scans.

184

RESULTS

185 Empirically Estimated Fieldmaps

There are small differences between the manufacturer and the measured field produced by the gradient coil. These are shown in Figure 1 in units of uT scaled by the intensity (mT/m) of the applied gradient (uT/(mT/m), or mm). On average the difference at a given voxel is approximately 1 uT/(mT/m) in the x and y magnetic field gradients and 2 uT/(mT/m) in the z gradient field within 135mm of isocenter. The difference maps indicate the presence of some structural artifacts. The average standard deviation at a given voxel after 40 acquisitions acquired throughout a year is approximately 4 uT/(mT/m) in the x and y fields and 6 uT/(mT/m) in the z field within 135mm of
isocenter.

194 **PVP phantom**

The mean absolute percent error within the phantom between the inferior scan and the isocenter scan is approximately 5% before correction. After correction using the manufacturer fields, this falls to approximately 1.6%. Correcting with the empirically derived fields leads to 0.9% mean error. Figure 3 shows most of the error before correction in the inferior regions of the phantom which were furthest from isocenter in the inferior scan.

When uncorrected, the mean absolute percent error within the phantom between the superior scan and the isocenter scan is approximately 4.9%. After correction using the manufacturer fields, this falls to approximately 2%. Correcting with the empirically derived fields leads to 1.3% mean error. Figure 4 shows most of the error before correction in the superior regions of the phantom which were furthest from isocenter in the superior scan.

205 Human repositioned

The intra-scanner sessions on scanner B result in a mean absolute percent error of 5.9% before correction within the brain volume excluding CSF regions. After correcting the scans using the empirically estimated fields, the mean error is reduced to 5.6% and further to 5.4% if the manufacturer specifications are used during correction. Just as in with the phantom, the error attributable to the gradient nonlinearities before correction appears in the superior regions of the brain which were furthest from isocenter during one of the sessions (Figure 5).

For the inter-scanner experiment, the mean absolute percent error before correction is 7.2% and is reduced 6.9% after correction using the estimated fields. Clearly the error that is accounted for in the correction is the superior regions of the brain which were furthest from isocenter during the session acquired on scanner B (Figure 6).

The intra-scanner sessions acquired on scanner A using a significantly higher number of gradient directions results in a mean absolute percent error of 4.6% when no correction is applied. After correction using the empirically estimated fields, the mean error is reduced to 4.2%. The difference can be seen in the inferior regions of the brain, specifically the cerebellum which was furthest from isocenter during one of the sessions (Figure 7). Figure 8 shows the mean absolute percent error across all voxels within the phantom and within the brain volume excluding cerebrospinal fluid (CSF) regions for each method.

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DISCUSSION

224 In comparing the empirically estimated fields to the fields specified by the manufacturer, we find 225 that our approximations are very similar. The largest differences are in the z gradient field which 226 corresponds to the largest variations in all the estimated fields across 40 oil phantom acquisitions. 227 In this study we use an average of fieldmaps across 10 acquisitions each acquired a week apart, 228 but this should not be necessary as the field produced by the gradient coil depends only on the coil 229 geometry and the current flowing in the coils. Unaltered system need only acquire the fields once 230 for this method, but further study on the stability of the empirical mapping may be necessary. 231 Additionally, further study on the stability of the fit of the spherical harmonics and the need for 232 higher order basis may be necessary. Appendix A shows the effects of different orders.

233 The experiments with the PVP phantom show in a large isotropic volume the impact of the gradient 234 nonlinearities within the magnet and the effectiveness of the correction. The small superior shift 235 of 4cm results in over 15% error in the superior voxels. In the case of a large inferior shift and a 236 smaller superior shift, the mean error is increased by a factor of two to five if these effects are not 237 accounted for. If we consider the experiments involving the human subject, we can see the impact 238 of this correction is reduced. This could in part due to imperfect registration which seems to have 239 contributed to error in the anterior regions of the brain. Results may vary depending on registration 240 strategy. We have tried multiple techniques with similar results. Though the absolute percent error 241 only changed by 0.3% to 0.4%, some small regions see a similar magnitude of improvement, and 242 it is qualitatively clear that the correction is impacting regions we expect. The differences between 243 resulting absolute percent error using the empirical fields and the manufacturer fields is varies 244 between the phantom and the human subject. The results for the phantom indicate that the 245 estimated fields improve performance of the method, but the human subject results show a small 246 advantage for using the manufacturer field directly.

247 Though all intra-scanner results on scanner B are compared against using the manufacturer field 248 directly, future work should investigate the sensitivity of our proposed method and compare with 249 other field mapping methods such as proposed by Janke et. al [24] even though these methods 250 require that the manufacturer provide the solid harmonic coefficients. In recent work, another 251 approach is proposed for correcting voxel-wise b-value errors. Instead of correcting for gradient 252 nonlinearities in the coil, this method directly estimates a voxel-wise b-value map that is used to 253 correct resulting diffusion metrics [40]. While this method could account for errors that stem from 254 other sources of deviation than just gradient nonlinearities, the model requires an estimation of

more parameters and likely it would be best practice to acquire a calibration scan along with every subject acquisition. In comparison to apply the approach proposed in this work, only a single calibration scan is necessary for each system.

258 While this method is successful in circumventing the need for manufacturer specifications which 259 are not always readily available, it should be noted that vendor-provided on-scanner gradient 260 nonlinearity correction is preferred for translation in a clinical environment. Additionally, when 261 working with any DICOM data coordinating world coordinate frame and patient frames can be 262 incredibly nuanced and should be considered carefully when applying any corrections post 263 acquisition. However, our approach remains as a solution to correct retroactively to enable the use 264 of acquired datasets which should be corrected for gradient nonlinearity effects for use in clinic 265 and in research.

266

CONCLUSION

This work shows that the errors caused by gradient nonlinearities is apparent in metrics derived from DW-MRI but can be reduced using the correction outlined by Bammer et al. Using empirically derived fields, we can achieve similar results without needing manufacturer specification of the hardware. In both phantom and in-vivo data, error in MD can be significantly reduced by applying this correction. We advocate for the use of gradient nonlinearity correction in standard diffusion preprocessing pipelines and provide a simple method for empirically measuring the fields necessary to account for the achieved b-values and b-vectors.

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Figure 1. Here we show the manufacturer specified fields (top), the averaged empirically estimated (directly measured) fields (middle-top), the difference between these (middle-bottom), and the standard deviation in the empirically estimated fields across time (bottom) in units of uT (per mT/m of applied gradient). The field of view is 384mm by 384mm, and a mask is applied to the fields according the usable regions within the oil phantom (135mm radius from isocenter). The x and y magnetic field gradients are shown as an axial slices at isocenter (192mm), and the z magnetic field gradient is shown as a sagittal slice at isocenter (192mm).



Figure 2. Gradient coil tensor L(r) (sagittal view) for each voxel position using 7th order spherical harmonic expansion using only odd order terms. This was generated using the coefficients estimated from the empirical field mapping procedure.



Figure 3. The absolute percent error (APE) in MD is shown for the PVP phantom with one session acquired at isocenter and another acquired 8cm inferior from isocenter. The top plot shows the sagittal and coronal view of the b0 from each session to demonstrate the shift within the scanner. The bottom plots show the APE for nine saggital slices before correction, after correction using the estimated fields, and after correction using the manufacturer specifications. The error before correction is most prominent in the inferior regions of the phantom as those were the furthest from isocenter during the second acquisition.

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Figure 4. The absolute percent error (APE) in MD is shown for the PVP phantom with one session acquired at isocenter and another acquired 4cm superior from isocenter. The top plot shows the sagittal and coronal view of the b0 from each session to demonstrate the shift within the scanner. The bottom plots show the APE for nine saggital slices before correction, after correction using the estimated fields, and after correction using the manufacturer specifications. The error before correction is most prominent in the superior regions of the phantom as those were the furthest from isocenter during the second acquisition.

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Figure 6. The absolute percent error (APE) in MD is shown for the human subject with one session acquired at isocenter on scanner A and another acquired 6cm superior from isocenter on scanner B. The top plot shows the sagittal and coronal view of the b0 from each session to demonstrate the shift within the scanner. The bottom plots show the APE for nine saggital slices before correction, after correction using the estimated fields, and after correction using the manufacturer specifications. The error before correction is most prominent in the superior regions of the phantom as those were the furthest from isocenter during the second acquisition.



Figure 7. The absolute percent error (APE) in MD is shown for the human subject with one session acquired at isocenter and another acquired 4cm inferior from isocenter on scanner A. These acquisitions were acquired with 384 directions. The top plot shows the sagittal and coronal view of the b0 from each session to demonstrate the shift within the scanner. The bottom plots show the APE for nine saggital slices before correction, after correction using the estimated fields, and after correction using the manufacturer specifications. The error before correction is most prominent in the inferior regions of the phantom as those were the furthest from isocenter during the second acquisition.





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APPENDIX A

392 The PVP phantom is corrected using fieldmaps estimated with various orders of solid harmonics.

393 Regardless of the order, both FA and MD reproducibility errors decrease when compared to the

- 394 uncorrected error. However, we find that a 3rd order basis results in the lowest FA error but a higher
- 395 MD error. Between the higher order basis, the 7th order solid harmonics achieves lower FA error.



Figure A.1: The reproducibility error in FA and MD for the PVP phantom are calculated using the estimated fieldmap utilizing different orders of solid harmonics. Orders higher than 3rd achieve lower MD RMSE but tend to have higher FA RMSE.