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Whole-Body Magnetic Resonance Imaging in the Large Population-Based German National Cohort Study: Predictive Capability of Automated Image Quality Assessment for Protocol Repetitions

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1 ABSTRACT

Background: Reproducible image quality is of high relevance for large cohort studies and
can be challenging for magnetic resonance imaging (MRI). Automated image quality
assessment may contribute to conducting radiologic studies effectively.

Purpose: The aims of this study were to assess protocol repetition frequency in populationbased whole-body MRI along with its effect on examination time and to examine the
applicability of automated image quality assessment for predicting decision-making
regarding repeated acquisitions.

9 Materials and Methods: All participants enrolled in the prospective, multicenter German National Cohort (NAKO) study who underwent whole-body MRI at 1 of 5 sites from 2014 to 10 11 2016 were included in this analysis (n = 11,347). A standardized examination program of 12 12 protocols was employed. Acquisitions were carried out by certified radiologic technologists, 13 who were authorized to repeat protocols based on their visual perception of image quality. 14 Eleven image quality parameters were derived fully automatically from the acquired images, 15 and their discrimination ability regarding baseline acquisitions and repetitions was tested. 16 **Results:** At least 1 protocol was repeated in 12% (n = 1359) of participants, and more than 17 1 protocol in 1.6% (n = 181). The repetition frequency differed across protocols (P < 0.001), imaging sites (P < 0.001), and over the study period (P < 0.001). The mean total scan time 18 19 was 62.6 minutes in participants without and 67.4 minutes in participants with protocol 20 repetitions (mean difference, 4.8 minutes; 95% confidence interval, 4.5-5.2 minutes). Ten of 21 the automatically derived image quality parameters were individually retrospectively 22 predictive for the repetition of particular protocols; for instance, "signal-to-noise ratio" alone 23 provided an AUC of 0.65 (P < 0.001) for repetition of the Cardio Cine SSFP SAX protocol. 24 Combinations generally improved prediction ability, as exemplified by "image sharpness" plus "foreground ratio" yielding an area under the curve of 0.89 (P < 0.001) for repetition of 25 26 the Neuro T1w 3D MPRAGE protocol, versus 0.85 (P < 0.001) and 0.68 (P < 0.001) as 27 individual parameters.

- 28 **Conclusion:** Magnetic resonance imaging protocol repetitions were necessary in
- approximately 12% of scans even in the highly standardized setting of a large cohort study.
- 30 Automated image quality assessment shows predictive value for the technologists' decision
- 31 to perform protocol repetitions, and has the potential to improve imaging efficiency.
- 32

33 Keywords

- 34 Population imaging; Large cohort study; Magnetic resonance imaging; Body imaging; Image
- 35 quality; Technical image analysis; Workflow optimization; Radiologic technologist;
- 36 Epidemiology
- 37

38 Abbreviations

39	AUC	Area under the curve
40	CI	Confidence Interval
41	DNN	Deep neural network
42	FOV	Field of view
43	MRI	Magnetic resonance imaging
44	NAKO	German National Cohort
45	UQI	Universal quality index
46	RT	Radiologic technologist
47	SNR	Signal-to-noise ratio

49 INTRODUCTION

50 Whole-body magnetic resonance imaging (MRI) has become a primary imaging technique in 51 population-based cohort studies due to its excellent spatial resolution and soft tissue 52 contrast, capacity for standardization, and lack of ionizing radiation. Several population-53 based cohort studies used this technique, such as the Multi-Ethnic Study of Arteriosclerosis 54 [1], the Framingham Heart Study [2], and the Study of Health in Pomerania [3], among 55 others. In addition, 2 large ongoing population-based studies, namely the UK Biobank 56 (100.000 participants) [4] and the German National Cohort (NAKO; 30,000 participants) [5]. rely on this modality. Their comprehensive databases that combine detailed imaging and 57 58 nonimaging phenotyping can provide valuable information about general health, and thus 59 are a valuable source for assessing potential risk markers, as well as identifying radiomic 60 features of subclinical disease states and personalized medicine [6], or ascertaining the 61 prevalence of incidental findings and outcomes [7].

62 Standardized and reproducible image acquisition is indispensable in such large 63 cohort studies to warrant a consistent basis for further postprocessing, including automated 64 segmentation tasks and data extraction [8]. The examination programs used in whole-body 65 MRI are usually complex and tedious for both radiologic technologists (RTs) and 66 participants, who may be immobilized for the duration of examination, depending on the 67 study [5]. The RTs' workflow generally involves reviewing all images immediately 68 postacquisition to assess quality and to repeat a series if deemed necessary. Common 69 reasons for repetition are image blurring (bulk or physiological motion artifacts), distortions 70 due to susceptibility artifacts, low signal and/or high noise, incorrect anatomical coverage, or 71 protocol-specific abnormalities such as fat-water swapping due to faulty shimming or center 72 frequency settings. This decision-making process is time-consuming and relies on subjective 73 visual perception and professional experience. Moreover, prolonged scan times can result in 74 additional discomfort and increase the likelihood of participant dropouts. In the context of 75 large cohort studies, this process then becomes both inefficient and costly. Developments

leading to automation in workflow and RTs' decision-making are therefore being undertaken
and could result in improved resource allocation and image quality, as well as less
discomfort for participants and patients [9, 10].

The fundamental role of RTs' perception of image quality and its influence on protocol repetition in a large cohort study, as a preceding factor to all further analyses, has not previously been established in the literature. Our study, therefore, aimed to assess protocol repetition frequency as well as the RTs' underlying decision-making process, to determine its correlation with quantitative image quality parameters, and to identify quantitative parameters that are predictive for protocol repetition, with a perspective to objectify and potentially automate this time-consuming task.

87 MATERIALS AND METHODS

88 <u>Study Design and Population</u>

89 Our project was designed as a predefined ad hoc analysis on data from the MRI study of the 90 NAKO. The NAKO is an ongoing, prospective, interdisciplinary, multicenter, population-91 based cohort study undertaken by a network of over 25 German institutions and spans 18 92 study centers across Germany. Its main goal is to investigate risk factors for the 93 development of common chronic diseases such as cancer, diabetes, cardiovascular, 94 neurodegenerative/psychiatric, respiratory, and infectious diseases [11]. For at least 25 95 years, more than 200,000 participants of the general population between the ages of 20 and 96 69 years will be examined in a baseline and follow-up studies. Besides interviews, 97 questionnaires, physical examinations, as well as the collection of biological samples, a 98 subgroup of 30,000 participants received a baseline whole-body MRI at 1 of 5 dedicated 99 imaging centers across Germany between May 2014 and April 2019 [5]. For our study, we 100 included all participants enrolled in the MRI substudy until December 31, 2016, which 101 resembles all currently available data. Participants were excluded from the analysis if they 102 were early dropouts (the examination was interrupted before the first protocol was fully 103 acquired) or if the participants withdrew consent. The scientific advisory board and the ethics 104 advisory board of the NAKO approved this study.

105

106 Image Acquisition

MRI was performed using identical 3 T whole-body scanners (MAGNETOM Skyra, Siemens Healthcare, Erlangen, Germany) installed at the 5 imaging centers, running an identical software version. A high field strength system was chosen for its improved signal-to-noise ratio and enhanced spatial resolution, despite higher costs and a higher likelihood of artifacts. The examination program comprised a set of 12 non-contrast agent-enhanced series, covering 4 anatomical domains, to form a whole-body examination (Figure 1). Aimed at the evaluation of neurological, cardiovascular, thoracoabdominal, and musculoskeletal

pathologies, as well as subclinical disease burden and physiological variants, the domains
were each allocated 15 minutes of scan time, with a total scan time of approximately 60
minutes. A detailed account of the rationale, design and technical background of the MRI
substudy has been given previously [5].

All acquisitions were carried out by RTs who received training and certification specifically for the MRI substudy and had to be recertified annually. Participants were briefed in detail on the scanning procedure and cautioned to minimize movement as well as follow breathing instructions. RTs were instructed to repeat measurements if anatomical coverage was not adhering to an internally defined standard, if severe artifacts were present, or if they considered image quality to be otherwise unsatisfactory.

124

125 <u>Automated Image Quality Assessment</u>

126 All acquired images were transferred "on-the-fly", parallel to the ongoing examination, from 127 the imaging centers to a central storage facility, dubbed the "imaging core", using a virtual 128 private network and the standard DICOM (Digital Imaging and Communications in Medicine) 129 format. Besides general data management, the imaging core provided basic automated 130 guality assurance, including checks for data completeness, conformity to the predefined protocol parameters, and data uniqueness. A set of 11 image-based quality parameters was 131 132 then calculated automatically from the acquired images (calculation time slightly below 60 133 minutes for 1 complete examination). It included a proprietary universal quality index (UQI), 134 sharpness, signal-to-noise ratio (SNR) and specific SNR, structured noise maximum and 135 average, N/2 ghosting level maximum and average, drift, variation over time, and foreground 136 ratio.

The UQI served as a first indication of image quality without giving specific
characteristics. It uses the original image, a noise-filtered, and an edge-filtered version to
determine a score that increases with image noise and decreases with image blur.
Sharpness was evaluated using an entropy focus criterion, based on previously published

work [12]. Further image quality assessment required the computation of binary foreground
and background masks; these were generated using established thresholding methods [1315], although robust mask generation was not possible with all protocols. SNR was then
calculated based on existing methodology [16]. Because its calculation considers the whole
image background, the result may be affected by artifacts there, for example, caused by
subject motion. For this reason, a specific SNR was additionally calculated using predefined
regions of interest to only consider real image noise.

For the detection of structured noise and nyquist-ghosting artifacts, iterative line/column correlation algorithms were used. The first examined the correlation between neighbored lines/columns to detect structure in the image background, which could be caused, for example, by streaking artifacts. The latter separated these lines/columns by half the size of the field of view (FOV) to detect nyquist-ghosting. We assessed the results by averaging over all correlations for general image quality, as well as selecting the maximum value for finding strong local artifacts.

The position of the binary foreground mask was used to evaluate the positioning of the subject. The ratio between foreground and background mask areas then represents the ratio of subject size to FOV choice. Lastly, low-frequency signal drifts are often observed in functional MRI with demanding imaging sequences [17]. A dedicated test module quantified this intensity drift for the Resting State EPI BOLD protocol by calculating the time-wise changes in mean signal intensity in the image foreground, as well as the signal variance over the time series.

For repeated measurements, RTs were required to select one acquisition that would primarily be used for reading and research, while provided with the information above through a Web-based thin client. An additional quality check was performed manually for each protocol by NAKO investigators (radiology residents and board-certified radiologists) while reading for incidental findings. Using a traffic light rating system, protocols were categorized based on completeness of anatomical coverage and severity of artifacts. Those labeled "red" were excluded from further analyses.

169 <u>Statistical Analysis</u>

- 170 Data are presented as mean with standard deviation for continuous variables and counts
- and percentages for categorical variables. Differences in image quality parameters between
- 172 initial protocol acquisitions with and without subsequent repetition were compared by
- 173 Student *t* test. Discrimination ability was assessed by receiver operating characteristic
- 174 curves and corresponding area under the curve (AUC) for single image quality parameters
- as well as for explorative combinations of multiple parameters. A *P* value of < 0.05 was
- 176 considered to denote statistical significance. Statistical analysis was performed using SAS
- 177 (version 9.4; SAS Institute Inc., Cary, NC, USA) and the R programming environment
- 178 (version 3.6.3; R Foundation, Vienna, Austria).

180 **RESULTS**

181 A total of 11,347 participants were examined in the specified time frame. A majority of 182 10,960 (96.6%) completed the full MRI examination. Recruitment across the 5 imaging 183 centers varied within a range of 1319 to 3093 participants per site. Most participants were 184 examined on weekdays; a minority of 350 participants (3.1%) underwent imaging on 185 Saturdays. Nearly equal distribution of participants was observed for the morning (07:00-186 11:59 AM) and afternoon (12:00–4:59 PM) scanning periods, accounting for 47.0% and 45.8% of the study sample, while the evening period (5:00 PM and later) comprised a 187 188 minority of 7.2%. Overall mean scan time was 63.1min and 63.7min, if restricted to participants with complete examinations. 189

190

191 MRI Protocol Repetition—A Per Participant Analysis

In 1359 participants (12.0%), a total of 1558 different protocols were repeated at least once based on the real-time decision of the RT. Repetitions were limited to only 1 protocol for the majority of 1178 participants (10.4%), whereas 2 or more different protocols were repeated in 181 participants (1.6%): 2 in 165, 3 in 14, and 4 protocols in 2 participants. The mean total scan time increased by 4.8 minutes (95% confidence interval (CI), 4.5–5.2 minutes; SD, ±5.9 minutes) from 62.6 to 67.4 minutes in participants with 1 or more protocol repetitions.

198 Repetition frequency in participants (from here on defined as the percentage of 199 participants with at least 1 protocol repetition) fluctuated over the enrollment period with a 200 range of 4.5% to 15.3% in 3-month intervals (P < 0.001), and simultaneously varied across 201 sites (range, 2.3–28.1% sitewide for the same periods, *P* < 0.001) (Figure 2). A higher 202 repetition frequency was observed on Saturdays (16.3%) versus weekdays (11.8%, 203 P = 0.01). Differences in repetition frequency between daytimes were not significant (11.6%) 204 vs 12.1% vs 14.2% for morning [7:00 to 11:59 AM], afternoon [12:00 to 4:59 PM], and 205 evening [5:00 to 9:30 PM], P = 0.11). In a multivariable analysis, only the study site and the 206 enrollment period (time since the first examination) remained predictive for differences in

207 repetition frequency, and notably, odds for protocol repetition increased by 1.25-fold
208 (95% CI, 1.13–1.39) per year of enrollment.

209

211

210 <u>MR Protocol Repetitions—A Per Protocol Analysis</u>

The frequency ranged between 0.1% and 3.6% per protocol; the most frequently repeated
protocols were Cardio Cine SSFP LAX nK (3.6% of initial acquisitions were repeated), Neuro
2D FLAIR axial (2.9%), and Body Multi-echo 3D VIBE (1.9%), while the least frequently
repeated protocols were Cardio MOLLI SAX (0.1%) and Neuro Resting State (0.1%)
(Figure 3). Merely 0.03% of initial protocol acquisitions were repeated more than once,
resulting in a total of 1606 repetition scans.

Of all initially acquired protocols (n = 134,239), 1.2% (n = 1558) were repeated at least once.

220 Ten of the automatically derived image quality parameters, if considered individually,

221 exhibited statistically significant differences between initial acquisitions with and without

subsequent repetition (Table 1 and Supplemental Table 1, exemplary images and

distribution plots; **Figure 4**). The 2 parameters demonstrating this discriminative behavior for

the largest number of protocols were image sharpness and SNR: in 9 of 14 protocols for

which it was measured, sharpness differed significantly between initial and repeated

protocols (P < 0.001 to P = 0.049), and SNR differed significantly in 6 of the 12 protocols for

which it was calculated (P < 0.001 to P = 0.003). For 3 protocols, statistically significant

228 differences between the 2 groups were not observed for any image quality parameter: MSK-

229 Spine T2w FSE sagittal, Body T2w HASTE axial, and Cardio MOLLI SAX.

Classification performance of the image parameters in terms of AUC was especially
high for the neurological protocols, with several parameters exhibiting areas greater than
0.75 when comparing initial acquisitions with and without subsequent repetition (Table 1).
Combinations of image quality parameters generally improved discriminative ability over
single parameters (Supplemental Table 2), although performance remained notably poor

for the musculoskeletal protocols MSK-Hip PDw FS 3D SPACE and MSK-Spine T2w FSE
sagittal (Figure 5).

237

238 <u>Setup Changes for Protocol Repetition and their Effect on the Discriminative Power of</u>

239 <u>Automatically Derived Image Quality Parameters</u>

240 Approximately half of the 1606 protocol repetitions (49.8%) were carried out with manual 241 adjustments to the technical setup, namely, radiofrequency (RF) coil configuration 242 (variations in RF coils and in selection of receive RF coil elements), FOV size, slice position 243 (FOV shifted along the x/y/z axis of the participant), or slice orientation (FOV rotated or 244 angled differently). A closer examination revealed that about one third of repetitions (32.4%) 245 involved changes to a single attribute, predominantly slice position (31.9%). If 2 attributes 246 were altered (14.1%), these concerned primarily slice position and slice orientation in 247 combination (9.5%). Field of view adjustments were always accompanied by additional 248 changes (Table 2, detailed breakdown between protocols: Supplemental Table 3). 249 Comparing these adjustments with the examination guidelines, they were generally made 250 due to an incorrect or at least inferior initial setup. 251 The other half of protocol repetitions (50.2%) was performed without any manual

adjustments to the technical setup, leading to the assumption that the RT operated solely on
the grounds of subjectively low image quality, and did not attribute their subpar visual
impression to any changeable technical factor.

Across all protocols, there was a clear negative correlation between the proportion of manual setup changes and maximum classification ability as measured by AUC (**Figure 6**). The correlation coefficient was r = -0.30 ($r^2 = 0.09$), and decreased to r = -0.67 (95% CI, -0.90 to -0.13; $r^2 = 0.46$) after removal of the outlying AUC for the functional Neuro Resting State protocol, thereby demonstrating that the fewer manual setup changes were performed, the better image quality parameters were able to discriminate between baseline acquisitions and repetitions.

263 **DISCUSSION**

In our sample of the MRI substudy of the population-based NAKO study, 11,347 participants underwent whole-body MRI. Of these, 10,960 (96.6%) completed the entire MRI sequence with a mean scan time of 63.1 minutes. In 1359 participants (12.0%), the RT decided to repeat at least 1 protocol, due to incorrect technical setup parameters or based on the subjective impression of insufficient image quality. Automatically derived image quality parameters were able to discriminate between the baseline acquisitions and repetitions.

270 In general, the decision for protocol repetition will substantially depend on the RTs' 271 level of experience and expertise, especially if subjectively low image quality occurs despite 272 an objectively error-free technical setup. In a clinical setting, achieving optimal examination 273 guality by well-selected protocol repetition is instrumental to patients receiving precise and 274 correct diagnoses, and may influence therapeutic outcomes. Moreover, protocol repetition is 275 a time-costly process, verified by our results, where the mean scan time increased by 4.8 276 minutes or 7.7% for examinations with protocol repetitions, which amounted to an additional 277 94.2 hours of scan time (or 11.8 days, considering an 8-hour workday). When considering 278 even larger cohort studies or routine clinical practice, both the time losses and the cost 279 burden become considerable. Equally important is the effect of increased scan time on study 280 participants and patients, who may already be in physical or psychological distress, and 281 could benefit from shorter examinations.

282 Interestingly, in a breakdown of 3-monthly intervals, the repetition frequency 283 fluctuated over the enrollment period, ranging from a low of 4.5% to a high of 15.3%, and 284 simultaneously varied across sites from 2.3% to 28.1%. These observations remained 285 significant in a multivariate analysis and yielded 1.25-fold per year increased odds for 286 protocol repetition. This may be explained by a continuous learning process, in which the RT 287 increased their ability to identify suboptimal image quality over time, and therefore more 288 likely performed protocol repetitions as they gained professional experience within the 289 NAKO study. A close site monitoring by the NAKO administration for quality assurance,

personalized feedback to RTs, and yearly training refreshers that highlighted common
quality issues may have been contributing factors as well. We did not, however, identify
specific factors for the variations between imaging sites.

293 The most frequently repeated protocol was Cardio Cine SSFP LAX nK, known to be 294 susceptible to off-resonance or banding artifacts as well as incorrect slice positioning, and 295 possibly additionally error-prone in this particular study due to being among the last 296 protocols in a time-intensive 60-minute scanning program. On the other hand, the least 297 frequently repeated protocols were Cardio MOLLI SAX and Neuro Resting State EPI BOLD; 298 both of which are not as intuitively assessable as the rest of the acquired protocols, which 299 possibly contributed to their low rate of repetition. We were, however, unable to support this 300 claim with the study data, as the RTs' reasoning regarding protocol repetition was not 301 explicitly documented.

We identified differences in the predictive ability of the automatically derived image quality parameters regarding protocol repetition and found parameter combinations to be more effective than single parameters. Their predictive ability was particularly good for protocols in the neurological domain—this could be the result of parameters being explicitly "pure" in a static body region not prone to motion (breathing) artifacts, although less obvious factors may have contributed as well. It could also imply an above-average proficiency of the RTs in neuroradiological imaging over the remaining domains.

309 Considering previous study findings as well as our own observations of the workflow 310 challenges faced by technologists, the advantages of automated image quality assessment 311 become more evident. In a substudy of the UK Biobank cohort, 100,000 participants 312 underwent cardiovascular MRI without the implementation of automated image quality 313 control. This was tedious and time-consuming for RTs, who had to undergo multiple training 314 sessions to ensure consistent quality assessment [18]. Five years later, in the same cohort, 315 an automated image quality control using machine learning methods was tested on the first 316 10,000 brain imaging datasets and evaluated against a validation set of manually assessed 317 examinations in 5816 participants [19]. The performance of the algorithm was satisfactory

318 and subsequently reduced the need for manual checking. Particularly algorithms that rely on 319 deep neural networks (DNN) are being increasingly employed in workflow automation and 320 processing of large datasets to improve quality assurance and cost-effectiveness. A recent 321 study by Kustner et al. [20] presented a DNN for automatic, reference-free quality 322 assessment in MRI studies of the head, thorax, abdomen, pelvis, as well as whole-body 323 examinations. Used in 2911 datasets obtained from 250 patients, their framework estimated 324 image quality accurately and efficiently. Another study demonstrated good feasibility and 325 accuracy for the automated, reference-free detection of motion artifacts in MRI studies of the 326 head and abdomen via a DNN [21]. These methods are of special interest for retrospective 327 quality control in large cohort studies and, similarly to our study, could provide insight into 328 how the RTs' decision-making correlated with the automated assessments. Their, as well as 329 our, methodologies could also be implemented as prospective quality assurance tools, either 330 by providing in-line quantitative feedback to RTs or by automatically "flagging" relevant 331 acquisitions for repetition, therewith bypassing the need for manual quality assessment 332 altogether, making it reader-independent and consequently experience-independent, and 333 thus speeding up the imaging process. Even automatic adaption of acquisition settings is 334 conceivable. In its current implementation for our study, the computation of quantitative 335 image quality parameters takes too long for this type of application. Deep neural networks 336 likely hold the potential to overcome this hurdle and may consequently pave the way for 337 clinical utilization, which is our long-term goal for this concept of automated image quality 338 assessment.

Our study has certain limitations. We did not examine whether or not professional experience relates to the individual RTs' decision for protocol repetition, due to these data not being available to researchers. Based on the available quantitative data alone, we were also unable to predict which types of artifacts were associated with specific protocol repetitions—further research into their relationship will be necessary, including a visual artifact rating for the initial acquisitions that were subsequently repeated. Although we did examine the effect of image quality on protocol repetition, we did not investigate the

reciprocal effect of protocol repetition on image quality—that is, if repetitions were indeed beneficial to the resulting image data, and in which instances image quality issues may have prevailed (and whether these were of clinical relevance). However, this will be the subject of a follow-up study. Lastly, the assessment of image quality via quantitative parameters was performed as exploratory research and has yet to be validated in a clinical setting with less standardization.

In conclusion, protocol repetitions in MRI are remarkably frequent even in the highly controlled setting of a large cohort study. Automated image quality assessment shows predictive value for the RTs' decision whether or not to perform protocol repetitions, and can objectivize this multifaceted process. Especially when used in conjunction with guided or automated planning tools, it has strong potential to ensure consistent examination quality in prospective MRI studies as well as clinical practice, while simultaneously improving timeefficiency and cost-efficiency.

359

360 Data Availability Statement

361 The data sets generated during and/or analyzed during the current study are not publicly

362 available. However, data are available upon request from NAKO Transferstelle

363 (https://transfer.nako.de/ or transferstelle@nako.de) by means of a project agreement, which

364 will be subject to approval by the NAKO board.

365

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- 375
- 376 Conflicts of Interest
- None declared.
- 378

379 **REFERENCES**

- Bild, D.E., et al., *Multi-Ethnic Study of Atherosclerosis: objectives and design.* Am J
 Epidemiol, 2002. **156**(9): p. 871-81.
- Mahmood, S.S., et al., *The Framingham Heart Study and the epidemiology of cardiovascular disease: a historical perspective.* Lancet, 2014. 383(9921): p. 999 1008.
- 385 3. Volzke, H., et al., *Cohort profile: the study of health in Pomerania.* Int J Epidemiol, 2011. 40(2): p. 294-307.
- 387 4. Sudlow, C., et al., UK biobank: an open access resource for identifying the causes of
 388 a wide range of complex diseases of middle and old age. PLoS Med, 2015. 12(3): p.
 389 e1001779.
- Bamberg, F., et al., *Whole-Body MR Imaging in the German National Cohort: Rationale, Design, and Technical Background.* Radiology, 2015. 277(1): p. 206-20.
- Schlett, C.L., et al., *Population-Based Imaging and Radiomics: Rationale and Perspective of the German National Cohort MRI Study.* Rofo, 2016. **188**(7): p. 652 61.
- 395 7. Schmidt, C.O., et al., *Quality standards for epidemiologic cohort studies : An*396 *evaluated catalogue of requirements for the conduct and preparation of cohort*397 *studies.* Bundesgesundheitsblatt Gesundheitsforschung Gesundheitsschutz, 2018.
 398 **61**(1): p. 65-77.
- Kart, T., et al., Deep Learning-Based Automated Abdominal Organ Segmentation in the UK Biobank and German National Cohort Magnetic Resonance Imaging Studies.
 Invest Radiol, 2021. 56(6): p. 401-408.
- 402 9. Stocker, D., et al., *Performance of an Automated Versus a Manual Whole-Body*403 *Magnetic Resonance Imaging Workflow.* Invest Radiol, 2018. **53**(8): p. 463-471.
- 404 10. Esser, M., et al., *Performance of an Automated Workflow for Magnetic Resonance*405 *Imaging of the Prostate: Comparison With a Manual Workflow.* Invest Radiol, 2020.
 406 55(5): p. 277-284.
- 407 11. Wichmann, H.E., et al., *The German National Cohort.* Bundesgesundheitsblatt
 408 Gesundheitsforschung Gesundheitsschutz, 2012. **55**(6-7): p. 781-7.
- 409 12. Wood, M.L. and R.M. Henkelman, *MR image artifacts from periodic motion.* Med
 410 Phys, 1985. **12**(2): p. 143-51.
- Huang, L.-K. and M.-J.J. Wang, *Image thresholding by minimizing the measures of fuzziness*. Pattern Recognition, 1995. 28(1): p. 41-51.
- 413 14. Kittler, J. and J. Illingworth, *Minimum error thresholding*. Pattern Recognition, 1986.
 414 **19**(1): p. 41-47.
- 415 15. Zack, G.W., W.E. Rogers, and S.A. Latt, *Automatic measurement of sister chromatid*416 *exchange frequency.* Journal of Histochemistry & Cytochemistry, 1977. 25(7): p. 741417 753.
- Firbank, M.J., et al., *A comparison of two methods for measuring the signal to noise ratio on MR images.* Physics in Medicine and Biology, 1999. 44(12): p. N261-N264.
- 420 17. Smith, A.M., et al., *Investigation of Low Frequency Drift in fMRI Signal.* NeuroImage,
 421 1999. 9(5): p. 526-533.
- Petersen, S.E., et al., *Imaging in population science: cardiovascular magnetic resonance in 100,000 participants of UK Biobank rationale, challenges and approaches.* J Cardiovasc Magn Reson, 2013. **15**: p. 46.
- 425 19. Alfaro-Almagro, F., et al., *Image processing and Quality Control for the first 10,000* 426 *brain imaging datasets from UK Biobank.* Neuroimage, 2018. **166**: p. 400-424.
- 427 20. Kustner, T., et al., *A machine-learning framework for automatic reference-free quality* 428 assessment in MRI. Magn Reson Imaging, 2018. **53**: p. 134-147.
- 429 21. Kustner, T., et al., Automated reference-free detection of motion artifacts in magnetic
 430 resonance images. Magma, 2018. **31**(2): p. 243-256.

431 APPENDIX

432 The German National Cohort MR Imaging Study Investigators

433 MR Study Site: Augsburg, Germany

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443 MR Study Site: Essen, Germany

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483 FIGURES

Figure 1. The standard acquisition sequence of MRI protocols in the German National
Cohort (NAKO) study covers 4 anatomical domains for epidemiologic research in the
neurological, cardiovascular, thoracoabdominal, and musculoskeletal fields. The planned
scan time without protocol repetitions is approximately 60 minutes.



Figure 2. Repetition frequency (here defined as the percentage of participants with at least 1
protocol repetition) varied statistically significantly across sites, as well as over the
enrollment period.



Figure 3. Repetition frequency by protocol across all participants (n = 11,347). As an

496 example, the Neuro 2D FLAIR protocol was repeated at least once in 2.9% of all

497 participants.



500 Figure 4. The upper row presents examples of a non-contrast, axial Neuro T1w 3D 501 MPRAGE protocol, acquired according to the German National Cohort (NAKO) study 502 specifications [5]. A, Ainitial acquisition of subjectively high quality. B, An initial acquisition of 503 subjectively low quality that was subsequently repeated, and a selection of image quality 504 parameters (universal image quality (UQI), signal-to-noise ratio (SNR), and sharpness) 505 confirmed the visual impression. The lower row provides frequency distributions for these 506 parameters in initial acquisitions of the axial Neuro T1w 3D MPRAGE protocol across all 507 participants (further grouped into cases with and without subsequent repetition).



- 510 **Figure 5.** Receiver operating characteristic curves from image quality parameter
- 511 combinations for each acquired protocol (Cardio Cine SSFP LAX nK represented by 2K).
- 512 Whichever combination provided the highest area under the curve (AUC) in Supplemental
- 513 Table 2 was selected for this overview.

515

516 Figure 6. Relationship between manual setup variations and classification ability of image 517 quality parameters. On the x axis: proportion of protocol repetitions without manual setup 518 changes of all repetitions (Cardio Cine SSFP LAX-2K representative for nK). On the y axis: 519 maximum area under the curve (AUC), given by the best discriminating combination of 520 quality parameters (for initial acquisitions with and without subsequent repetition, cf. 521 Supplemental Tables 2 and 3). Red line: line of linear correlation for all protocols (correlation 522 coefficient: r = -0.30). Gray area, 95% confidence interval. In summary, the more often 523 manual setup changes occurred in the context of protocol repetitions, the less were image 524 quality parameters able to discriminate between baseline acquisitions and repetitions.

526 **TABLES**

- 527 **Table 1.** Direction of Change of Automatically Derived Image Quality Parameters for Initial Acquisitions With and Without Subsequent
- 528 Repetition (From "Without Repetition" to "With Repetition"), Along With Statistical Significance and Area Under the Curve.

Protocol	UQI	Sharpness	SNR	Specific SNR	Structured Noise Max	Structured Noise Average	N/2 Ghosting Max	N/2 Ghosting Average	Drift	Variation Over Time	Foreground Ratio
	***	▼***	▼***	▼***	***	***	A ***	▲ ***	NIA	NIA	A ***
TTW JD WIFRAGE	0.65	0.85	0.85	0.68	0.73	0.76	0.66	0.64	INA	NA	0.68
2D FLAIR	▲ ***	NS	▲ ***	▼***	▲ ***	▲ ***	▲ ***	▲ ***	NA	NA	NS
	0.58		0.83	0.74	0.69	0.73	0.69	0.67			
Resting State EPI BOLD	▲*** 0.05	▲ ***	▼***	▼***	NS	NS	▼*	▼*	▼***	NS	NS
5	0.95	0.82	0.97	0.81			0.63	0.69	0.72		
MRA 3D SPACE STIR	▼** 0.56	▼*** 0.62	NA	NA	NA	NA	NA	NA	NA	NA	NA
	0.00	▼***									
Cine SSFP LAX 2K	NS	0.57	NS	NA	NS	NS	NS	NS	NA	NA	NS
Cine SSED LAV 2K	NC	▼***	▼***	NIA	▲ ***	***	▲*	▲*	NIA	NIA	* **
CITE SOLL TAY 24	NO.	0.60	0.61	NA	0.60	0.60	0.54	0.54	INA	INA	0.54
Cine SSEP LAX /K	NS	▼*	▼**	NΔ	▲ ***	▲ ***	▲*	▲*	ΝΔ	ΝΔ	▼**
	NO	0.53	0.55		0.60	0.60	0.58	0.58			0.53
Cine SSFP SAX	NS	NS	▼***	NA	* **	***	NS	A *	NA	NA	A **
			0.63		0.59	0.62		0.55			0.56
MOLLI SAX	NS	NS	NS	NA	NS	NS	NS	NS	NA	NA	NS
T2w HASTE	NS	NS	NS	NA	NS	NS	NS	NS	NA	NA	NS
	NC	▼***	NC	NIA	NC	NC	▼**	▼**	NIA	NIA	***
	NO.	0.64	113	NA	113	113	0.59	0.58	INA	INA	0.63
Multiecho 3D VIBE	NS	▼***	NS	NΔ	NS	NS	NS	NS	ΝΔ	NΔ	▲ ***
	NO	0.57	NO	1473	NO	No	110	No	1473	1473	0.58
PDw FS 3D SPACE	▲ **	▲ *	NA	NA	NA	NA	NA	NA	NA	NA	NA
	0.55	0.54									
T2w 2D FSE	NS	NS	NS	NS	NS	NS	NS	NS	NA	NA	NS

⁵²⁹

530 Asterisks denote ranges of *P* values from Student *t* test. Numerical values represent area under the curve.

531 *P < 0.05. **P < 0.01. ***P < 0.001. NS, not significant; NA, not available; UQI, universal quality index; SNR, signal-to-noise ratio.

532 **Table 2.** Overview of the Technical Adjustments Performed Manually by RTs for Protocol Repetitions (Total N = 1606).

Protocol Repetition	n	%
Without change	807	50.2
Change in 1 category	521	32.4
RF coil configuration	8	0.5
Slice position	513	31.9
Slice orientation	0	0
FOV	0	0
Changes in 2 categories	227	14.1
RF coil configuration + slice position	65	4.0
Slice position + slice orientation	152	9.5
Slice position + FOV	10	0.6
Changes in 3 categories	51	3.2
Slice Position + slice orientation + RF coil configuration	8	0.5
Slice Position + slice orientation + FOV	43	2.7
Changes in 4 categories	0	0

533

534 Four categories of change were analyzed: RF coil configuration, slice position, slice orientation, field of view.

535 RF, radiofrequency; FOV, field of view.

536 SUPPLEMENTAL MATERIAL

537 **Supplemental Table 1.** Frequency of protocol repetitions and differences in automated

538 image quality parameters.

Protocol	All initial acquisitions	Initial acquis with re	itions	Initial acquisi without	tions repetition	р	AUC
mage quality parameter	Ν	N	Value	N	Value		
T1w 3D MPRAGE (Neuro)							
UQI	11,310	114	0.55	11,196	0.53	<.001	0.65
Sharpness			96.5		103.7	<.001	0.85
SNR			30.2		41.9	<.001	0.85
Specific SNR			130.7		155.4	<.001	0.68
Structured Noise Max			0.69		0.63	<.001	0.73
Structured Noise Avg			0.50		0.44	<.001	0.76
N/2 Ghosting Max			0.49		0.43	<.001	0.66
N/2 Ghosting Avg			0.26		0.24	<.001	0.64
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,310	114	0.40	11,196	0.38	<.001	0.68
2D FLAIR (Neuro)							
UQI	11,294	326	0.42	10,968	0.40	<.001	0.58
Sharpness			71.9		71.7	.46	0.59
SNR			25.2		33.7	<.001	0.83
Specific SNR	•	•	40.6		54.3	<.001	0.74
Structured Noise Max	11,291	324	0.74	10,967	0.67	<.001	0.69
Structured Noise Avg	•	•	0.55		0.49	<.001	0.73
N/2 Ghosting Max	•	•	0.55		0.46	<.001	0.69
N/2 Ghosting Avg	•	•	0.31	•	0.26	<.001	0.67
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,294	326	0.62	10,968	0.63	.29	0.52
Resting State EPI BOLD (Neuro)							
UQI	11,266	7	0.57	11,259	0.47	<.001	0.95
Sharpness	•	•	106.3	•	94.5	<.001	0.82
SNR	•	•	33.9	•	114.9	<.001	0.97
Specific SNR	•	•	38.4	•	115.4	<.001	0.81
Structured Noise Max	11,251	7	0.581	11,244	0.583	.95	0.49
Structured Noise Avg	•	•	0.50	•	0.54	.33	0.60
N/2 Ghosting Max	11,261	7	0.53	11,254	0.60	.04	0.63
N/2 Ghosting Avg	11,262	8	0.42	11,254	0.51	.04	0.69
Drift	11,266	7	0.002	11,259	0.006	<.01	0.72
Variation Over Time		•	0.22		0.25	.62	0.62
Foreground Ratio		•	0.72	•	0.74	.25	0.45
MRA 3D SPACE STIR (Cardio)							
UQI	11,145	51	0.37	11,094	0.38	.002	0.56
Sharpness		•	56.7	•	58.9	<.001	0.62
SNR	NA	NA	NA	NA	NA	NA	NA

Specific SNR	NA	NA	NA	NA	NA	NA	NA
Structured Noise Max	NA	NA	NA	NA	NA	NA	NA
Structured Noise Avg	NA	NA	NA	NA	NA	NA	NA
N/2 Ghosting Max	NA	NA	NA	NA	NA	NA	NA
N/2 Ghosting Avg	NA	NA	NA	NA	NA	NA	NA
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	NA	NA	NA	NA	NA	NA	NA
Cine SSFP LAX 2K (Cardio)							
UQI	11,054	392	0.294	10,662	0.295	.47	0.51
Sharpness			43.2		43.9	<.001	0.57
SNR			122.6		113.7	.17	0.53
Specific SNR	NA	NA	NA	NA	NA	NA	NA
Structured Noise Max	1.758	39	0.79	1.719	0.74	.05	0.64
Structured Noise Ava			0.79		0.74	.05	0.64
N/2 Ghosting Max	1.793	43	0.19	1.750	0.18	.32	0.53
N/2 Ghosting Avg			0.19		0.18	.32	0.53
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,054	392	0.87	10,662	0.86	.35	0.56
Cine SSFP LAX 3K (Cardio)	,			,			
UQI	11,053	392	0.282	10,661	0.284	.46	0.52
Sharpness	•		49.0	•	50.6	<.001	0.60
SNR			160.0		198.9	<.001	0.61
Specific SNR	NA	NA	NA	NA	NA	NA	NA
Structured Noise Max	10,407	350	0.69	10,057	0.65	<.001	0.60
Structured Noise Avg			0.69		0.65	<.001	0.60
N/2 Ghosting Max	10,503	352	0.22	10,151	0.21	.04	0.54
N/2 Ghosting Avg			0.22		0.21	.04	0.54
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,053	392	0.75	10,661	0.73	.004	0.54
Cine SSFP LAX 4K (Cardio)							
UQI	11,053	392	0.2787	10,661	0.2786	.95	0.50
Sharpness			46.4		46.8	.049	0.53
SNR			264.6		296.7	.003	0.55
Specific SNR	NA	NA	NA	NA	NA	NA	NA
Structured Noise Max	2,623	110	0.64	2,513	0.59	<.001	0.60
Structured Noise Avg			0.64		0.59	<.001	0.60
N/2 Ghosting Max	2,757	119	0.16	2,638	0.14	.010	0.58
N/2 Ghosting Avg	•		0.16	•	0.14	.010	0.58
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,053	392	0.87	10,661	0.88	.001	0.53
Cine SSFP SAX (Cardio)							
UQI	11,020	115	0.312	10,905	0.306	.06	0.52
Sharpness			49.0	•	49.6	.08	0.57
SNR	•		116.3	•	152.0	<.001	0.63
Specific SNR	NA	NA	NA	NA	NA	NA	NA
Structured Noise Max	10,686	107	0.82	10,579	0.79	.003	0.59
Structured Noise Avg			0.72		0.68	<.001	0.62

N/2 Ghosting Max	10,715	107	0.275	10,608	0.266	.31	0.52
N/2 Ghosting Avg			0.17		0.16	.026	0.55
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,020	115	0.80	10,905	0.78	.009	0.56
MOLLI SAX (Cardio)							
UQI	11,022	11	0.269	11,011	0.267	.90	0.54
Sharpness			42.6		43.5	.55	0.57
SNR			21.1		24.5	.66	0.56
Specific SNR	NA	NA	NA	NA	NA	NA	NA
Structured Noise Max	5,137	7	0.91	5,130	0.85	.20	0.67
Structured Noise Avg			0.91		0.85	.20	0.67
N/2 Ghosting Max	4,145	5	0.14	4,140	0.18	.31	0.63
N/2 Ghosting Avg			0.14		0.18	.31	0.63
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,022	11	0.51	11,011	0.57	.49	0.59
T2w HASTE (Body)							
UQI	11,182	82	0.20	11,100	0.21	.47	0.53
Sharpness			61.0		62.4	.23	0.55
SNR			129.1		131.4	.57	0.52
Specific SNR	NA	NA	NA	NA	NA	NA	NA
Structured Noise Max	11,182	82	0.6828	11,100	0.6832	.94	0.50
Structured Noise Avg			0.465		0.470	.27	0.54
N/2 Ghosting Max			0.35		0.36	.25	0.55
N/2 Ghosting Avg			0.22	•	0.23	.76	0.50
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,182	82	0.56	11,100	0.54	.06	0.56
T1w 3D VIBE Dixon (Body)							
UQI	11,194	94	0.35	11,000	0.36	.52	0.53
Sharpness			71.6		74.1	<.001	0.64
SNR			74.9	•	78.7	.084	0.55
Specific SNR	•		NA	•	NA	NA	NA
Structured Noise Max	•		0.66	•	0.67	.20	0.55
Structured Noise Avg	•		0.43	•	0.42	.25	0.53
N/2 Ghosting Max	•	•	0.37	•	0.41	.001	0.59
N/2 Ghosting Avg	•	•	0.20	•	0.22	.005	0.58
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,194	94	0.56	11,000	0.52	<.001	0.63
Multiecho 3D VIBE (Body)							
UQI	11,174	216	0.237	10,958	0.236	.73	0.53
Sharpness		•	57.3	•	59.1	<.001	0.57
SNR	•		115.8		113.6	.30	0.52
Specific SNR	NA	NA	NA	NA	NA	NA	NA
Structured Noise Max	9,836	186	0.577	9,650	0.577	.97	0.50
Structured Noise Avg		•	0.466	•	0.462	.50	0.52
N/2 Ghosting Max	9,869	186	0.296	9,683	0.290	.41	0.53
N/2 Ghosting Avg			0.191		0.187	.49	0.53
Drift	NA	NA	NA	NA	NA	NA	NA

Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,174	216	0.77	10,958	0.74	<.001	0.58
PDw FS 3D SPACE (MSK)							
UQI	11,346	117	0.314	11,229	0.306	.008	0.55
Sharpness	•	•	52.3		51.6	.022	0.54
SNR	NA	NA	NA	NA	NA	NA	NA
Specific SNR	NA	NA	NA	NA	NA	NA	NA
Structured Noise Max	NA	NA	NA	NA	NA	NA	NA
Structured Noise Avg	NA	NA	NA	NA	NA	NA	NA
N/2 Ghosting Max	NA	NA	NA	NA	NA	NA	NA
N/2 Ghosting Avg	NA	NA	NA	NA	NA	NA	NA
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	NA	NA	NA	NA	NA	NA	NA
T2w 2D FSE (MSK)							
UQI	11,232	33	0.354	11,199	0.351	.71	0.53
Sharpness	•		68.5		67.8	.37	0.55
SNR	•	•	28.8		29.1	.88	0.52
Specific SNR	•	•	335.5		353.1	.40	0.54
Structured Noise Max	•	•	0.47		0.46	.50	0.51
Structured Noise Avg	•	•	0.451		0.449	.78	0.49
N/2 Ghosting Max	•		0.18		0.19	.75	0.48
N/2 Ghosting Avg	•	•	0.162		0.164	.68	0.48
Drift	NA	NA	NA	NA	NA	NA	NA
Variation Over Time	NA	NA	NA	NA	NA	NA	NA
Foreground Ratio	11,232	33	0.406	11,199	0.415	.36	0.55

540 Note: Centered dot denotes the same value as above. Probability values from Student's t-

test. Bold denotes statistical significance at level p<.05. AUC: area under the curve.

Protocol	Model Number												
	1	2	3	4	5	6	7	8	9	10	11		
T1w 3D MPRAGE	0.88	0.88	0.88	0.87	0.87	0.89	0.89	0.85	0.73	0.84	0.85		
2D FLAIR	0.66	0.84	0.84	0.83	0.83	0.84	0.51	0.84	0.79	0.73	0.83		
Resting State EPI BOLD	0.82	0.97	0.97	0.97	0.96	0.95	0.77	0.97	0.81	0.88	0.99		
MRA 3D SPACE STIR	0.62	NA											
Cine SSFP LAX 2K	0.61	0.61	NA	NA	0.57	NA	0.57	0.54	NA	0.63	0.63		
Cine SSFP LAX 3K	0.64	0.66	NA	NA	0.65	NA	0.60	0.64	NA	0.64	0.62		
Cine SSFP LAX 4K	0.55	0.56	NA	NA	0.56	NA	0.56	0.55	NA	0.60	0.60		
Cine SSFP SAX	0.65	0.66	NA	NA	0.63	NA	0.56	0.66	NA	0.62	0.65		
MOLLI SAX	0.59	0.58	NA	NA	0.58	NA	0.57	0.59	NA	0.69	0.62		
T2w HASTE	0.55	0.54	NA	NA	0.54	NA	0.57	0.56	NA	0.57	0.55		
T1w 3D VIBE Dixon	0.64	0.65	NA	NA	0.64	NA	0.64	0.64	NA	0.64	0.60		
Multiecho 3D VIBE	0.58	0.59	NA	NA	0.59	NA	0.58	0.58	NA	0.59	0.54		
PDw FS 3D SPACE	0.55	NA											
T2w 2D FSE	0.54	0.54	0.59	0.58	0.55	0.57	0.55	0.55	0.55	0.59	0.58		

Protocol					Мо	del Numb	er (continu	ued)				
	12	13	14	15	16	17	18	19	20	21	22	23
T1w 3D MPRAGE	0.87	0.85	0.85	0.88	0.76	0.66	0.73	0.79	0.79	0.88	0.81	NA
2D FLAIR	0.84	0.70	0.83	0.84	0.72	0.69	0.69	0.73	0.72	0.84	0.76	NA
Resting State EPI BOLD	0.99	0.85	0.98	0.97	0.80	0.65	0.72	0.69	0.82	0.99	0.92	0.98
MRA 3D SPACE STIR	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA
Cine SSFP LAX 2K	0.62	0.57	0.58	0.59	0.64	0.53	0.64	0.64	0.64	0.63	NA	NA
Cine SSFP LAX 3K	0.66	0.61	0.61	0.65	0.60	0.54	0.61	0.61	0.61	0.66	NA	NA
Cine SSFP LAX 4K	0.61	0.58	0.60	0.61	0.60	0.58	0.60	0.60	0.60	0.61	NA	NA
Cine SSFP SAX	0.65	0.60	0.64	0.65	0.62	0.56	0.59	0.62	0.62	0.65	NA	NA
MOLLI SAX	0.71	0.69	0.81	0.81	0.67	0.63	0.77	0.77	0.77	0.81	NA	NA
T2w HASTE	0.57	0.56	0.56	0.56	0.54	0.56	0.54	0.54	0.58	0.58	NA	NA
T1w 3D VIBE Dixon	0.65	0.64	0.62	0.65	0.58	0.60	0.59	0.60	0.61	0.65	NA	NA
Multiecho 3D VIBE	0.59	0.58	0.53	0.60	0.53	0.53	0.53	0.52	0.53	0.60	NA	NA
PDw FS 3D SPACE	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA
T2w 2D FSE	0.59	0.56	0.51	0.56	0.61	0.52	0.48	0.45	0.56	0.55	0.56	NA

545

546 Note: Predictors are composed of parameter combinations as specified below. AUC values are shown for "initial acquisition with

547 subsequent repetition" versus "initial acquisition without subsequent repetition". Bold marks the maximum area under the curve

548 (AUC) value for each protocol.

549	Legend fo	r model numbers and predictors:
550	1	UQI + Sharpness
551	2	UQI + Sharpness + SNR
552	3	UQI + Sharpness + SNR + Specific SNR
553	4	Sharpness + SNR + Specific SNR
554	5	Sharpness + SNR
555	6	Sharpness + SNR + Specific SNR + Foreground Ratio
556	7	Sharpness + Foreground Ratio
557	8	SNR + Foreground Ratio
558	9	Specific SNR + Foreground Ratio
559	10	Sharpness + Structured Noise Max + Structured Noise Avg
560	11	SNR + Structured Noise Max + Structured Noise Avg
561	12	Sharpness + SNR + Structured Noise Max + Structured Noise Avg
562	13	Sharpness + N/2 Ghosting Max + N/2 Ghosting Avg
563	14	SNR + N/2 Ghosting Max + N/2 Ghosting Avg
564	15	Sharpness + SNR + N/2 Ghosting Max + N/2 Ghosting Avg
565	16	Structured Noise Max + Structured Noise Avg
566	17	N/2 Ghosting Max + N/2 Ghosting Avg
567	18	Structured Noise Max + N/2 Ghosting Max
568	19	Structured Noise Avg + N/2 Ghosting Avg
569	20	Structured Noise Max + Structured Noise Avg + N/2 Ghosting Max + N/2 Ghosting Avg
570	21	Sharpness + SNR + Structured Noise Max + Structured Noise Avg + N/2 Ghosting Max + N/2 Ghosting Avg
571	22	Specific SNR + Structured Noise Max + Structured Noise Avg + N/2 Ghosting Max + N/2 Ghosting Avg
572	23	UQI + Sharpness + SNR + Specific SNR + Structured Noise Max + Structured Noise Avg + N/2 Ghosting Max + N/2
573		Ghosting Avg + Drift + Variation Over Time + Foreground Ratio
574		

575 **Supplemental Table 3.** Detailed analysis of the technical adjustments performed manually by the radiologic technologist for protocol

576 repetitions (including any first, and, if acquired, second and third repetition).

			1 Change 2 Changes										3 Changes				
Protocol	Repetitions	No Change		Coil Conf.		Slice Pos.		Coil Slic	Conf. e Pos.	F Slice	OV e Pos.	Slice Slice	e Pos. Orient.	Coil Slice Slice	Conf. e Pos. Orient.	F Slice Slice	OV e Pos. Orient.
	N	Ν	(%)	Ν	(%)	Ν	(%)	Ν	(%)	Ν	(%)	Ν	(%)	Ν	(%)	Ν	(%)
All	1,606	807	(50.2)	8	(0.5)	513	(31.9)	65	(4.0)	10	(0.6)	152	(9.5)	8	(0.5)	43	(2.7)
T1w 3D MPRAGE	114	87	(76.3)	4	(3.5)	16	(14)	2	(1.8)	0		3	(2.6)	2	(1.8)	0	
2D FLAIR	335	191	(57.0)	0		45	(13.4)	1	(0.3)	0		96	(28.7)	2	(0.6)	0	
Resting State EPI BOLD	7	1	(14.3)	0		1	(14.3)	0		0		4	(57.1)	1	(14.3)	0	
MRA 3D SPACE STIR	51	20	(39.2)	0		31	(60.8)	0		0		0		0		0	
Cine SSFP LAX nK	417	322	(77.2)	0		41	(9.8)	0		2	(0.5)	43	(10.3)	0		9	(2.2)
Cine SSFP SAX	115	63	(54.8)	0		44	(38.3)	0		8	(7.0)	0		0		0	
MOLLI SAX	11	6	(54.5)	0		1	(9.1)	0		0		0		0		4	(36.4)
T2w HASTE	83	30	(36.1)	2	(2.4)	35	(42.2)	15	(18.1)	0		0		1	(1.2)	0	
T1w 3D VIBE Dixon	98	53	(54.1)	1	(1.0)	43	(43.9)	1	(1.0)	0		0		0		0	
Multiecho 3D VIBE	225	30	(13.3)	0		189	(84.0)	6	(2.7)	0		0		0		0	
PDw FS 3D SPACE	117	2	(1.7)	1	(0.9)	66	(56.4)	40	(34.2)	0		6	(5.1)	2	(1.7)	0	
T2w 2D FSE	33	2	(6.1)	0		1	(3.0)	0		0		0		0		30	(90.9)

577

578 Note: Four categories of change were analyzed: RF Coil Configuration, Slice Position, Slice Orientation, field of view (FOV). If more

579 than one repetition was performed, changes were evaluated against the initial acquisition.