

Medical image analysis: Introduction

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Outline

- What and why
- Main tasks in medical image analysis
- Image formats
- Toolkits

Outline

- What and why
 - Medical imaging
 - What is medical image analysis?
 - Why do we need medical image analysis?
- Main tasks in medical image analysis
- Image formats
- Toolkits

Medical Imaging



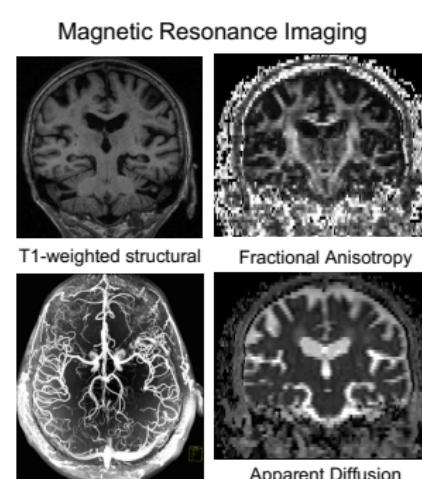
Anatomy and physiology



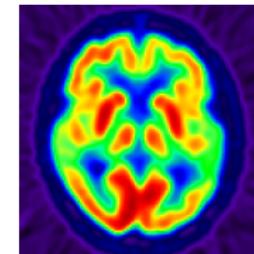
Computed Tomography



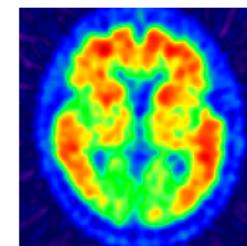
Computed Tomography
Angiography
[H. Alkhadi]



[J. Polimeni]



FDG-PET

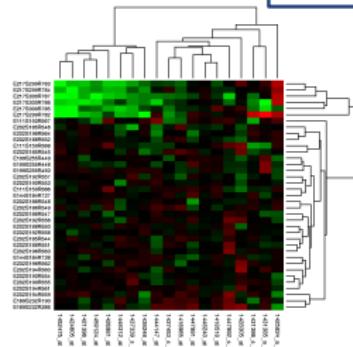
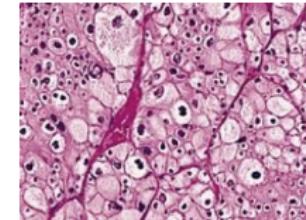
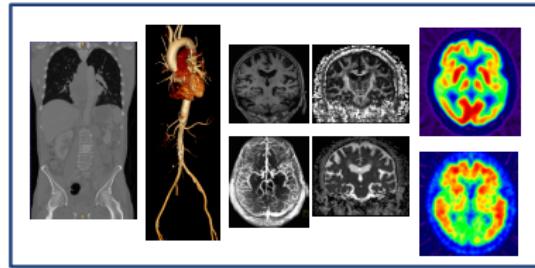
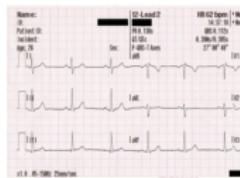


PIB-PET

└ What and why

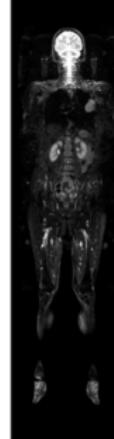
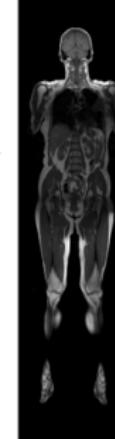
└ Medical imaging

One of many observations



Patient name:	Date:
GPCOG Screening Test	
Instructions: Please answer only the relevant questions.	
Name and Address for achievement and next of kin:	
1. I am going to give you a name and address. After I have said it, I want you to repeat what I have said back to me. John Brown, 42 West Street, London, EC1M 4BW. Please repeat what I have said.	
2. What is the date? (today's date)	
3. Clock Drawing: —— draw a circle. —— draw a cross inside the circle. —— draw a clock face with numbers 1 through 12. —— draw hands on the clock face to indicate the time of 10 o'clock. —— draw a hand pointing to the number 12. —— draw a hand pointing to the number 10.	
Information: —— ask the following questions if appropriate to the person's health. —— ask the following questions if appropriate to the person's health.	
4. What was the name and address I asked you to remember?	
Brown West Street London	
5. Name (if known):	
Hannah	
6. Date of birth (if known):	
1980	
7. Did you smoke? —— Yes (if yes, ask the following questions): —— ask the following questions if appropriate to the person's health.	
8. Patient score: Is it a significant cognitive impairment and further testing not necessary? Patient score 0-4: more information required. Proceed and do up to 6 relevant sections. Patient score 5-8: significant cognitive impairment. General practitioner or specialist advice required.	

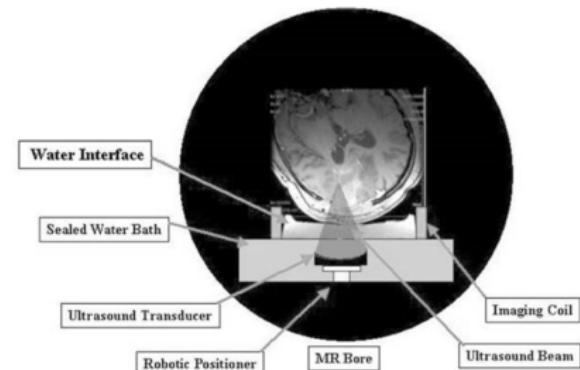
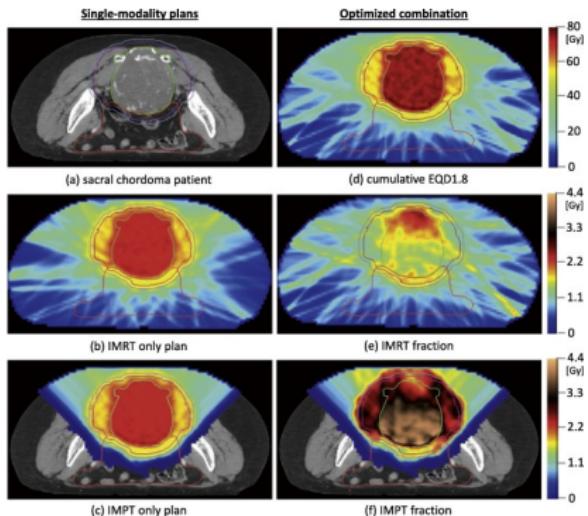
Life cycle of a medical image for diagnosis



MR Protocol Name:	MR Protocol Name:
T1, T2, FSE, PD, GRE, and T1-weighted images, and T2-weighted proton density images, and the suppressed T2 signal in the cerebellum, high-resolution T1-weighted images are obtained in sagittal, coronal and axial planes.	T1, T2, FSE, PD, GRE, and T1-weighted proton density images, and the suppressed T2 signal in the cerebellum, high-resolution T1-weighted images are obtained in sagittal, coronal and axial planes.
Field strength:	1.5T
Contrast agent:	None
Contrast agent dose:	None
Other information:	(Leave blank for collection.)
Comments:	(Leave blank.)
Study:	
Protocol:	
Protocol ID:	
Location:	(Leave blank.)
Site:	(Leave blank.)
Right:	1
Depression or elevation:	(Leave blank.)
Asymmetry:	(Leave blank.)
Adjacent structure:	(Leave blank.)
Findings:	(Leave blank.)
Organ and structures: results (Leave blank.)	(Leave blank.)
Notes:	
Long bone:	(Leave blank.)
Liver:	(Leave blank.)
Abdomen:	(Leave blank.)
Pelvis:	(Leave blank.)
Muscle:	(Leave blank.)
Breast (Leave blank.)	(Leave blank.)
Endocrine system (Leave blank.)	(Leave blank.)
Lymphatic system:	(Leave blank.)
Urinary tract:	(Leave blank.)
Female genital:	(Leave blank.)
Bone and soft tissue:	(Leave blank.)
Interpretation:	(Leave blank.)



Medical imaging in intervention



MR guided high-intensity focused ultrasound surgery

[Ram et al. Neurosurgery. 2006;59(5):949-956]

Treatment planning for photon and proton treatment

[Unkelbach et al. Radiotherapy and Oncology 2018]

[Unispital Zurich / University of Zurich]

Imaging modalities - not a complete list at all

Structural information

- Anatomical and microstructural information
- X-ray - 2D
- Ultrasound - 2D and 3D
- Computed Tomography (CT) - 3D
- Magnetic Resonance Imaging (MRI) - 2D / 3D
 - T1-weighted, T2-weighted, FLAIR, ...
 - Diffusion-weighted
 - Angiography
 - Spectroscopy

Temporal / Dynamic information

- Ultrasound
- Dynamic CT
- Dynamic MRI

Physiologic and metabolic information

- Perfusion imaging
- Flow imaging
- Functional MRI
- Positron Emission Tomography (PET)
- Single-photon Emission Tomography (SPECT)

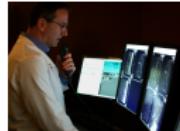
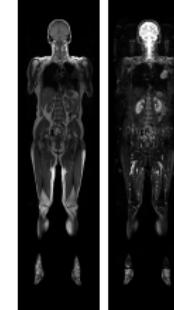
Microscopic information

- Microscopic images - histology
- Optical Coherence Tomography (OCT)
- Micro CT (μ CT)

Very rich in terms of imaging modalities
They are different sources of information
Different glasses for the same underlying tissue

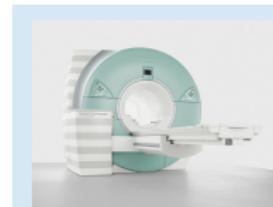
What is medical image analysis?

Algorithms for automatic interpretation of biomedical images



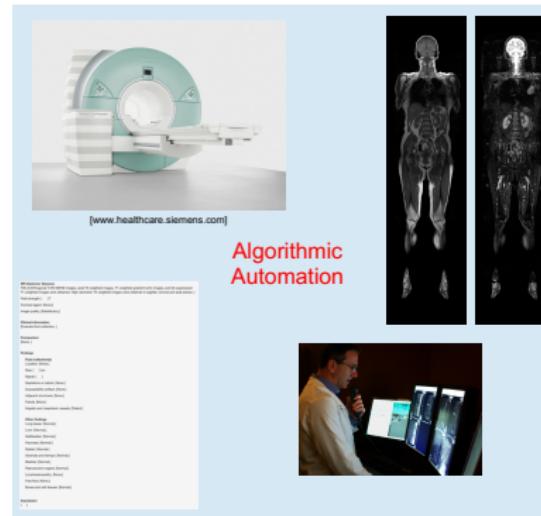
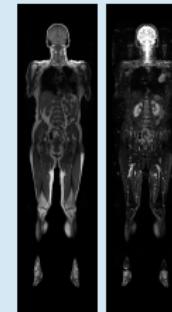
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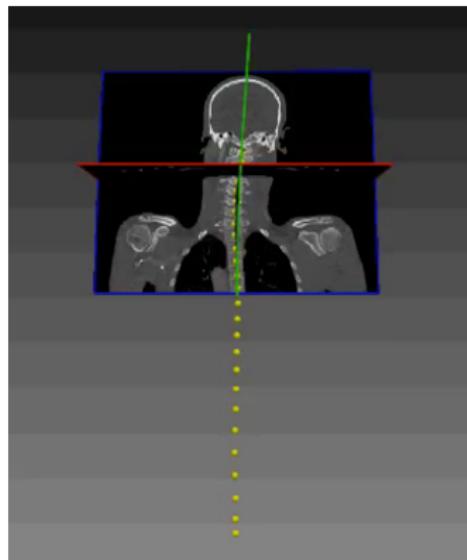


[www.healthcare.siemens.com]

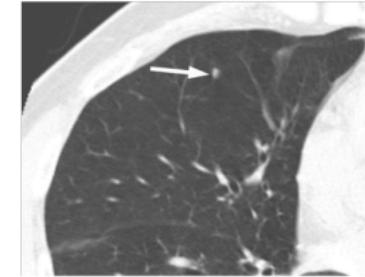
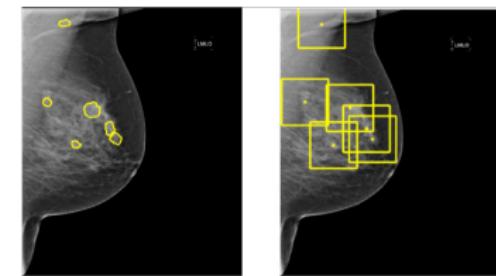
Algorithmic
Automation



Detection and localization



[Kooi et al. Medical image analysis 2017]



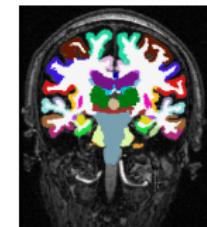
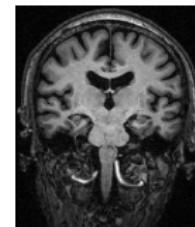
[Jeon et al. Academic Radiology 2013]

- Automatic positioning and recognition
- Visualization
- Transferring over network, search

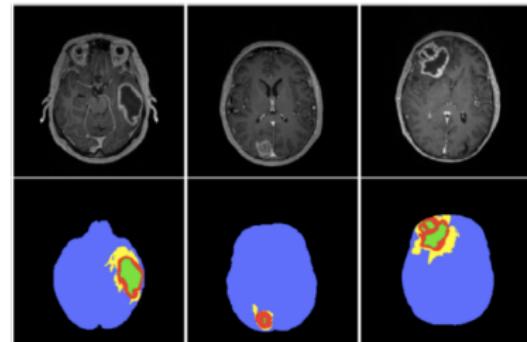
Morphological assessment and parcellation



Entire organs



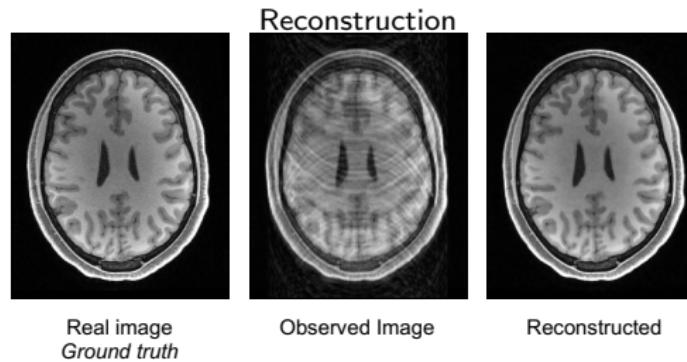
Finer scale substructures



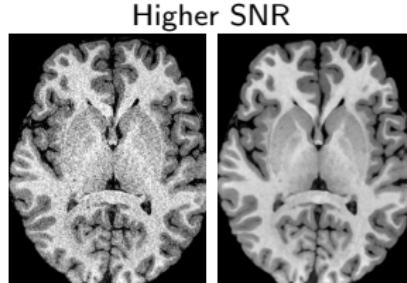
Abnormal lesions

- Morphological measurements
- Quantification for diagnosis, staging
- First step in treatment planning

Image reconstruction and enhancement



[Tezcan et al. TMI 2019]

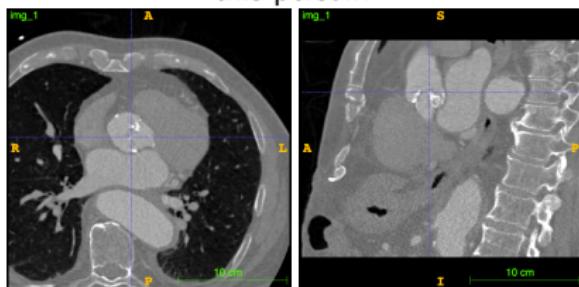


[Konukoglu et al. MICCAI 2013]

- Remove reconstruction artifacts
- Accelerate acquisition
- Improve resolution
- Improve signal-to-noise ratio (SNR)

Prediction

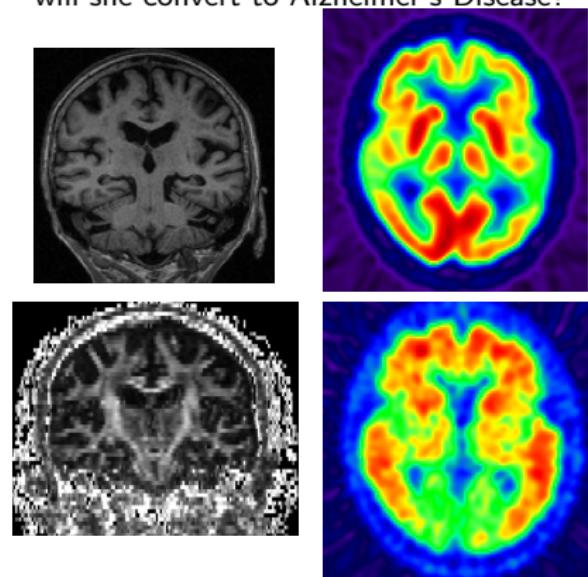
Cardiac CT with aortic calcification
Would the treatment cause a problem for
this person?



[Dan Linh, Hatem Alkhadi, Felix Tanner]

- Predicting diagnosis
- Predicting risk score
- Predicting treatment outcome

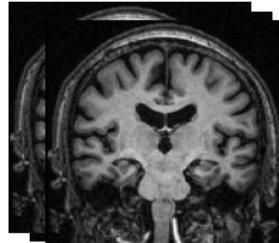
This person has Mild Cognitive Impairment
will she convert to Alzheimer's Disease?



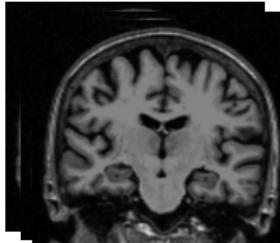
[ADNI dataset]

Statistical analysis

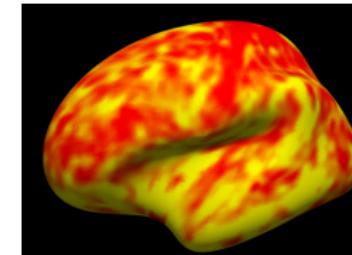
Data from population of both groups



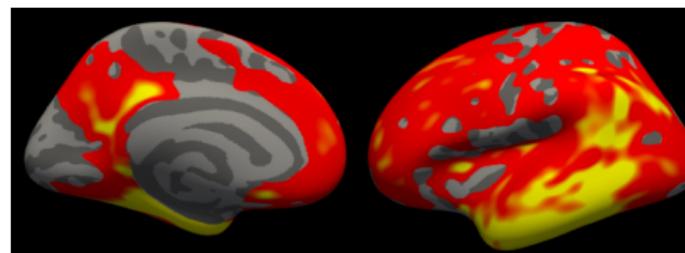
Healthy Controls



Cases



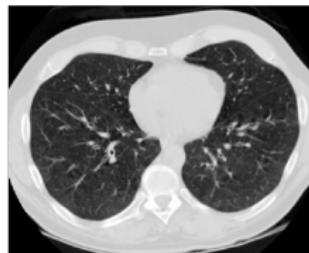
Cortical thickness



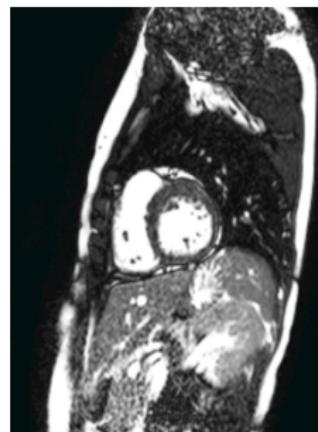
Population-wide maps showing group differences
AD effects computed from 290 subjects

Identifying anatomical / metabolic footprints of conditions

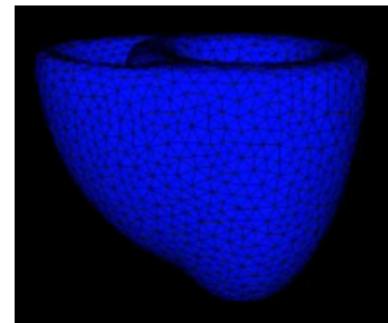
Many others: longitudinal analysis, spatio-temporal modeling,...



Has the nodule grown
between these images?
Is the change predictive
of disease?
Longitudinal analysis

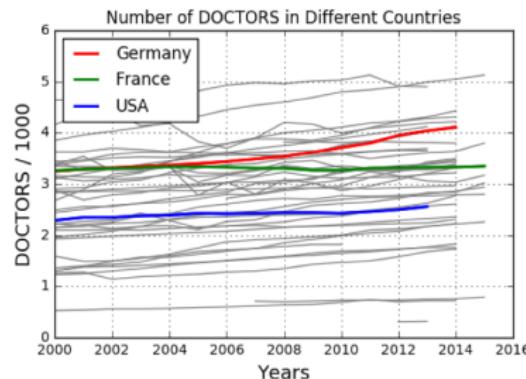
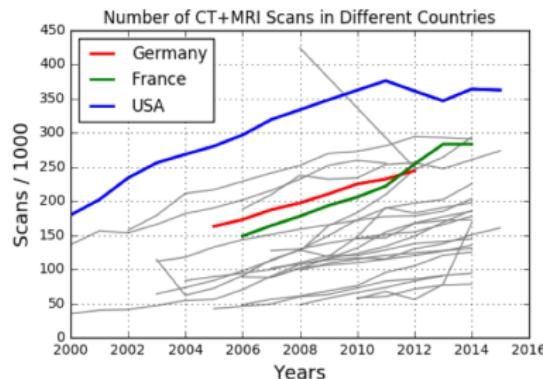


Extracting deformation fields to
understand motion



Building mathematical
models to describe and
simulate physiological
phenomena

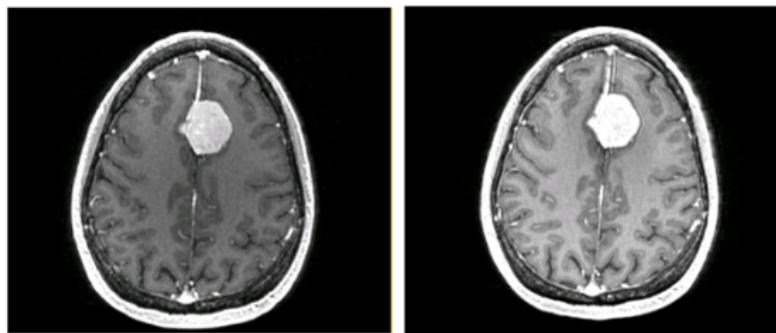
Why medical image analysis - throughput



[<https://data.oecd.org>]

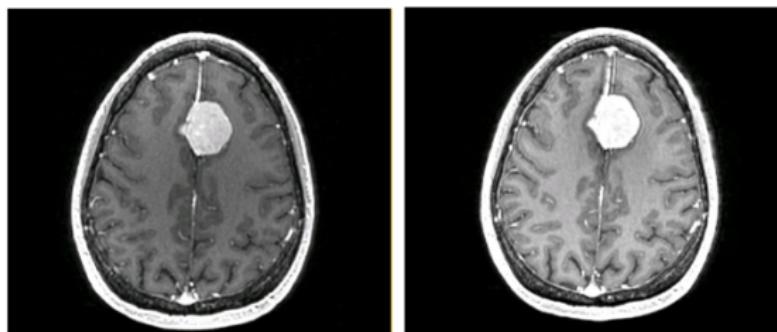
- Higher number of scans
- Finer resolution
- Higher number of modalities
- A lot of images to look at!
- Higher throughput

Why medical image analysis - quantitative analysis



1 year apart

Why medical image analysis - quantitative analysis



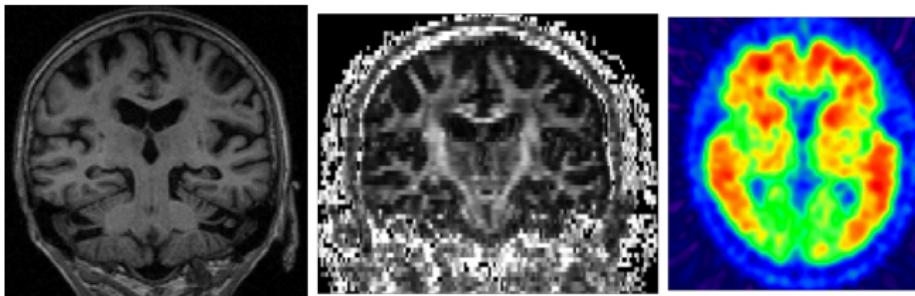
1 year apart

Growth	1%	3%	5%	11%	16%	22%
Accuracy of Detection (mean \pm std)	$8 \pm 8\%$	$6 \pm 6\%$	$28 \pm 11\%$	$44 \pm 9\%$	$52 \pm 24\%$	$88 \pm 12\%$

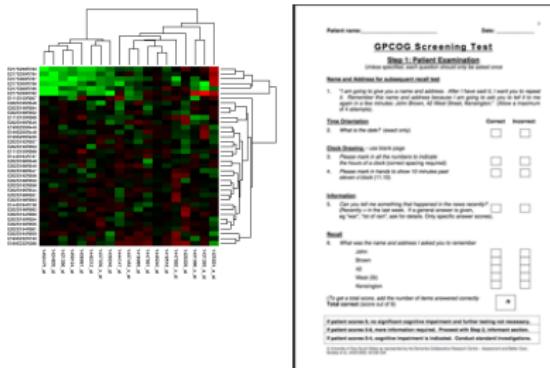
[Pohl, Konukoglu, et al. 2011]

- Subtle changes for earliest detection
- Highly accurate measurements
- Repeatability
- Changes not visible to human eye, e.g. texture

Why medical image analysis - fusion of information



different physical and metabolic characteristics



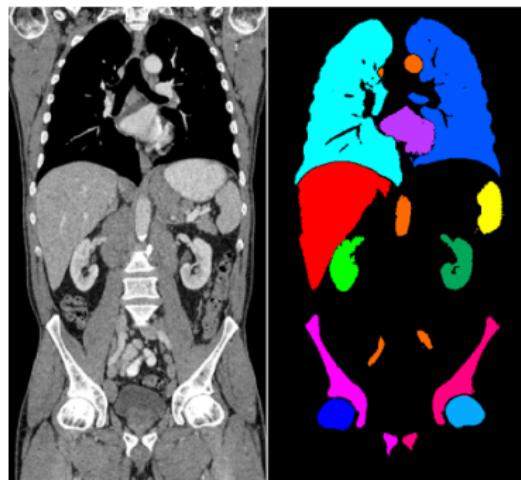
- combining information from different images
- image → numerical matrix
- fusion with other quantitative sources

Why do you want to learn medical image analysis?

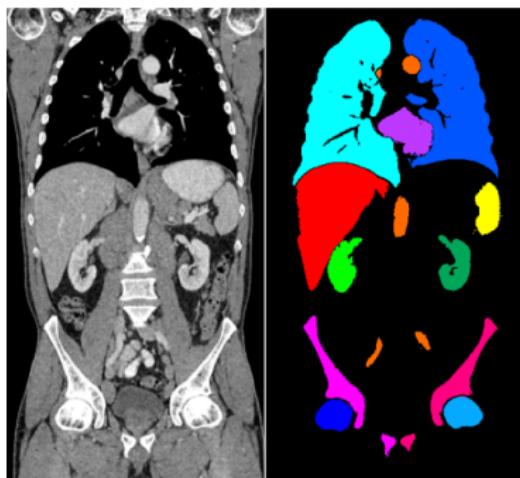
Outline

- What and why
- Main tasks in medical image analysis
 - Segmentation
 - Registration
 - Classification
- Image formats
- Toolkits

Segmentation principle



Segmentation principle



- At each pixel / voxel assign a label
- Labels can be
 - Organ: liver, spine, ...
 - Part of organs: individual vertebrae, ...
 - Lesions: tumors, pathologies, ...
- Information at each pixel
 - Intensity at and around the pixel
 - Intensity at different modalities

$$x \in \Omega \subset \mathbb{R}^d, d = 2 \text{ or } d = 3$$

$f(x) = [I(x), I(\mathcal{N}(x)), J(\mathcal{N}(x)), \dots]$: Features

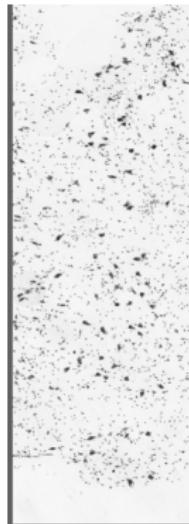
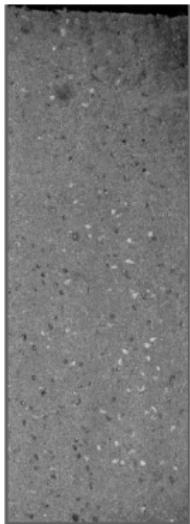
$L(x) \in \{0, \dots, N\}$: Labels

$L(x) \approx S(f(x))$: Segmentation approximating labels

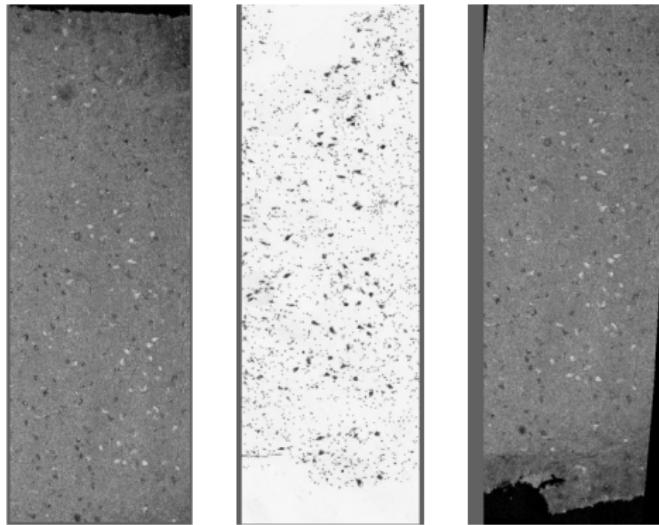
Segmentation techniques overview

- Thresholding and histogram
- Clustering
 - K-means
 - Unsupervised learning
 - Non-parametric modeling
 - ...
- Graph partitioning methods
 - Watershed
 - Graph-cuts
 - Random walker
 - Minimum spanning forest
 - ...
- Region growing
- Variational and PDE-based
 - Chan-Vese model
 - Mumford-Shah model
 - Active-shape models
 - Level sets
 - Fast marching
 - ...
- Discriminative modeling based
 - Random forest
 - Conditional random fields
 - Supervised convolutional neural networks
 - ...
- Generative modeling based
 - Expectation-maximization
 - Markov random fields
 - Atlas-based segmentation
 - Variational inference
 - ...

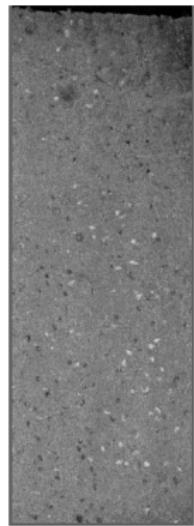
Registration principle



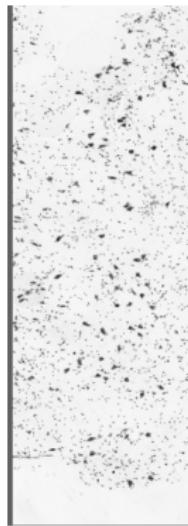
Registration principle



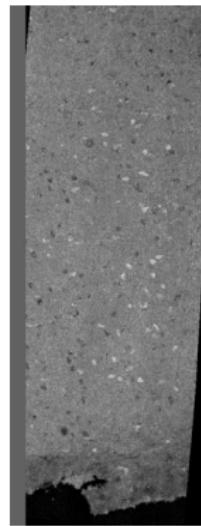
Registration principle



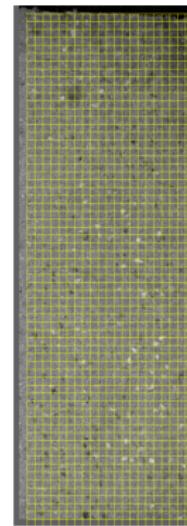
Moving



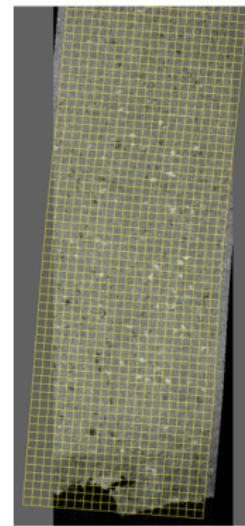
Target



Aligned

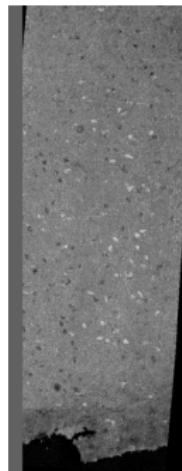
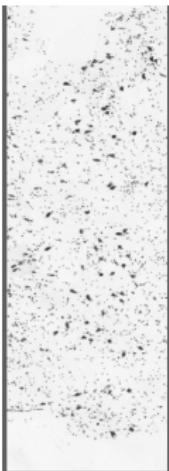
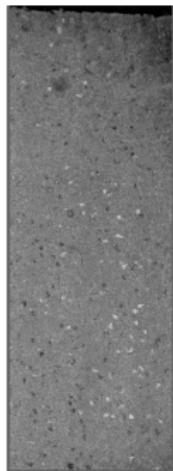


Grid



Deformed

Registration principle



Moving

Target

Aligned

- Determine spatial transformation between two images
 $I(x), J(x)$: the two images
 $T(x) : \Omega \rightarrow \mathbb{R}^d$: spatial transformation
 $J(T(x)) = I(x)$: mapping the images
- Creates a correspondence between points
- $T(\cdot)$ can be linear or non-linear
- Often the problem might be ill-posed and regularization is used
 $R(T(x))$: regularization imposing some physical constraint

Registration techniques overview

Problem-based taxonomy

- Intra-subject
- Inter-subject
- Unimodal
- Multimodal

Transformation-based taxonomy

- Linear
 - Rigid
 - Similarity
 - Affine
- Non-linear / Deformable

- Landmark based registration
- Linear basis function modeling

- B-spline registration
- RBF registration
- ...

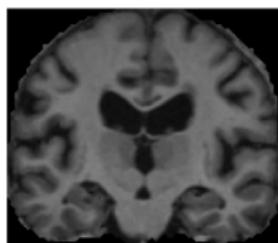
- Dense field models

- Demons, Diffeomorphic demons
- Large deformation diffeomorphic metric mapping (LDDMM)
- Poly-affine
- ...

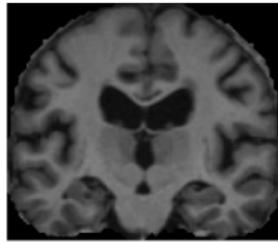
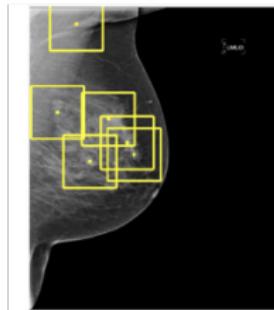
- Discrete optimization (DROP,...)

- Feature based (HAMMER,...)

Classification / Regression principle



How old is she?



Is she at risk for AD?



- Mapping from image to label
- Classification:
 $F : \Omega_c \subseteq \Omega \rightarrow \{0, \dots, N\}$
- Regression:
 $F : \Omega_c \subseteq \Omega \rightarrow \mathbb{R}^N$
- Labels can be
 - Diagnosis: Disease/Healthy
 - Outcome prediction:
Success/Complication
 - Presence of an object
- Ω_c can be the entire image, i.e. $F(I)$, for diagnosis, outcome prediction
- Ω_c can be part of an image, $F(I_c)$, for localization
- Segmentation / registration can be thought as classification / regression tasks with special structure, i.e. per pixel/voxel prediction

Classification techniques overview

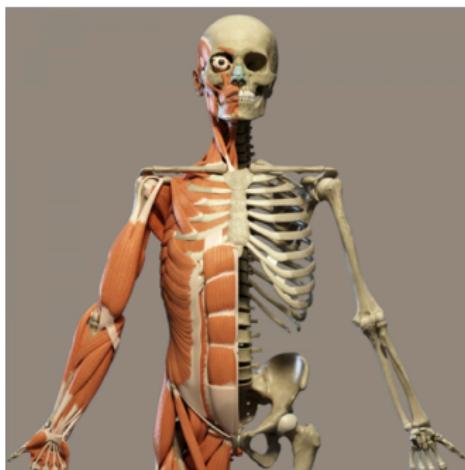
- Feature extraction
 - morphological descriptors
 - texture features
 - SIFT, HOG, ...
 - multimodal
- Supervised classification
 - Linear regression: both prediction and statistical analysis
 - Support vector machines, Kernel machines, ...
 - Random forests, Adaboost, ...
 - Neural networks

Outline

- What and why
- Main tasks in medical image analysis
- Image formats
 - Anatomical axes and basic properties
 - DICOM and volumetric formats
- Toolkits

Anatomical axes

Superior



Right

Superior



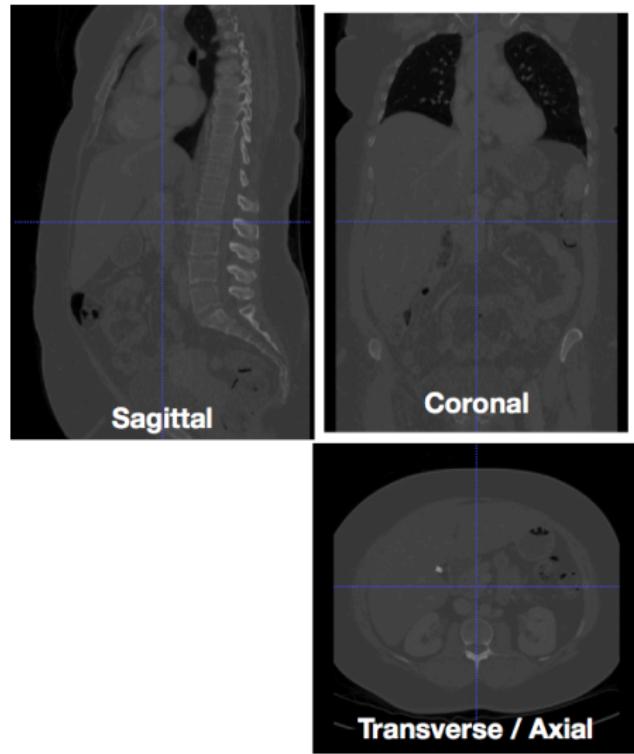
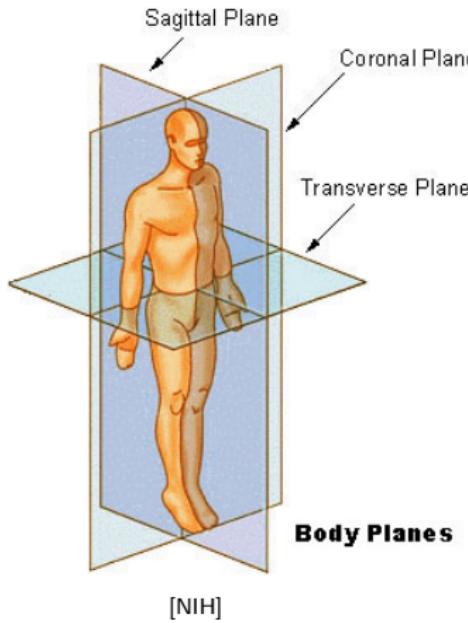
Posterior

Anterior

Inferior

Inferior

Cross sections in a volumetric image

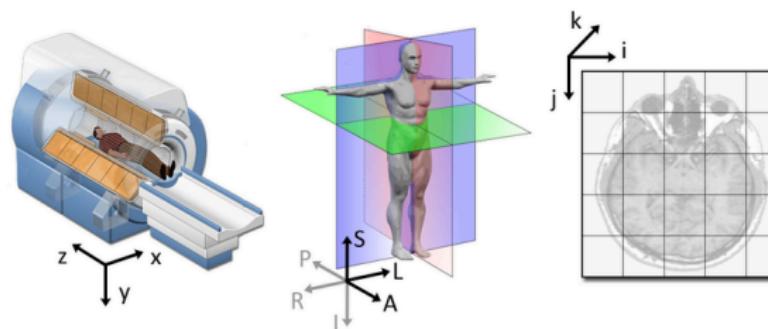


Basic image properties - volume

- Image size
 - number of pixels / voxels
 - 2 or 3 integers
 - e.g. 256x256x128
- Pixel / Voxel size
 - the size a pixel / voxel occupies in the real world
 - 2 or 3 real numbers
 - e.g. $0.6 \times 0.6 \times 3.5 \text{ mm}^3$
 - Sometimes called resolution, however, pixel size and resolution are two different things for acquisition people
- Field-of-view (FOV)
 - the size an entire image occupies in the real world
 - 2 or 3 real numbers
 - product of image size and pixel/voxel size
 - e.g. $153.6 \times 153.6 \times 448 \text{ mm}^3$

Basic image properties - beyond

- Temporal information
 - Additional dimension in the image, e.g. a volumetric image would have 4 dimensions with time
 - Additional number indicating number of images
 - Temporal resolution
 - functional MRI, perfusion, dynamic cardiac imaging, ...
- Metabolic information
 - Additional dimension
 - Additional number indicating number of metabolites
 - MR spectroscopy
- Transformations
 - Transformation matrices between voxel coordinates and real world coordinates - patient position on the table



[https://www.slicer.org/wiki/File:Coordinate_systems.png]

DICOM

- Need to keep the image and all the basic information
- Need to keep additional information
 - Scanner
 - Scan type
 - Ordering physician, conducting physician
 - Patient information
 - ...
- Digital Imaging and Communications in Medicine (DICOM)
- Standard for storing and transmitting data
- Most commonly used format in practice
- All vendors, all centers, all doctors
- Used in clinical practice
- Turns out to be difficult to manage for processing

Name
DICOMDIR
README.TXT
DICOM
0000BE1F
AA8751BC
AA549D59
000034F4
EE0AD5EB
EE0AE431
EE0B143B
EE0BD21F
EE0D97E7
EE0DC541
EE0DE6D7
EE0E4C51
EE0EEBD2
EE1A213A
EE1C7BDA
EE1CC591
EE1D19FC
EE1EBF9E
EE2C8BBB
EE2CA5CB
EE2D6D72
EE2D2720
EE2DE720
EE2E514D
EE3AB5EE
EE3B64D4
EE4C88B1

Volumetric formats

- Luckily people came up with many volumetric formats
- Sometimes institutional proprietary formats
- Analyze file format from Mayo Clinic
[https://en.wikipedia.org/wiki/Analyze_\(imaging_software\)](https://en.wikipedia.org/wiki/Analyze_(imaging_software))
- NIFTI (Neuroimaging Informatics Technology Initiative)
replacement for Analyze
<https://nifti.nimh.nih.gov/nifti-1/>
- NRRD (Nearly raw raster data)
<https://en.wikipedia.org/wiki/Nrrd>
- Which format to use depends on your preferences
- Useful for engineering purposes but need to be able to deal with DICOM if you want to work with clinicians

Outline

- What and why
- Main tasks in medical image analysis
- Image formats
- Toolkits
 - Visualization and simple manipulations
 - Processing

Visualization and simple manipulations

■ ITK-snap

<http://www.itksnap.org/pmwiki/pmwiki.php>

Great tool for basic visualization of volumetric data

Simple routines for image segmentation

■ OSIRIX

<http://www.osirix-viewer.com/>

Popular, especially among clinicians

■ Freeview

<https://surfer.nmr.mgh.harvard.edu/>

Visualization tool from freesurfer

■ Medinria

<http://med.inria.fr/>

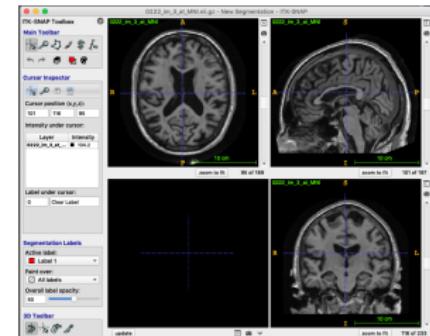
■ Mango

<http://ric.uthscsa.edu/mango/>

■ Paraview

<https://www.paraview.org/>

Scientific visualization, great for meshes



ITK-Snap

Processing

■ Libraries

- ITK (Insight Toolkit), <https://itk.org/>
- VTK (Visualization Toolkit), <https://www.vtk.org/>
- Tensorflow
- PyTorch
- Scikit-learn
- Nibabel - to read medical volumes in python
- Iso2Mesh - to generate mesh from volumetric data
- GitHub resources...

■ Software suites - combination of executables

- Freesurfer <https://surfer.nmr.mgh.harvard.edu/>
- FSL <https://fsl.fmrib.ox.ac.uk/fsl/fslwiki/>
- Camino <http://camino.cs.ucl.ac.uk/>
- SPM <https://www.fil.ion.ucl.ac.uk/spm/>

■ Visualization and processing with GUI

- Slicer3D <https://www.slicer.org/>

■ Many more...