

Service User Guide

AMRA[®] BCP Scan

Powered by AMRA Profiler 4

AMRA MEDICAL DEVICE

Europe AMRA Medical AB, Badhusgatan 5, SE-582 22 Linköping, Sweden

US AMRA Medical Inc, 150 North Michigan Avenue, Suite 1950, Chicago, IL 60601-7550, USA

Table of Contents

| | |
|---------------------------------------------------------------------------|----|
| About this Service User Guide | 5 |
| Introduction | 5 |
| Version History | 5 |
| About AMRA Profiler 4 | 6 |
| Legal Manufacturer Information | 6 |
| Intended Purpose (US) | 7 |
| Indications for Use | 7 |
| Medical Indications | 7 |
| Patient Population | 7 |
| Contraindications | 7 |
| How to use AMRA® BCP Scan service | 8 |
| Imaging Requirements | 8 |
| Data Export | 8 |
| Input of patient height on the MRI console in case of GE or Philips | 9 |
| Data Transfer | 9 |
| Upload via Ambra Gateway | 9 |
| Set-up | 9 |
| De-identification | 10 |
| Patient ID | 10 |
| Upload via Ambra share code | 11 |
| Set-up | 11 |
| De-identification | 11 |
| Patient ID | 11 |
| Manual upload via Ambra web portal | 12 |
| Set-up | 12 |
| De-identification | 12 |
| Patient ID | 12 |
| Browser Requirements | 12 |
| Registration | 13 |
| Logging In | 13 |
| Uploading Studies | 13 |

| | |
|---------------------------------------------------------|----|
| Download and View Reports..... | 14 |
| Cybersecurity..... | 15 |
| Support..... | 15 |
| The AMRA® BCP Scan report..... | 16 |
| Underlying concepts..... | 16 |
| Reference Population..... | 16 |
| Virtual Control Group..... | 16 |
| Definitions of measurements and anatomical regions..... | 17 |
| Anterior thigh muscle group..... | 17 |
| Posterior thigh muscle group..... | 17 |
| Muscle Fat..... | 17 |
| Muscle Volume..... | 17 |
| Visceral Fat..... | 17 |
| Subcutaneous Fat..... | 17 |
| Liver Fat..... | 17 |
| The different elements of the report..... | 18 |
| 1: Patient data..... | 19 |
| 2: Body composition..... | 19 |
| 3: Coronal images..... | 20 |
| 4: Detailed muscle composition..... | 21 |
| 5: Liver image..... | 21 |
| Issue categories explained..... | 22 |
| MRI data issues..... | 22 |
| Issue 1: Metal artifact..... | 22 |
| Issue 2: Body movement during scan..... | 22 |
| Issue 3: Region partly outside field of view..... | 22 |
| Issue 4: Faulty alignment of adjacent slabs..... | 23 |
| Issue 5: Fat/water signal swap..... | 23 |
| Issue 6: Positive fat bias..... | 23 |
| Issue 10: Affected by strong negative signal..... | 23 |
| Issue 99: Other uncategorized issue..... | 23 |
| Patient Data issues..... | 24 |
| Issue 11: Missing patient data..... | 24 |
| Issue 12: Unreasonable height or weight..... | 24 |

| | |
|-----------------------------------------------------------------|----|
| Reference Population Characteristics | 25 |
| Performance Specification..... | 26 |
| Appendix A: Patient Height Input - GE and Philips Consoles..... | 27 |

About this Service User Guide

Introduction

This Service User Guide and the *AMRA Profiler 4 DICOM Conformance Statement* are provided as electronic Instructions for Use at <https://amramedical.com/user-guides> upon your subscription purchase of AMRA® BCP Scan service.

This Service User Guide is a standalone document to be used for safe and effective operation of the AMRA® BCP Scan service powered by AMRA Profiler 4. This document describes how to use AMRA® BCP Scan, a service that is used to quantify global and regional fat volumes, fat fractions, and lean tissue volumes.

Version History

| Version Number | Date of Revision | Updates since last version |
|----------------|------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| 1.0 | 2022-02-21 | First version |
| 2.0 | 2022-05-03 | <p>Updated sections:</p> <ul style="list-style-type: none"> • Data Export (Subsection <i>Input of patient height on the MRI console in case of GE or Philips</i> added) • Appendix A (New) <p>General: Minor clarifications in text and typo corrections.</p> |

About AMRA Profiler 4

Legal Manufacturer Information

| | |
|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <p>Product Name: AMRA Profiler 4</p> <p>AMRA Profiler 4 is an FDA cleared medical device.</p> | <p>Manufacturer: AMRA Medical Badhusgatan 5 SE-582 22 Linköping SWEDEN www.amramedical.com support@amramedical.com</p> |
| <p>Clinical diagnosis should not be based solely on results shown in AMRA Profiler 4 reports.</p> <p>Results may vary slightly in different versions of AMRA Profiler 4.</p> <p>Any serious incident that has occurred in relation to the device should be reported to AMRA at support@amramedical.com and the competent authority of the Member State in which the healthcare institution and/or patient is established.</p> <p>Federal law restricts this device to sale by or on the order of a licensed physician.</p> | |

Intended Purpose (US)

Indications for Use

Indicated for use as a magnetic resonance diagnostic device software application for non-invasive fat and muscle evaluation that enables the generation, display and review of 2D magnetic resonance medical image data.

Designed to utilize DICOM 3.0 compliant magnetic resonance image datasets, acquired from compatible MR Systems, to display the internal structure of the body including the liver. Other physical parameters derived from the images may also be produced.

Provides a number of quantification tools, such as Region of Interest (ROI) placements, to be used for the assessment of regions of an image to quantify liver tissue characteristics, including the determination of fat fraction in the liver, T2* and muscle volume.

These images and the physical parameters derived from the images, when interpreted by a trained clinician, yield information that may assist in diagnosis.

Medical Indications

- Health and wellness

Patient Population

- Adults

Contraindications

Not applicable, software only.

How to use AMRA[®] BCP Scan service

Imaging Requirements

AMRA[®] BCP Scan service has the following requirements for use:

- Ambra Health platform (DG PACS, K152977), a web-based tool for data transfer, set-up to communicate with AMRA
- Pdf-reader installed on the computer in order to read/view the AMRA[®] BCP Scan Report

AMRA[®] BCP Scan service has the following requirements for the input DICOM data acquired with the MRI scanner:

- MRI scanner from one of the following manufacturers: GE, Siemens, or Philips
- MRI images scanned with 1.5 T or 3.0 T main magnetic field strength
- 2-point Dixon data including fat, water, in-phase and opposed-phase images
- Images acquired with inherent transmit/receive body coil unless otherwise specified
- Neck-to-knee coverage in the head-feet direction, and field-of-view that covers subcutaneous tissue of the abdomen and thighs
- A voxel volume of minimum 1.75³ mm³
- A voxel volume of maximum 4.5³ mm³
- An overlap between neighboring slabs of at least 3 cm
- T1-weighted images with a flip angle of 10 degrees and short repetition time
- DICOM files as an archive compliant with the *DICOM Conformance Statement*
- Fat, water, in-phase and opposed-phase image separation identifiable by Series Description or Image Type DICOM tags
- For liver fat assessment: Scanner manufacturer's clinical liver application images acquired with factory settings including fat fraction, fat, water and T2*/R2* images, or a multi-echo gradient-echo acquisition of the liver with at least 3 in-phase echo times

For detailed information on the DICOM image requirements please refer to the *DICOM Conformance Statement* provided at <https://amramedical.com/user-guides> .

Data Export

- Body-slabs: Export fat, water, in-phase and opposite-phase images
- Liver image acquisition for liver fat: Export fat, water, fat fraction and R2* images, as well as in-phase/opposite-phase (GE/Philips) or Original echoes (Siemens)
- Note that compression or enhanced DICOM are not supported
- Transfer complete datasets, including body slabs and liver slabs as one study
- If alternative liver image acquisition for liver fat, T2s (T2*), is used, export all corresponding images (multi-echo series or separate spoiled GRE images with different TEs)

Exported DICOM files must comply with the *DICOM Conformance Statement* that can be found at <https://amramedical.com/user-guides>. Note, that information on the patient's sex, height and weight needs to be included in the exported DICOM data. This information is typically typed in at patient registration on the MRI console.

Input of patient height on the MRI console in case of GE or Philips

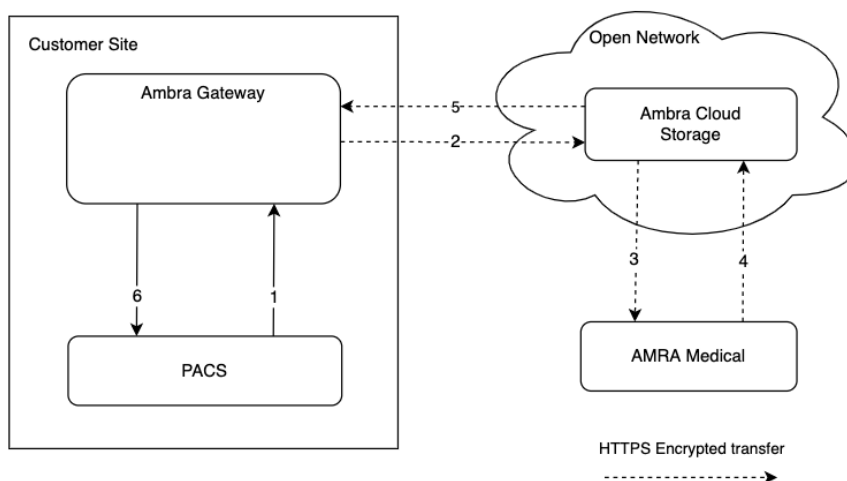
In case of GE or Philips scanners, there is no dedicated field to input patient height on the MRI console. The recommended solution is then to add the height information in the tags Study Comments (Philips) or Additional Patient History (GE) instead. See Appendix A in this document for a printable specification on how the information then needs to be typed in, in order for the information to be correctly interpreted and used by AMRA. If manual data transfer is done (see next chapter), the patient height can be input in the Ambra web portal instead.

Data Transfer

AMRA uses the Ambra Health platform for data transfer. AMRA recommends that one of the following three transfer solutions are used to upload the de-identified data to the Ambra Cloud; direct upload from PACS via Ambra Gateway, transfer between Ambra accounts via share code, or manual upload to the Ambra Web Portal. In this chapter, “report” refers to the BCP Scan Report.

Upload via Ambra Gateway

This is the flow-chart of the setup when using upload from PACS via Ambra Gateway.



Automatic data transfer via Ambra Gateway contains the following steps:

1. Push of images, including sex, weight and height of the patient, from PACS to the Ambra Gateway
2. Automatic upload of de-identified DICOM images to Ambra Cloud Storage
3. Automatic download of DICOM images from Ambra Cloud Storage to AMRA
4. Automatic upload of a report to Ambra Cloud Storage
5. Automatic download of the report to Ambra Gateway
6. Automatic push of the report to the patient exam in PACS (based on Study Instance UID)

SET-UP

For setup of the Ambra Gateway solution, contact your PACS or IT administrator.

The following procedures are needed for the setup of Ambra Gateway.

1. The Ambra Gateway is installed on a machine on the local network of the site
2. The Ambra Gateway is connected to the site's PACS and the PACS is configured to push images to the Ambra Gateway (either manually or automatically)
3. The Ambra Gateway is configured in the AMRA account of Ambra Cloud Storage

4. Configuration is added to convert the PDF report into a DICOM secondary capture, including Study Instance UID metadata.

Note that it is possible to get a notification if data was not successfully transferred from Ambra Gateway to the Ambra cloud. It is recommended that this notification is enabled at all times and that the receiver(s) of this notification is specified and communicated to Ambra Health at set-up.

DE-IDENTIFICATION

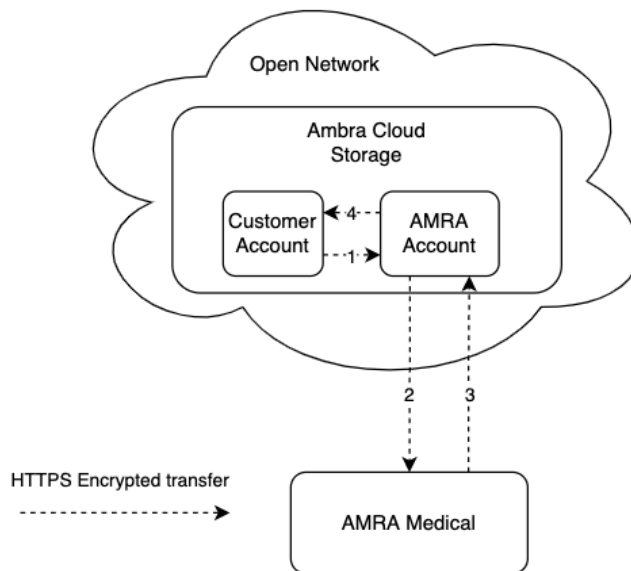
The data shall be de-identified before upload, so that it does not contain any unnecessary personal information. The site is responsible for de-identification of the data. When using the Ambra Gateway solution, the de-identification of data can be done either before the data is pushed to the Ambra Gateway, or within the Ambra Gateway. If it is done within the Ambra Gateway, the site needs to agree with Ambra at set up, on which DICOM tags shall be de-identified. AMRA will not actively screen for unnecessary personal information in the received data. If AMRA detects unnecessary personal information, the whole dataset will be deleted and the site asked to resend the images. The *DICOM Conformance Statement* states which DICOM tags are required in order to generate the BCP Scan Report output.

PATIENT ID

When using upload via Ambra Gateway, the Patient ID tag needs to be populated with a string or number, unique for the examination, that is easily accessible for the site, such as the Accession Number (0008,0050). The information in the Patient ID tag will be presented in the report, in the Patient Data section, and it will also be used to identify a specific dataset when AMRA communicates with you. Always refer to the de-identified Patient ID if there is a need to contact AMRA support about a specific dataset.

Upload via Ambra share code

This transfer option can be selected if the site is already using Ambra and have an account. The data transfer can then be done via share code in the Ambra cloud. This is the flow-chart when using upload via share code.



Automatic data transfer via Ambra share code contains the following steps:

1. De-identified DICOM images, including sex, weight and height of the patient, are shared to AMRA's Account, via share code and the routing rule set up
2. Automatic download of DICOM images to AMRA
3. Automatic upload of the de-identified report to AMRA's Account
4. Automatic share of the report to the Customer's Account, via share code and routing rule set up

SET-UP

For set-up of the share code solution, a share code and routing rule needs to be setup for the Customer Account as well as for the AMRA account in Ambra cloud. Contact your local PACS or IT administrator.

DE-IDENTIFICATION

The data shall be de-identified before upload, so that it does not contain any unnecessary personal information. The site is responsible for de-identification of the data. When using the share code solution, the de-identification of data shall be done before data is pushed to the AMRA Account. The site needs to agree with Ambra at set up, on which DICOM tags shall be de-identified. AMRA will not actively screen for unnecessary personal information in the received data. If AMRA detects unnecessary personal information, the whole dataset will be deleted and the site asked to resend the images. The DICOM Conformance Statement states which DICOM tags are required in order to generate the BCP Scan Report output.

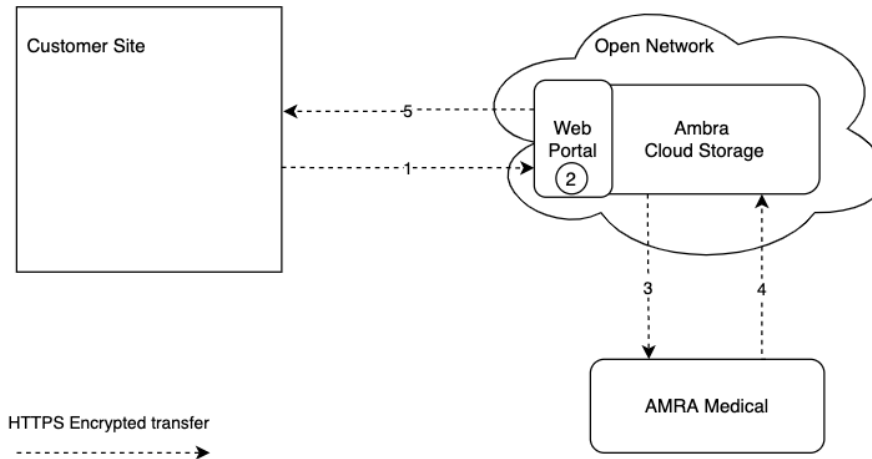
PATIENT ID

When using upload via share code, the Patient ID tag needs to be populated with a string or number, unique for the examination, that is easily accessible for the site, such as the Accession Number (0008,0050).

The information in the Patient ID tag will be presented in the report, in the Patient Data section, and it will also be used to identify a specific dataset when AMRA communicates with you. Always refer to the de-identified Patient ID if there is a need to contact AMRA support about a specific dataset.

Manual upload via Ambra web portal

This is the flow-chart for manual upload to Ambra web portal.



Manual data transfer via Ambra Web Portal contains the following steps:

1. De-identified DICOM images, including sex, weight and height of the patient, are manually uploaded to the Ambra Web Portal
2. Patient ID, sex, height and weight can be modified in the web portal, if needed
3. Automatic download of DICOM images to AMRA
4. Automatic upload of the de-identified report to Ambra Cloud Storage
5. Manual download of the report from the Ambra Web Portal

SET-UP

The only requirement to upload images manually to Ambra is a computer with Internet access. Browser requirements and instructions for the different steps (registration, log in, upload, download and view reports) are described later in this chapter.

DE-IDENTIFICATION

The data shall be de-identified before upload, so that it does not contain any unnecessary personal information. The site is responsible for de-identification of the data. AMRA will not actively screen for unnecessary personal information in the received data. If AMRA detects unnecessary personal information, the whole dataset will be deleted and the site asked to resend the images. The *DICOM Conformance Statement* states which DICOM tags are required in order to generate the BCP Scan Report output.

PATIENT ID

When using the manual upload solution, a unique identifier must be used for each patient, in the Patient ID tag. The information in the Patient ID tag will be presented in the report, in the Patient Data section, and it will also be used to identify a specific dataset when AMRA communicates with you. Always refer to the de-identified Patient ID if there is a need to contact AMRA support about a specific dataset.

BROWSER REQUIREMENTS

Recommended browser: Google Chrome

Other compatible browsers: Microsoft Internet Explorer 9 or later, Microsoft Edge, Apple Safari and Firefox.

REGISTRATION

You will be sent an e-mail invite where you will be asked to register for the Ambra platform. The invite is sent from support@amramedical.com, please check your spam folder if you haven't received the invite after service subscription startup.

LOGGING IN

To log in, go to: <https://amra.ambrahealth.com/>
Your email address will be your Login.

Sign In

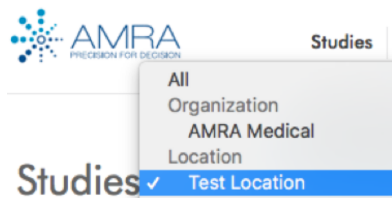
Login

Password

[Forgot your password?](#)

UPLOADING STUDIES

1. From the Studies drop-down, select the appropriate location/site. You will only see the locations that are relevant to your studies. In the startup email, you will receive information about the location name.



2. Click 'Add Study' then 'Upload Studies'.



3. During manual upload you will have the chance to update specific DICOM tags. The resulting BCP Scan report will be based on the given metadata in these fields, so make sure that they are correct.

Choose studies to upload

[Choose File...](#)

| Description | Images | Patient Name | DOB | Modality | Study Date | |
|--------------------------------------------------------|--------|--------------|------------|----------|------------|--|
| <input checked="" type="checkbox"/> Certification scan | 96 | Anon | 1990-01-01 | MR | 2015-09-29 | |

Patient ID

New Patient ID

Patient Size

New Patient Size

Enter the patient's size (##.#) in meters.

Patient Sex

New Patient Sex

Patient Weight

New Patient Weight

Enter the patient's weight (###.#) in kilograms.

[Upload Selected Studies](#) [Preview Selected Studies](#) [Cancel](#)

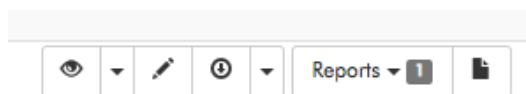
Please note, that any changes done after upload will not automatically be transferred to us. Please email support@amramedical.com alerting us to any changes.

For more help uploading, please consult Ambra's Help Center: support.ambrahealth.com

DOWNLOAD AND VIEW REPORTS

When patient data has been analyzed and results are available, you will receive an e-mail notification. You can then log-in and view/download the report. This is best accomplished with Chrome.

1. From Studies drop-down, select the appropriate site/location.
2. To view or download, click on the Reports icon from the Actions on the right-hand side of the study in the worklist.



3. Double click the desired report to download.

Cybersecurity

The BCP Scan report requires an installed pdf-reader on a computer in order to read/view the Report. This computer should have an operating system and a viewer kept up to date as well as an updated anti-virus software. Please contact your local system administrator should you have any questions regarding cybersecurity measures or if you suspect any cybersecurity incidents.

Support

If you have any questions, please contact AMRA at support@amramedical.com.

The AMRA[®] BCP Scan report

This chapter provides the user of the AMRA[®] BCP Scan report with more detailed information on the underlying concepts, measurement definitions, how to interpret the different elements of the report, and how issues in input data are handled.

Underlying concepts

Reference Population

In the AMRA[®] BCP Scan, a reference population of more than 30,000 individuals is used. Each individual in this reference population has its own set of data (height, weight, sex) and body composition measurements calculated by AMRA. The reference population is used to put the patient in a relevant context that is helpful when assessing his/her data.

Virtual Control Group

The patient specific virtual control group is extracted from the reference population by finding a group of individuals with the same sex and similar body size ($BMI \pm 2 \text{ kg/m}^2$) as the patient. With this, each patient that gets a BCP Scan will get their own individualized reference group. The virtual control group is created based on the patient data (height, weight and sex information) that is provided in conjunction with the upload of MRI data to Ambra Health.

The patient specific virtual control group is used in the report to present an individualized statistical reference (blue field) in the bar plots in the report. The blue field represents the interquartile range (25th-75th percentile) of the virtual control group and is given as a reference to help assess if the patient's value is within the statistically expected range, given its sex and body size.

Note: The patient marker being within the blue field, does not indicate if the patient is healthy or unhealthy.

Note: If a patient has been scanned at two different timepoints, and a comparison of reports between these timepoints are made, the virtual control group and thus the blue field in bar plots can differ between the reports. For instance, if the patient has increased his/her weight between the two scans, the individuals included in the patient specific virtual control group will be slightly different between the timepoints and thus the resulting blue field will have a different distribution.

Definitions of measurements and anatomical regions

Anterior thigh muscle group

Includes quadriceps femoris, sartorius and tensor fascia latae.

Posterior thigh muscle group

Includes gluteus muscles, iliacus, adductor muscles and hamstring muscles.

Muscle Fat

Muscle Fat is the fraction of adipose tissue in the muscle, also known as muscle fat infiltration or intramuscular fat. Defined as the average proton density fat fraction in the muscle. Muscle Fat is measured in percentages (%).

Muscle Volume

The fat-free muscle volume, i.e. the volume of active tissue in the muscle. Defined as the volume of all voxels with fat fraction <50%. The Muscle Volume is measured in liters (L).

Visceral Fat

Defined as the volume of adipose tissue within the abdominal cavity, excluding adipose tissue outside the abdominal skeletal muscles and adipose tissue and lipids within the cavity and posterior of the spine and back muscles. The Visceral Fat is measured in liters (L).

Subcutaneous Fat

Defined as the subcutaneous adipose tissue volume in the abdomen from the top of the femoral heads to the top of the thoracic vertebrae, T9. The Subcutaneous Fat is measured in liters (L).

Liver Fat

Defined as the average proton density fat fraction in regions of interest in the liver. Liver Fat is measured in percentages (%).

The different elements of the report

The BCP Scan report contains several different elements, which are described in this section.

AMRA® BCP Scan Report

1 PATIENT DATA
 Patient ID: -
 Acquisition Date: 1901-01-01 Height: 1.93 m
 Sex: Male Weight: 105.7 kg
 Age: 38 years BMI: 28.4 kg/m²

2 BODY COMPOSITION

| Measurement | Value | Percentile |
|------------------|---------|------------|
| Visceral Fat | 5.41 L | ~50 |
| Subcutaneous Fat | 9.43 L | ~75 |
| Muscle Fat | 5.46 % | ~25 |
| Muscle Volume | 15.97 L | ~75 |
| Liver Fat | 13.65 % | ~75 |

3 **Image Legend**

- Visceral Fat
- Subcutaneous Fat
- Left Anterior Thigh Muscle
- Right Anterior Thigh Muscle
- Left Posterior Thigh Muscle
- Right Posterior Thigh Muscle

How to read the AMRA® BCP Scan Report

This report is generated by AMRA® Profiler 4, which is a tool that measures thigh muscle composition and abdominal fat distribution using magnetic resonance imaging (MRI) data. The report provides patient specific Body Composition Profile (BCP) measurements; subcutaneous and visceral fat volume, muscle fat, muscle volume and liver fat. It also visualizes the patient's data in comparison to reference data, in order to enable assessment of the results in a relevant context.

MEASUREMENTS AND BAR PLOTS
 Each measurement is visualized in a bar plot, where the patient's location (black dot) is presented in relation to the distribution within a general sex-specific reference population. The bar also shows the expected range for the patient, given the patient's sex and body size (blue bar).

IMAGE QUALITY ISSUES
 In rare cases, there can be one or more image quality issues present in the MRI data that is used as input to calculate one or more measurements presented in this report. As a consequence, the quality of the measurements cannot be guaranteed to be within the stated performance and the affected measurements are therefore not reported. In the event of an image quality issue in the MRI data, a short description of the identified issue will be presented.

Additional measurements and visualizations

4 DETAILED MUSCLE COMPOSITION

| Measurement | Value | Percentile |
|-----------------------------------------|--------|------------|
| Left Anterior Thigh Muscle Fat (%) | 6.50 % | ~50 |
| Right Anterior Thigh Muscle Fat (%) | 4.41 % | ~25 |
| Left Posterior Thigh Muscle Fat (%) | 9.84 % | ~75 |
| Right Posterior Thigh Muscle Fat (%) | 8.87 % | ~50 |
| Left Anterior Thigh Muscle Volume (L) | 2.91 L | ~50 |
| Right Anterior Thigh Muscle Volume (L) | 2.84 L | ~50 |
| Left Posterior Thigh Muscle Volume (L) | 5.11 L | ~50 |
| Right Posterior Thigh Muscle Volume (L) | 5.11 L | ~50 |

5 PLACEMENT OF ROIS USED FOR CALCULATION OF LIVER FAT

FOR MORE INFORMATION ON

- Underlying concepts and definitions
- Reference population used
- Performance specification
- Image quality issue categories

PRODUCT INFORMATION
 Generated by AMRA® Profiler 4, version 2021.10.32 (release: 3116d4568).
 Reference population ID: 10.
 Results may vary slightly in different versions of AMRA® Profiler 4. Release updates can be requested for details.
 AMRA® Profiler 4 is an FDA cleared medical device.
 Clinical diagnosis should not be based solely on results shown in this report.

BCP Scan report, with references to the following descriptions of each element.

1: Patient data

The Patient Data element presents the patient specific metadata. This data is used as ground for the visual elements presented in the report.

| Information | Source of information (DICOM tag or other) | Possible to modify in case of manual upload (YES/NO) |
|------------------|------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------|
| Patient ID | (0010,0020) Patient ID | YES |
| Acquisition Date | (0008,0022) Acquisition Date Attribute | NO |
| Sex | (0010,0040) Patient's Sex | YES |
| Age | (0010,1020) Patient's Age | NO |
| Height | (0010,1020) Patient's Size (m) | YES |
| Weight | (0010,1030) Patient's Weight (kg) | YES |
| BMI | Weight / Height ² (kg/m ²) Calculated by AMRA, given the received Weight and Height numbers. | NO |

In case any of the Patient Data was omitted at upload, the value is replaced by a dash “–” in the Patient Data element.

2: Body composition

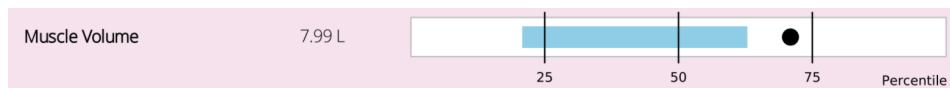
Measurements

The body composition measurements give an objective measure of the fat distribution and muscle composition within the patient's body, allowing for assessment of body composition changes. The measurements may assist in diagnosis, prediction and prognosis.

The Muscle Fat measurement is calculated as the mean muscle fat infiltration in the anterior thigh muscle groups in both legs. A high value means there is a higher fraction of fat in the muscles, which indicates a lower muscle quality. The Muscle Volume measurement is calculated as the thigh fat-free muscle volume of both the anterior and posterior thigh muscle groups. A low value may indicate that there is low quantity of fat-free muscle tissue.

Bar plots

Each body composition measurement is visualized in a bar plot to help put the patient's measurement in a relevant context. The patient's measurement (black dot) is compared to the distribution of measurements within a sex specific reference population (white bar). The 25th, 50th and 75th percentiles of this reference population are presented as solid lines. The distribution within the patient's virtual control group (blue bar) shows the expected range of the measurement, given the patient's sex and body size at time of scan.



Example: The patient's Muscle Volume is high compared to individuals in the reference population with the same sex (close to the 75th percentile). It is also above the expected, given the patient's sex and body size (well to the right of the blue field).

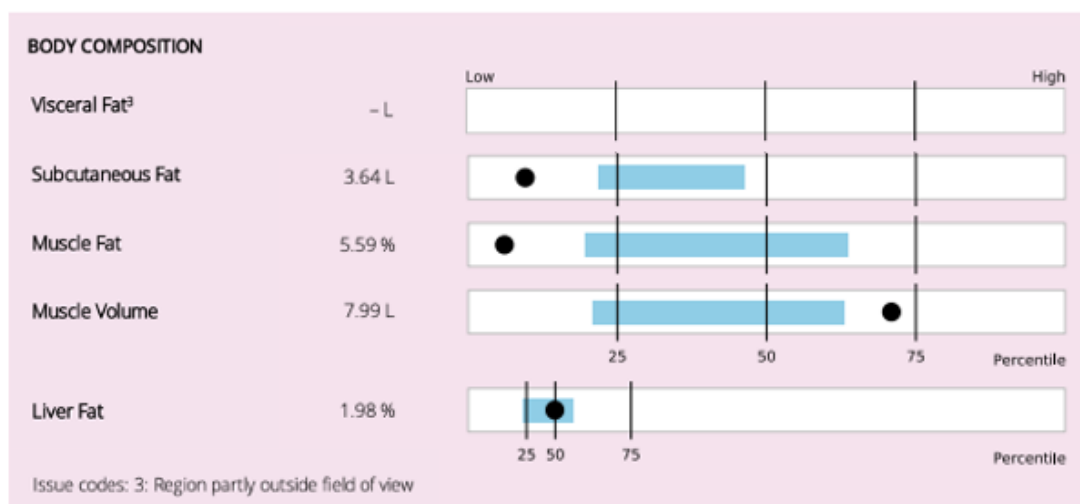
Liver fat has a skewed distribution in a general population, where most individuals have little liver fat. Therefore, the Liver Fat bar plot is scaled differently than the other bars plots, for easier interpretation.

Note: The patient marker being within the blue field, does not indicate if the patient is healthy or unhealthy.

Handling of identified input data issues

If issues in the MRI data are identified, the quality of the measurements cannot be guaranteed to be within the stated performance and the affected measurement(s) are therefore not reported. This is indicated by a dash, “-”, instead of a value and an empty bar plot will be output, with no patient marker or expected field visualized.

If there is an issue identified with the provided Patient Data, values will be presented, but an empty bar plot will be output, with no patient marker or expected field visualized. A notification is presented at the bottom of the bar plot element, with a short description of the identified issue.



Example: Issues with the MRI scan resulting in that the Visceral Fat measurement could not be calculated within stated performance. A dash is output instead of a value and an empty bar plot is presented.

Single leg used in case of identified issue in one of the legs

Note: If there is an issue identified for one of the legs, only the leg without issues is used for muscle composition calculations in this element (Muscle Fat and Muscle Volume with corresponding bar plots). In such cases, a notification is presented, on which leg was used. This information can be of importance if comparing two BCP Scan reports from two different timepoints for the same patient. When this happens, it is also clear from the Detailed muscle composition section on page 2, that one of the legs could not be assessed.

3: Coronal images

Representative coronal fat and water images of the patient, where the segmented volumes used to calculate the measurements are visualized with different colors. The color of each segmentation is explained in the image legend next to the images. This gives the reader an overview of the muscle and fat status within the patient's body.

4: Detailed muscle composition

Measurements

The detailed muscle composition section presents the separate muscle composition values for the anterior and posterior thigh muscle groups as complementary information, allowing for assessment of differences in muscle composition between the legs.

Bar plots

Each muscle composition measurement is visualized in a bar plot to help put the patient's measurement in a relevant context. The patient's measurement (black dot) is compared to the distribution of measurements within a sex specific reference population (white bar). The 25th, 50th and 75th percentiles of this reference population are presented as solid lines. The distribution within the patient's virtual control group (blue bar) shows the expected range of the muscle measurement, given the patient's sex and body size at time of scan.

Note: The patient marker being within the blue field, does not indicate if the patient is healthy or unhealthy.

Handling of identified input data issues

If issues in the MRI data are identified, the quality of the measurements cannot be guaranteed to be within the stated performance and the affected measurement(s) are therefore not reported. This is indicated by a dash, “-”, instead of a value and an empty bar plot will be output, with no patient marker or expected field visualized.

If there is an issue identified with the provided Patient Data, values will be presented, but an empty bar plot will be output, with no patient marker or expected field visualized. A notification is presented at the bottom of the bar plot element, with a short description of the identified issue.

5: Liver image

A representative slice of the liver is presented, with the ROIs placed in this slice visualized. The ROIs are a subset of the total number of ROIs used for calculation, which are evenly distributed within the liver. The image is either a Proton Density Fat Fraction (PDFF) image or a fat image, depending on which image was provided for Liver Fat calculations.

Issue categories explained

Input data issues can occur either in the MRI data or in the Patient Data. The following sections explain more about the different issue categories. AMRA proposes remedial actions and how issues can be prevented going forward.

MRI data issues

| Issue code | Short description given in report |
|------------|-------------------------------------|
| 1 | Metal artifact |
| 2 | Body movement during scan |
| 3 | Region partly outside field of view |
| 4 | Faulty alignment of adjacent slabs |
| 5 | Fat/water signal swap |
| 6 | Positive fat bias |
| 10 | Affected by strong negative signal |
| 99 | Other uncategorized issue |

Table of issue categories used in AMRA® BCP Scan report to identify issues in the MRI data.

ISSUE 1: METAL ARTIFACT

The measurable region contains an imaging artifact specific to the presence of metal within the imaging volume, with an adverse effect on image quality. This can occur if the patient has a metal implant/prosthesis/device, metal in their clothing or jewelry/piercings have not been removed.

Remedial action: Rescan all sequences if the metal is removable. If the metal is an immovable implant, prosthesis, or device, no rescan of the patient shall be done.

To prevent recurrence: Ensure all patients are free from metallic objects, jewelry and piercings and that clothing is metal free, prior to MRI acquisition.

ISSUE 2: BODY MOVEMENT DURING SCAN

The measurable region contains motion imaging artifacts, with an adverse effect on image quality. This can occur if any part of the patient's body moves during the scan.

Remedial action: Rescan all sequences.

To prevent recurrence: Ensure the patient remains as still as possible during the scan and repeat affected sequences at MRI acquisition, paying particular attention to the patient's compliance with breath-hold instructions.

ISSUE 3: REGION PARTLY OUTSIDE FIELD OF VIEW

The measurable region is wholly or partly outside of the imaging field of view. This can occur if the patient is not positioned correctly or has moved during or in-between sequences.

Remedial action: Rescan all sequences.

To prevent recurrence: Ensure the patient is positioned centered and straight on the scanner table. Ensure the arms and legs are not abducted from the body and air gaps between the limbs are minimized as far as site procedures allow. Take extra care to ensure the patient is not given any support underneath the legs. Prior to

finalizing the scan, ensure the images contain all measurable anatomy. Repeat all sequences when improved patient positioning is deemed necessary.

ISSUE 4: FAULTY ALIGNMENT OF ADJACENT SLABS

The measurable region encompasses several incongruent or misaligned imaging slabs. This can occur when the MRI sequences are manually planned/angled, table positions altered, slice thickness/number altered, or the patient has changed position during or in-between sequences.

Remedial action: Rescan all sequences.

To prevent recurrence: Ensure table positions match those provided at protocol installation. Ensure the slabs are not manually planned or angled. Ensure the patient does not move or change position and complies with breath-hold instructions.

ISSUE 5: FAT/WATER SIGNAL SWAP

The measurable region contains an imaging artifact specific to a fat and water signal swap, with an adverse effect on image quality.

Remedial action: Rescan all sequences paying particular attention to MRI imaging artifacts at acquisition. If a fat/water swap is identified at acquisition, rescan any affected sequences.

To prevent recurrence: Ensure the patient is positioned centered and straight on the scanner table. Ensure the arms and legs are not abducted from the body and air gaps between the limbs are minimized as far as site procedures allow. Take extra care to ensure the patient is not given any supports underneath the legs and clothing is metal free.

ISSUE 6: POSITIVE FAT BIAS

The measurable region contains an area of fat bias which results in an erroneously elevated fat fraction measurement. This occurs mainly with Philips' scanners with software versions < 5.4 (not supported), but can, on rare occasions, happen with other scanners.

Remedial actions: If this issue occurs, please contact AMRA support for further guidance.

To prevent recurrence: If this issue occurs, please contact AMRA support for further guidance.

ISSUE 10: AFFECTED BY STRONG NEGATIVE SIGNAL

The measurable region contains a strong negative signal close to the region of measurement, which results in erroneously lowered volume measurements. This issue can occur if there is iron present in the abdomen/intestines.

Remedial actions: None. The patient shall not be rescanned.

To prevent recurrence: None.

ISSUE 99: OTHER UNCATEGORIZED ISSUE

There is an issue with the MRI data that does not fit into any of the above categories. For more information on the issue that has occurred, please contact AMRA support.

Patient Data issues

| Issue code | Short description given in report |
|------------|-----------------------------------|
| 11 | Missing patient data |
| 12 | Unreasonable height or weight |

Table of issue categories used in AMRA® BCP Scan report to identify issues in the Patient Data.

ISSUE 11: MISSING PATIENT DATA

Required patient data height, weight, or sex, was not provided at upload to Ambra Health. The omitted patient data can be determined in the 'Patient Data' section of the report.

Remedial action: Provide the missing patient data, either by re-uploading the images with corrected metadata (in case of upload is done via Ambra Gateway or share code) or add it manually to the examination in Ambra Webportal (in case of manual upload). Notify AMRA support that modification has been done, so that a re-analysis can be triggered and a new report generated. Always refer to the de-identified Patient ID when communicating with AMRA about a specific dataset.

Note that the patient shall not be rescanned.

To prevent recurrence: Ensure patient data is complete and accurate at upload. If any further guidance is needed, please contact AMRA support.

ISSUE 12: UNREASONABLE HEIGHT OR WEIGHT

BMI calculated from the input height and weight value is outside the interval of 10-100 kg/m². This can occur if incorrect values are input at upload to Ambra Health or the values are not represented in conventional SI units.

Remedial action: Check the height and weight values presented in the 'Patient Data' section of the report and update the numbers to the correct values, either by re-uploading the examination with corrected metadata (in case of upload via Ambra Gateway or share code) or modify it manually for the examination in Ambra Webportal (in case of manual upload). Notify AMRA support that patient data has been modified, so that a re-analysis can be triggered and a new report generated. Always refer to the de-identified Patient ID when communicating with AMRA about a specific dataset.

Note that the patient shall not be rescanned.

To prevent recurrence: Ensure patient height and weight are correct at upload. If any further guidance is needed, please contact AMRA support.

Reference Population Characteristics

| | Total | Females | Males |
|-----------------------------------------|----------------|---------------|---------------|
| N | 30176 (100.0%) | 15631 (51.8%) | 14545 (48.2%) |
| Age (years) | 64.7 (7.5) | 64.1 (7.3) | 65.4 (7.6) |
| Race [White] (No.) | 29136 (96.6%) | 15099 (96.6%) | 14037 (96.5%) |
| Race [Black] (No.) | 229 (0.8%) | 130 (0.8%) | 99 (0.7%) |
| Race [Asian] (No.) | 354 (1.2%) | 134 (0.9%) | 220 (1.5%) |
| Race [Chinese] (No.) | 95 (0.3%) | 60 (0.4%) | 35 (0.2%) |
| Race [Mixed] (No.) | 139 (0.5%) | 90 (0.6%) | 49 (0.3%) |
| Race [Other] (No.) | 154 (0.5%) | 86 (0.6%) | 68 (0.5%) |
| Weight (kg) | 76.0 (15.2) | 68.9 (13.1) | 83.6 (13.4) |
| Height (cm) | 169.5 (9.2) | 163.0 (6.2) | 176.4 (6.6) |
| Body mass index (kg/m ²) | 26.4 (4.4) | 25.9 (4.7) | 26.8 (3.9) |
| Waist circumference (cm) | 88.7 (12.8) | 83.1 (12.0) | 94.6 (10.9) |
| Visceral Fat (L) | 3.8 (2.3) | 2.7 (1.5) | 4.9 (2.4) |
| Subcutaneous Fat (L) | 6.9 (3.2) | 7.9 (3.4) | 5.9 (2.5) |
| Liver Fat (%) | 4.3 (4.0) | 3.9 (3.8) | 4.8 (4.1) |
| Muscle Volume (L) | 10.2 (2.5) | 8.2 (1.2) | 12.3 (1.8) |
| Muscle Fat (%) | 8.3 (2.0) | 9.0 (1.9) | 7.6 (1.9) |
| Left Anterior Thigh Muscle Volume (L) | 1.7 (0.5) | 1.3 (0.2) | 2.1 (0.4) |
| Right Anterior Thigh Muscle Volume (L) | 1.7 (0.5) | 1.3 (0.2) | 2.1 (0.4) |
| Left Posterior Thigh Muscle Volume (L) | 3.4 (0.8) | 2.7 (0.4) | 4.0 (0.6) |
| Right Posterior Thigh Muscle Volume (L) | 3.4 (0.8) | 2.8 (0.4) | 4.1 (0.6) |
| Left Anterior Thigh Muscle Fat (%) | 8.4 (2.1) | 9.1 (2.0) | 7.6 (1.9) |
| Right Anterior Thigh Muscle Fat (%) | 8.3 (2.1) | 9.0 (2.0) | 7.6 (2.0) |
| Left Posterior Thigh Muscle Fat (%) | 12.2 (2.5) | 12.9 (2.4) | 11.4 (2.4) |
| Right Posterior Thigh Muscle Fat (%) | 11.9 (2.5) | 12.6 (2.4) | 11.2 (2.4) |
| Metabolic disease free (No.) | 7278 (24.1%) | 3774 (24.1%) | 3504 (24.1%) |
| Non-alcoholic fatty liver disease (No.) | 3860 (12.8%) | 1814 (11.6%) | 2046 (14.1%) |
| Diabetes, type 2 (No.) | 1557 (5.2%) | 510 (3.3%) | 1047 (7.2%) |
| Cardiovascular disease (No.) | 2004 (6.6%) | 535 (3.4%) | 1469 (10.1%) |
| Coronary heart disease (No.) | 1586 (5.3%) | 402 (2.6%) | 1184 (8.1%) |
| Cancer (No.) | 3177 (10.5%) | 1747 (11.2%) | 1430 (9.8%) |

Reference population characteristics. Values are mean (standard deviations).

Performance Specification

| Measurement | Accuracy | Precision | Reproducibility |
|----------------------------------------------------|----------|-----------|-----------------|
| Liver Fat | ±2.0 pp | 1.0 pp | 2.0 pp |
| Visceral Fat | ±0.5 L | 0.10 L | 0.15 L |
| Subcutaneous Fat | ±1.0 L | 0.20 L | 0.25 L |
| Muscle Volume | ±1.0 L | 0.15 L | 0.20 L |
| Left/Right Anterior/Posterior Thigh Muscle Volumes | ±0.5 L | 0.05 L | 0.10 L |
| Muscle Fat | ±2.0 pp | 0.5 pp | 1.0 pp |
| Left/Right Anterior/Posterior Thigh Muscle Fat | ±3.0 pp | 0.5 pp | 1.0 pp |

L – Liter, pp – percentage point, SD – Standard Deviation.

Performance is derived from in-vivo experiments and represents limits of performance - measured performance is normally within the limits. Liver Fat represents AMRA's fat-referenced Liver Fat method. Accuracy was assessed by comparison with established reference methods, precision using a test-retest design, and reproducibility by scanning the same subjects on multiple different MRI systems. Accuracy is reported as outer limits of the limits of agreement, while precision and reproducibility are reported as upper limits of the within-subject standard deviation.

Appendix A: Patient Height Input - GE and Philips Consoles

When registering the patient for the AMRA® BCP Scan Protocol, please insert the patient's height into the following fields on the MRI console:

GE: Additional Patient History (0010,21B0)

Philips: Study Comments (0032,4000)

Input the height in either meters or inches:

Meters: Type "HEIGHT *number* METERS", for example "HEIGHT 1.84 METERS"

Inches: Type "HEIGHT *number* INCHES", for example "HEIGHT 72.4 INCHES"

To convert feet into inches, multiply height in feet with 12.

For support, contact support@amramedical.com