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Mach F²
Vallée J-P¹

High time-resolved cardiac functional imaging with temporal regularization in mouse on a clinical 3T scanner

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BASP Workshop - Villars 09.08.2011

Cardiac imaging on rodents on clinical scanners

- Importance of research on rodents
- Dedicated scanners not always available
- Importance of translational research for drug or medicine studies
- Problem of hardware limitations (gradient amplitude and slew-rate)



Cardiac imaging on rodents on clinical scanners

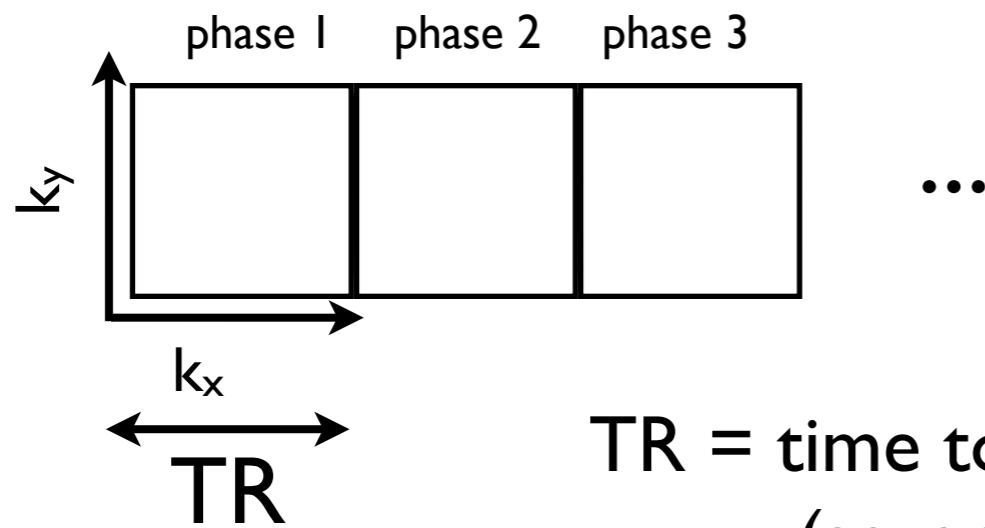
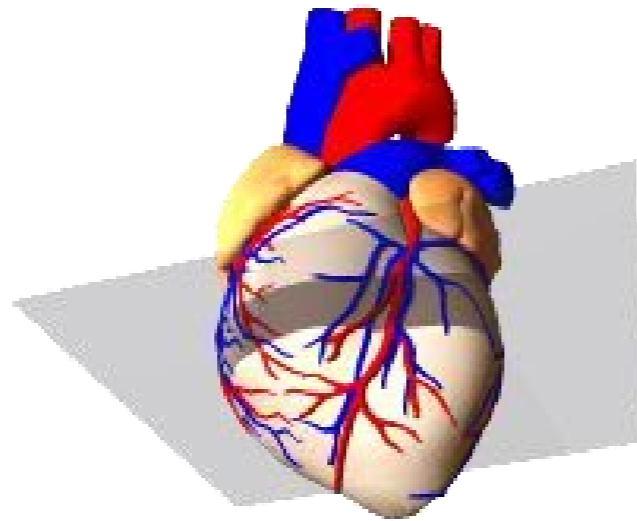
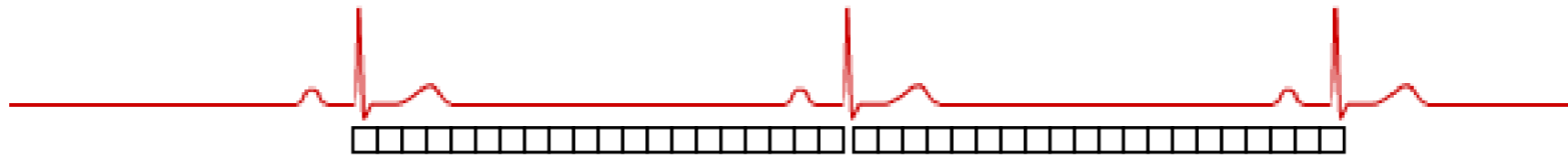
- Importance of research on rodents
- Dedicated scanners not always available
- Importance of translational research for drug or medicine studies
- Problem of hardware limitations (gradient amplitude and slew-rate)

New cine sequence

Compatible with resolutions needed in stress studies in mice (TR=8.6ms)¹

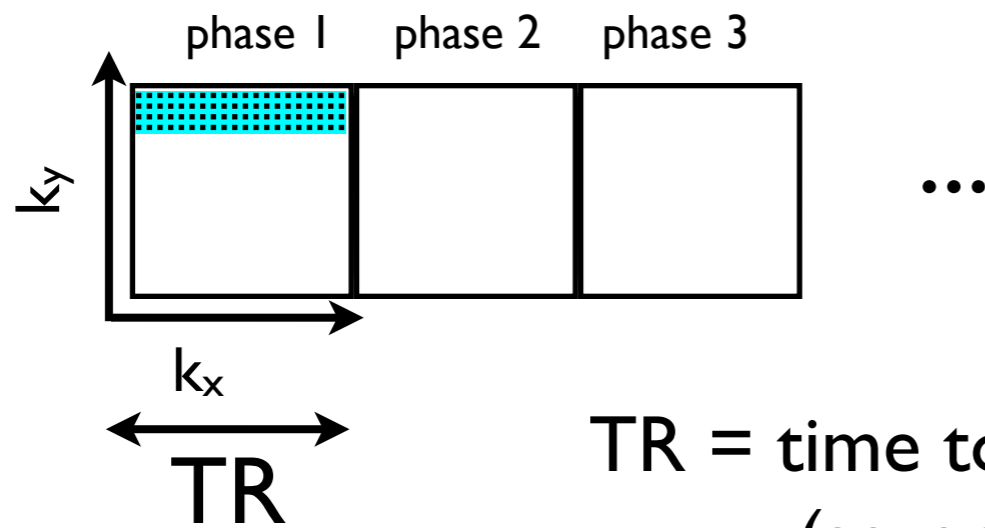
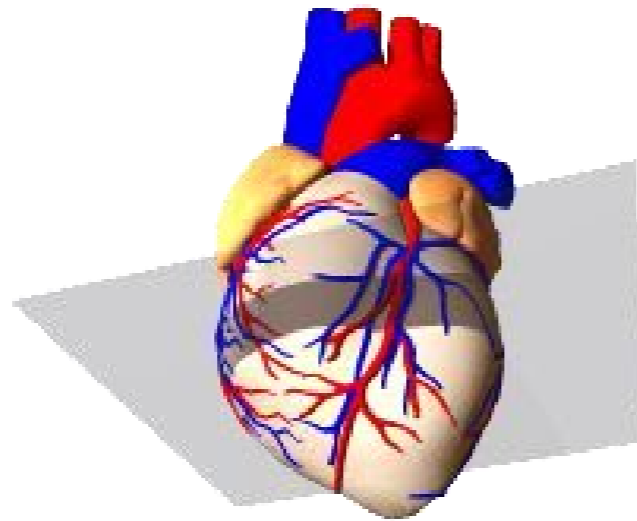
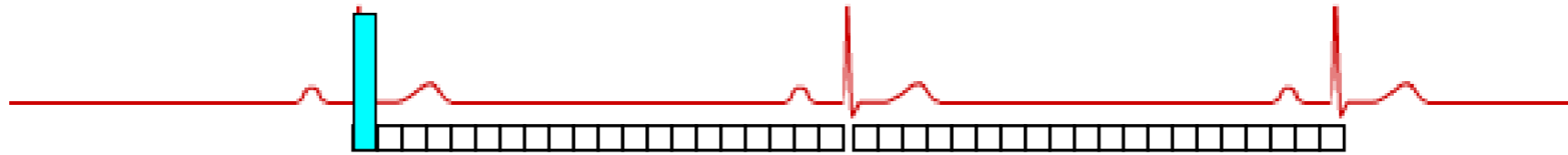


Cine acquisition



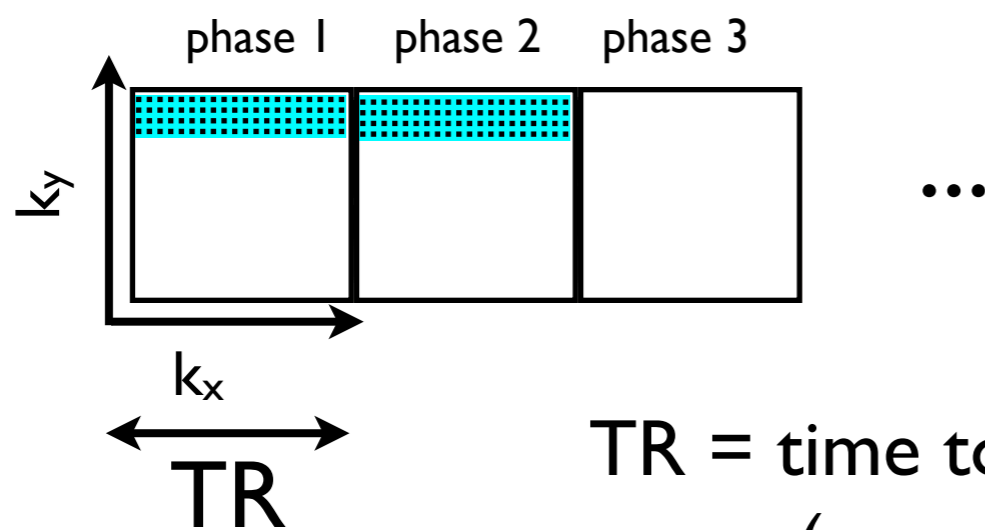
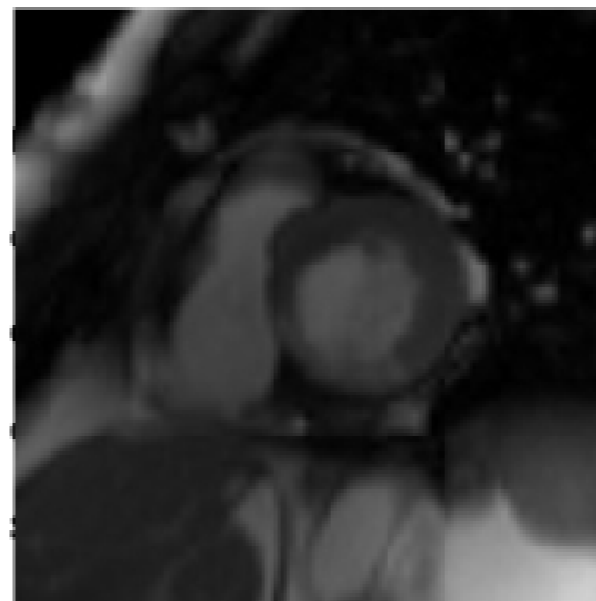
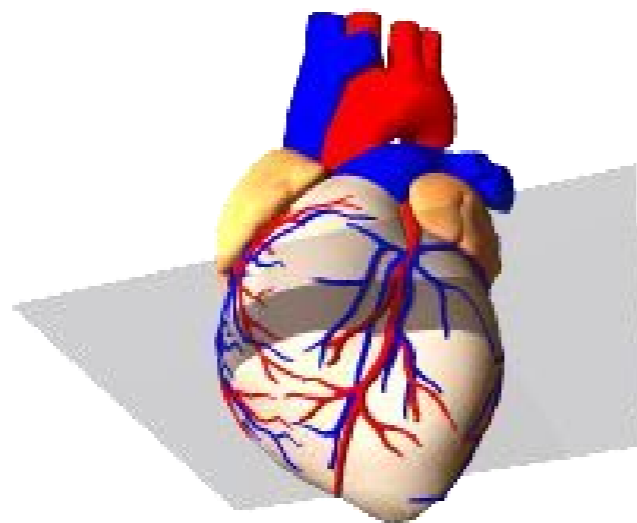
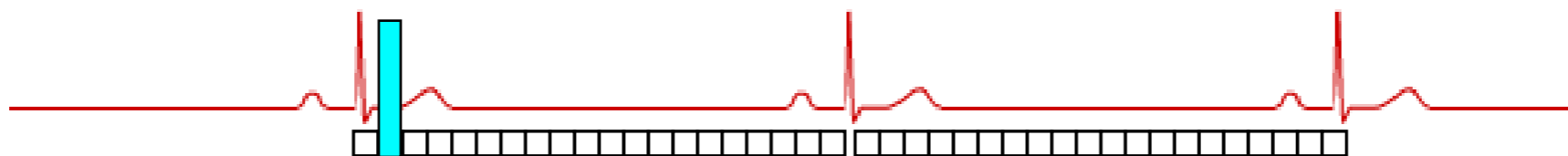
TR = time to acquire one segment
(several k -space lines)

Cine acquisition



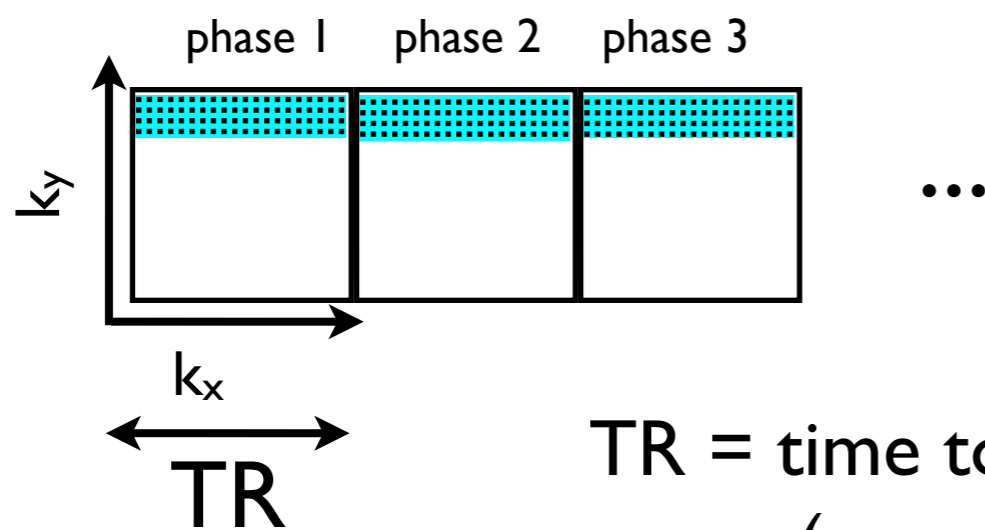
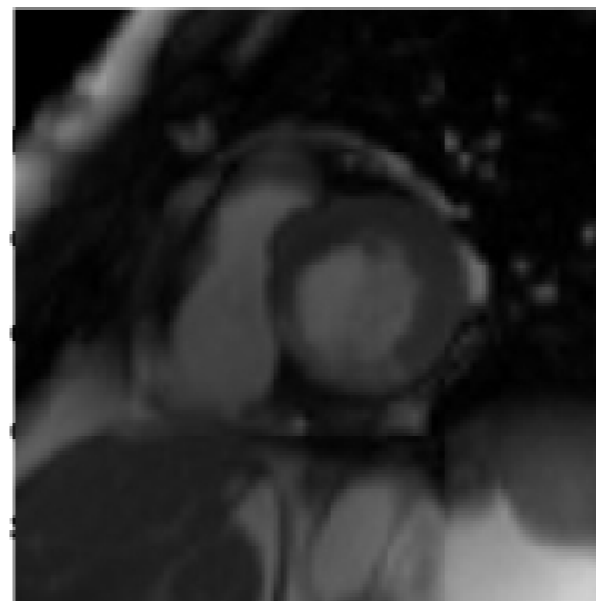
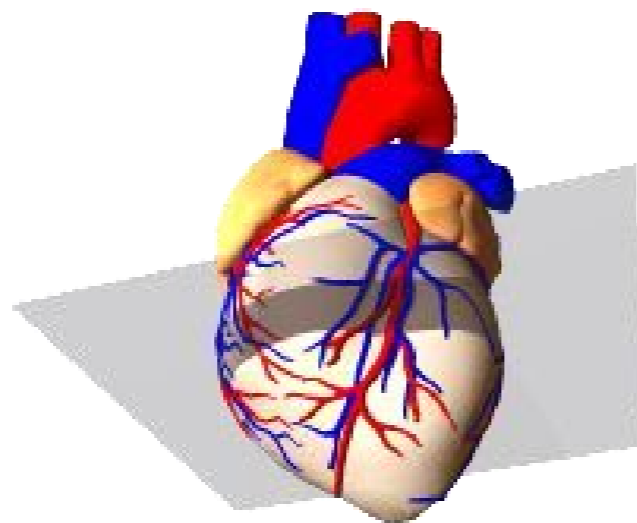
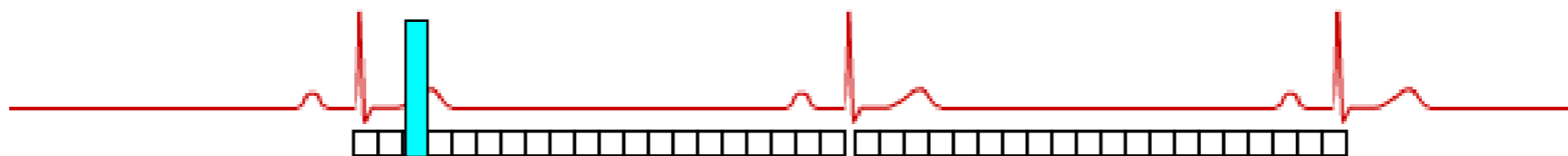
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Cine acquisition



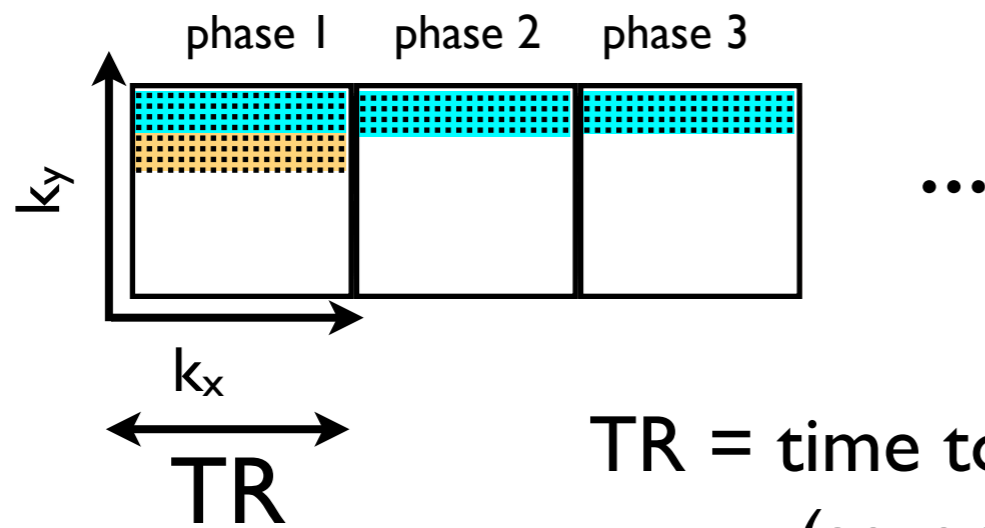
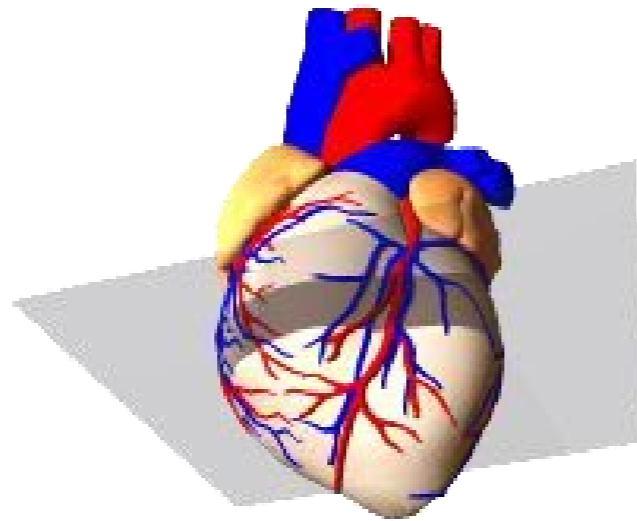
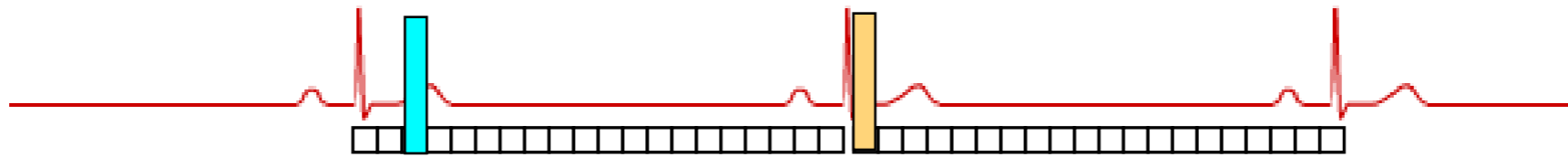
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Cine acquisition



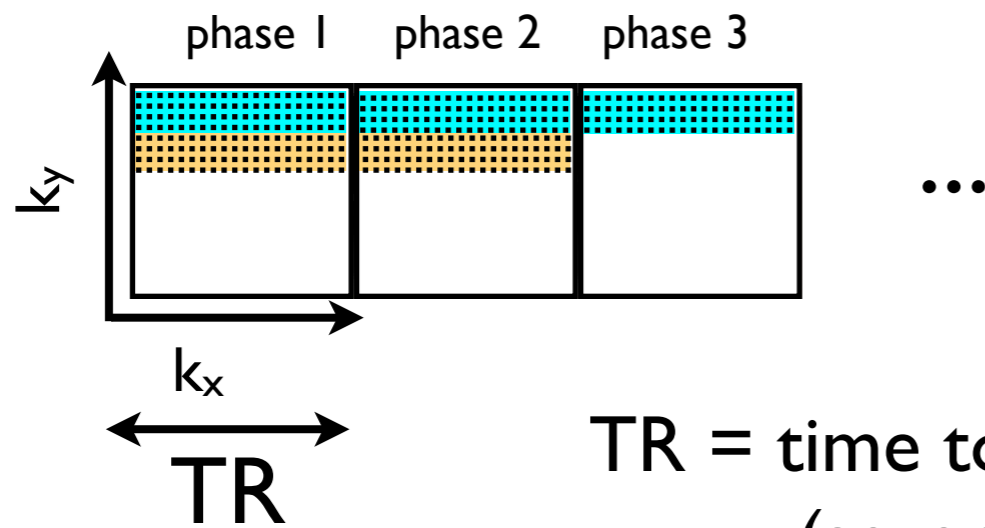
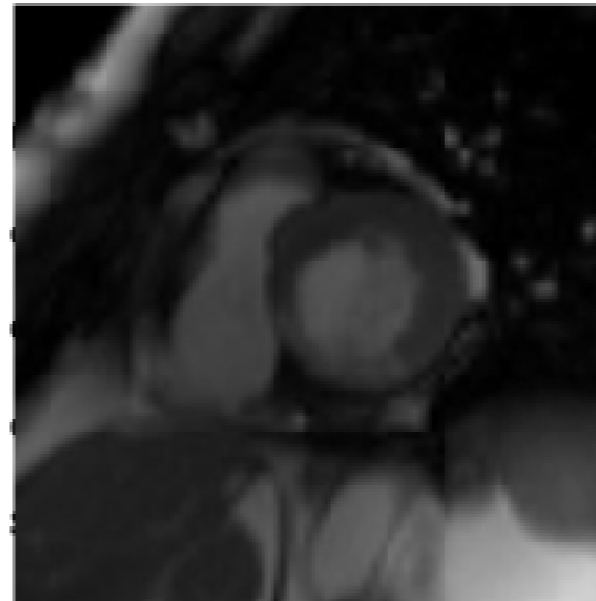
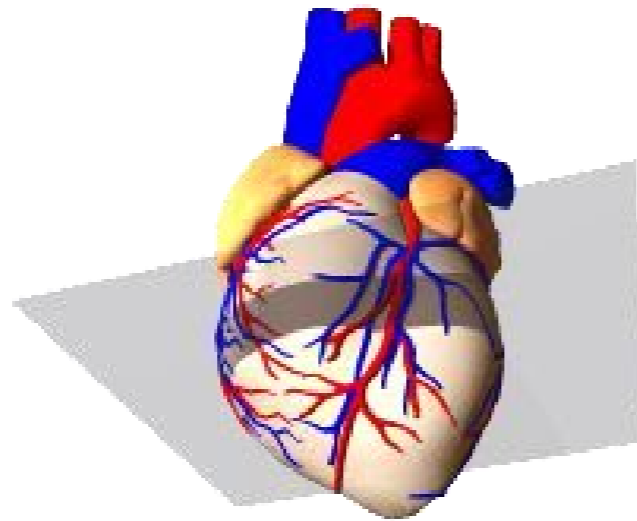
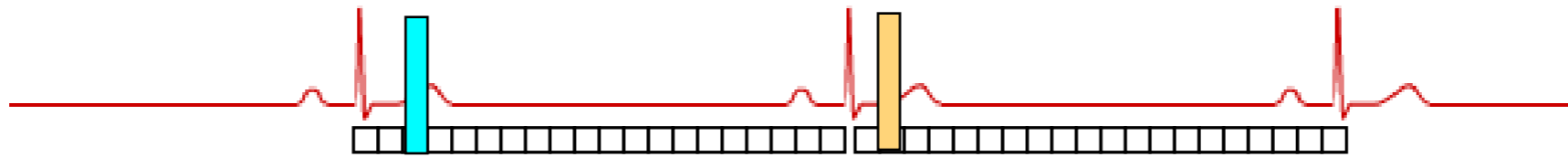
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Cine acquisition



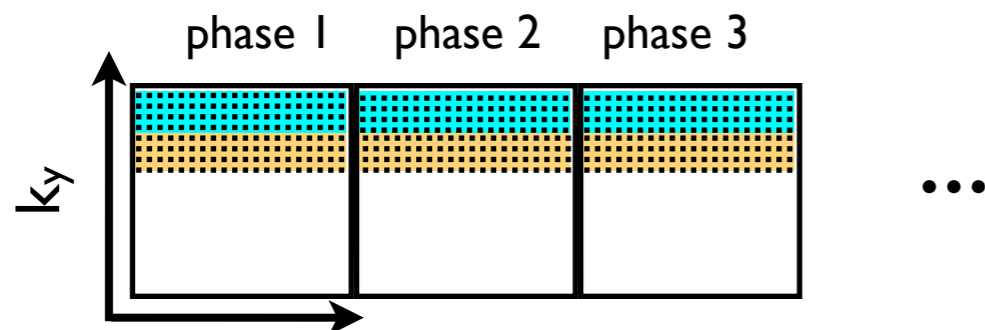
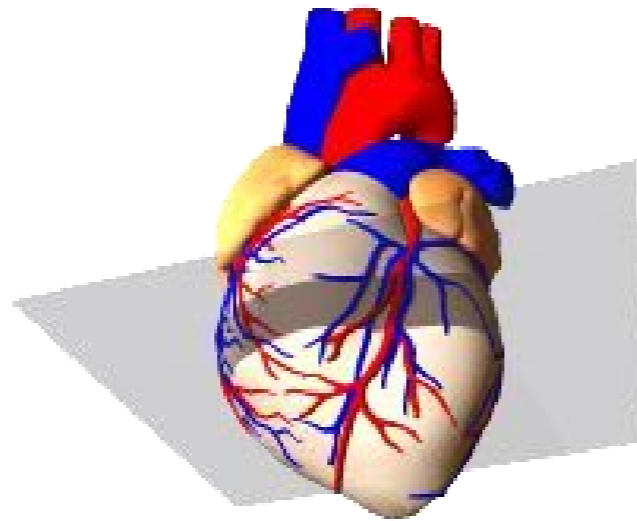
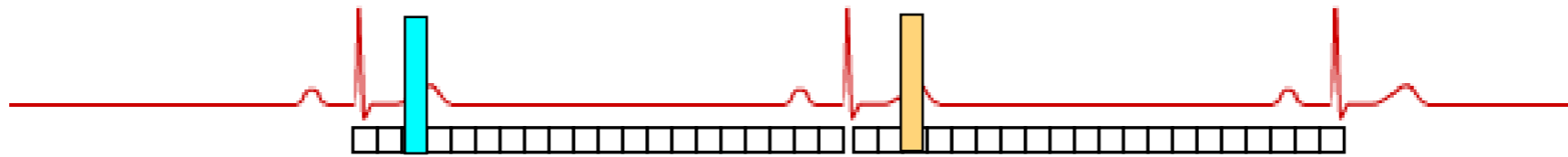
TR = time to acquire one segment
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Cine acquisition



TR = time to acquire one segment
(several k -space lines)

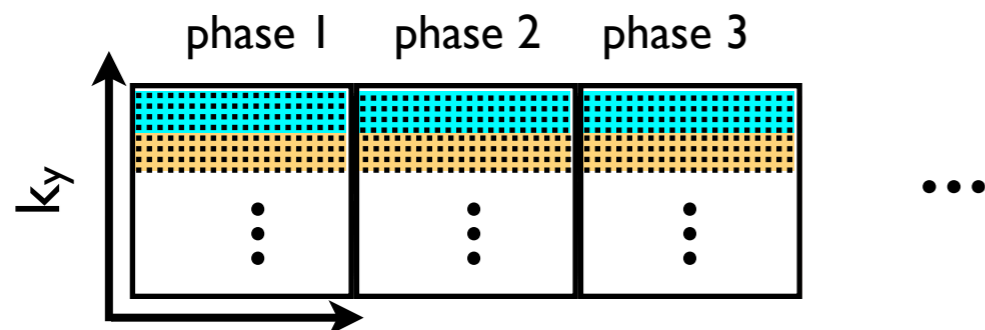
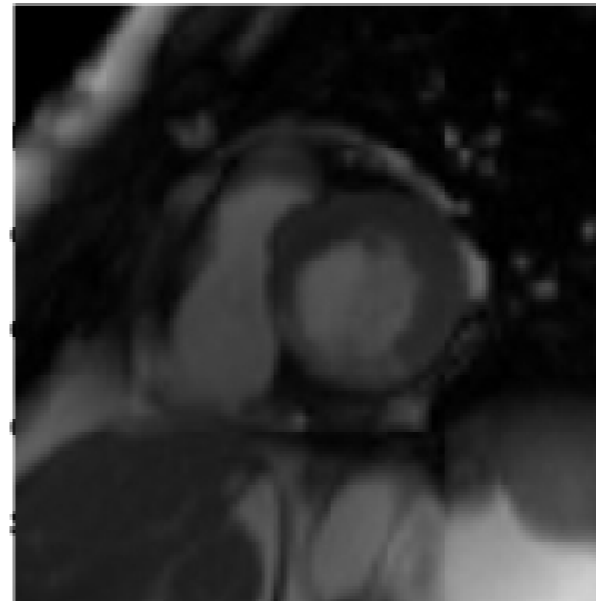
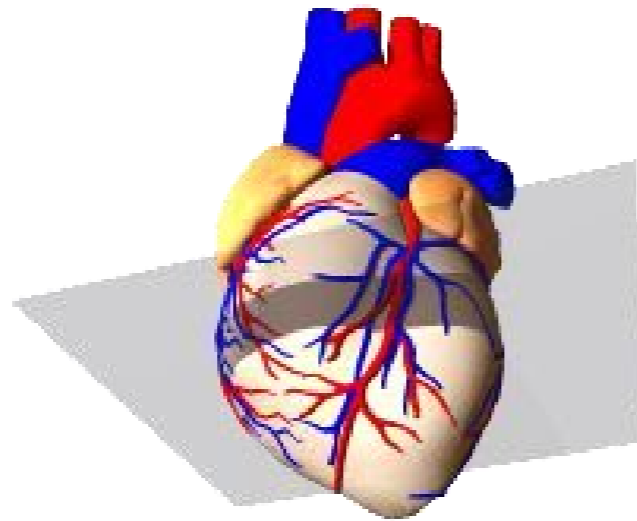
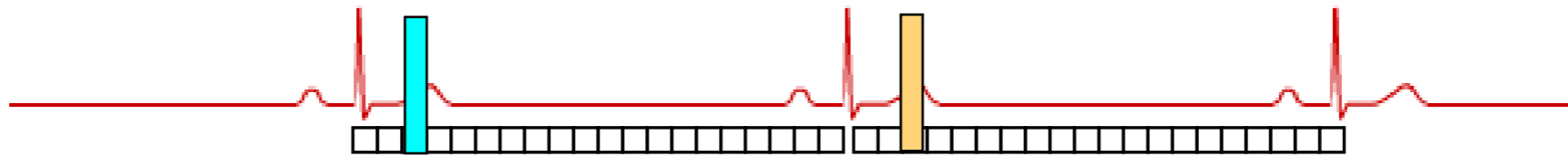
Cine acquisition



k_x
TR

TR = time to acquire one segment
(several k -space lines)

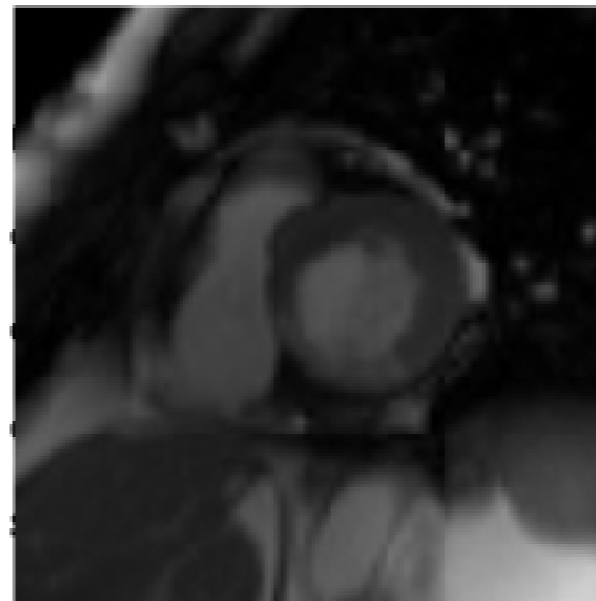
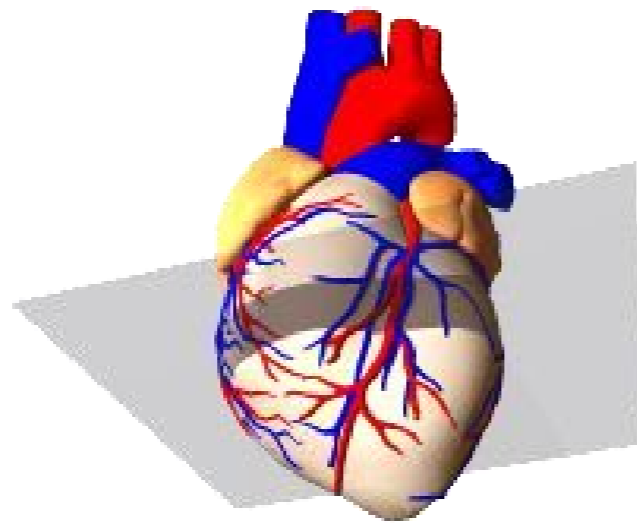
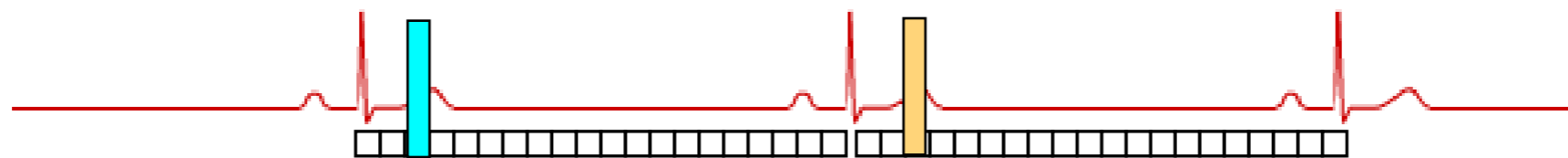
Cine acquisition



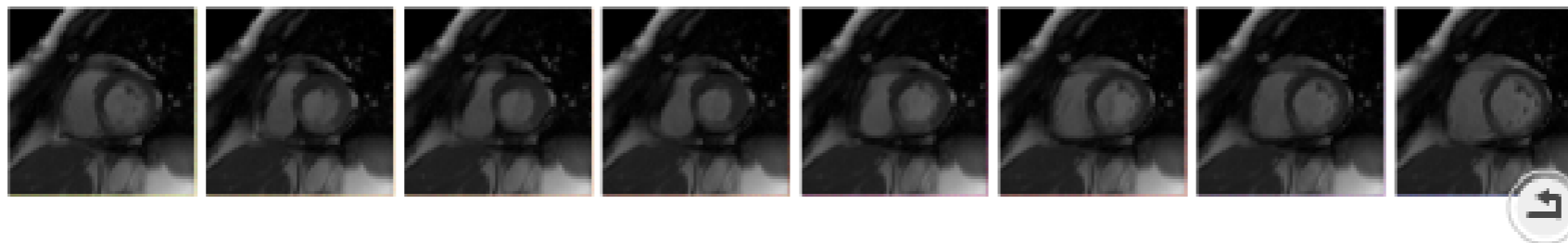
k_x
TR

TR = time to acquire one segment
(several k -space lines)

Cine acquisition



phase 1 phase 2 phase 3



← TR →

TR = time to acquire one segment
(several k -space lines)

Robust function evaluation: 11 phases per cycle (Roussakis et al., JCMR 2004)

Cine acquisition



Human
15 cm
60 bpm

Cine acquisition



Human
15 cm
60 bpm

11 phases

Cine acquisition



Human
15 cm
60 bpm



11 phases
**TR should be
around 9 ms**

Mouse
1 cm
600 bpm

Cine acquisition



Human
15 cm
60 bpm



11 phases
**TR should be
around 9 ms**

Mouse
1 cm
600 bpm

On dedicated scanner: in-plane resolution 117 μm^2 , TR **8.4 ms**¹

The best we can do: in-plane resolution 257 μm^2 , TR 13.5 ms

Cine acquisition



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15 cm
60 bpm



11 phases
**TR should be
around 9 ms**

Mouse
1 cm
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On dedicated scanner: in-plane resolution 117 μm^2 , TR **8.4 ms**¹

The best we can do: in-plane resolution 257 μm^2 , TR 13.5 ms

Best is not enough...

Cine acquisition



Human
15 cm
60 bpm



11 phases
**TR should be
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1 cm
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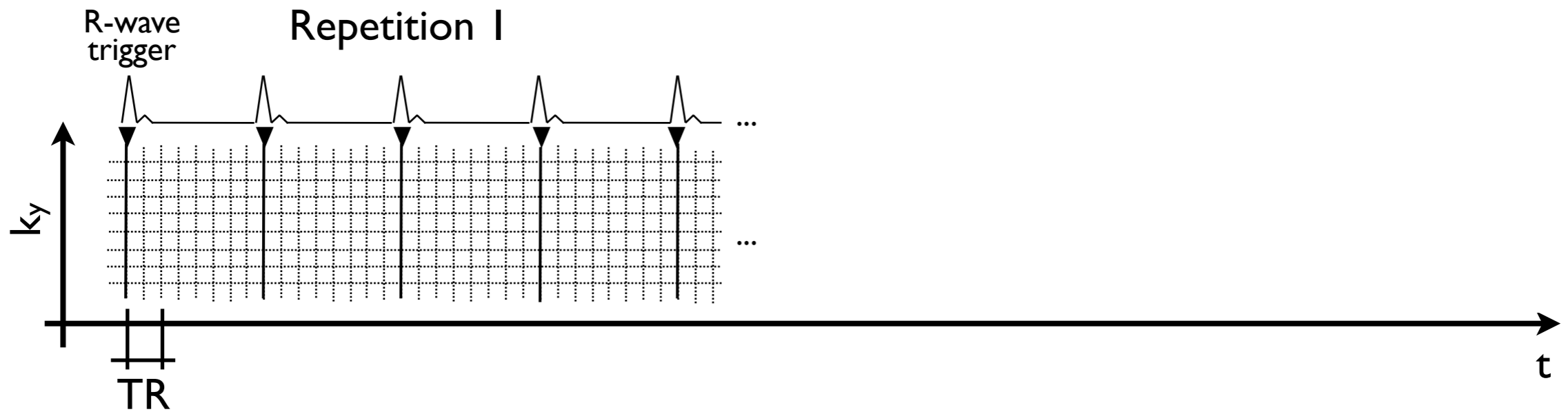
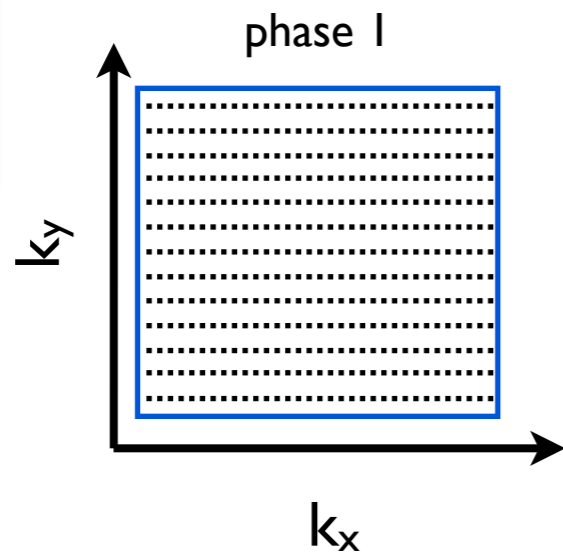
On dedicated scanner: in-plane resolution $117 \mu\text{m}^2$, TR **8.4 ms**¹

The best we can do: in-plane resolution $257 \mu\text{m}^2$, TR 13.5 ms

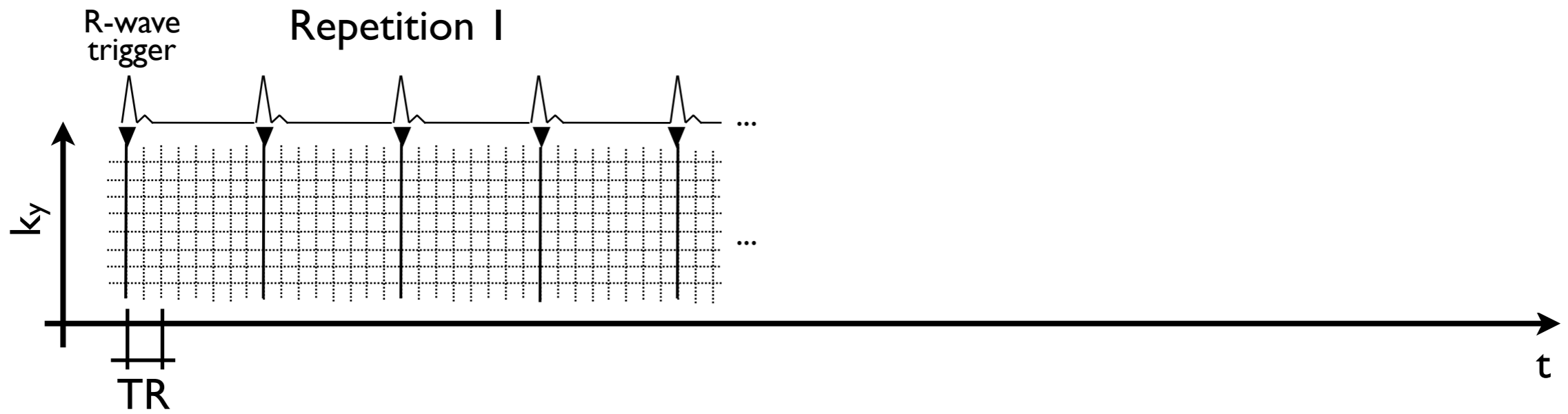
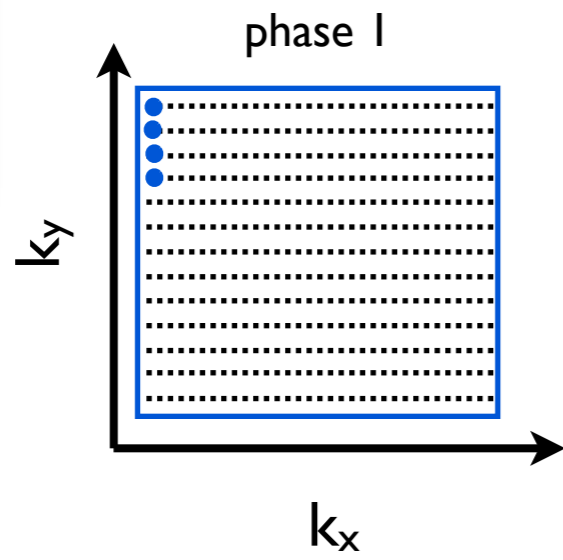
Best is not enough...

- ▶ 10 times higher gradient strength
- ▶ Modify sequence scheme

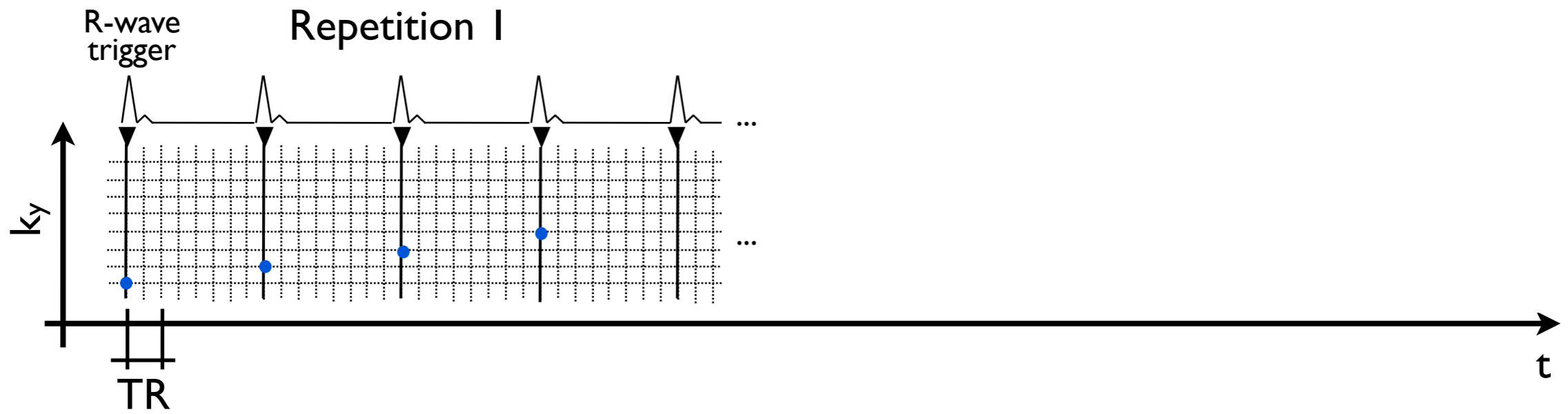
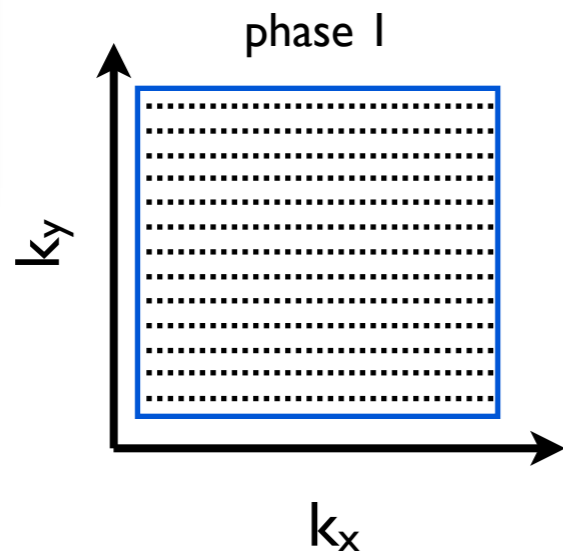
Basic cine



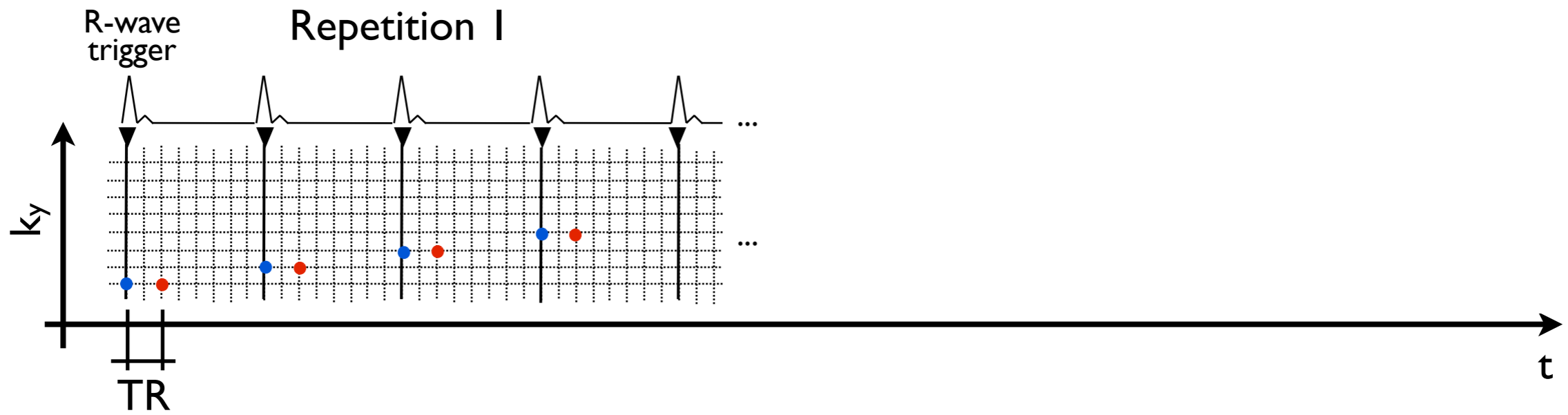
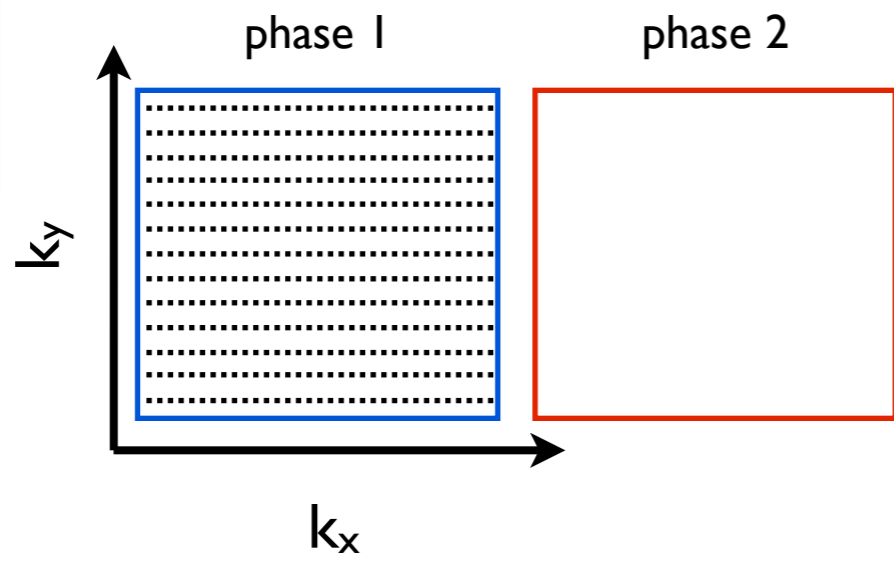
Basic cine



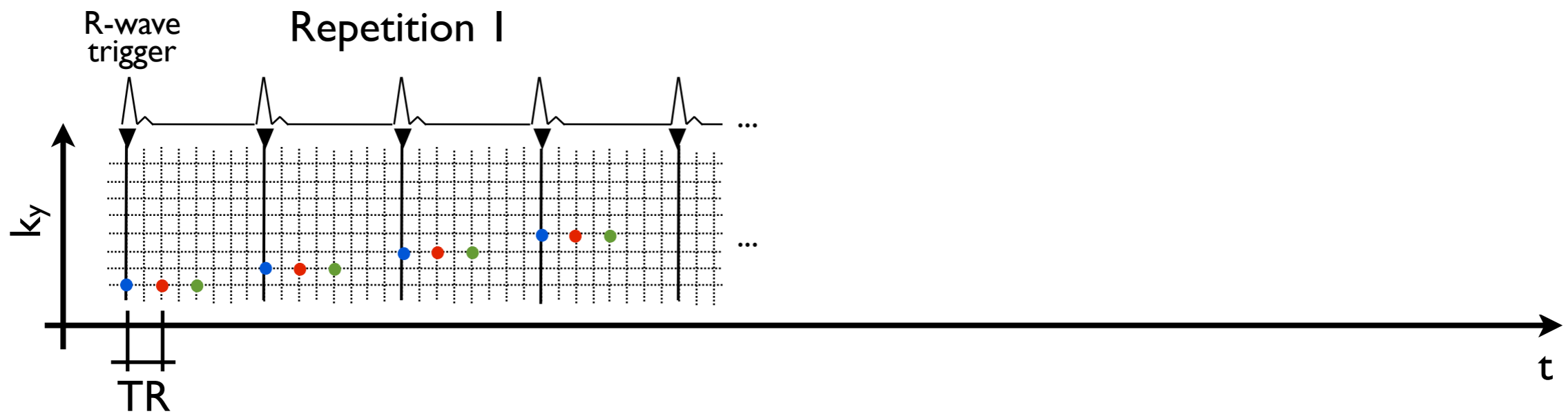
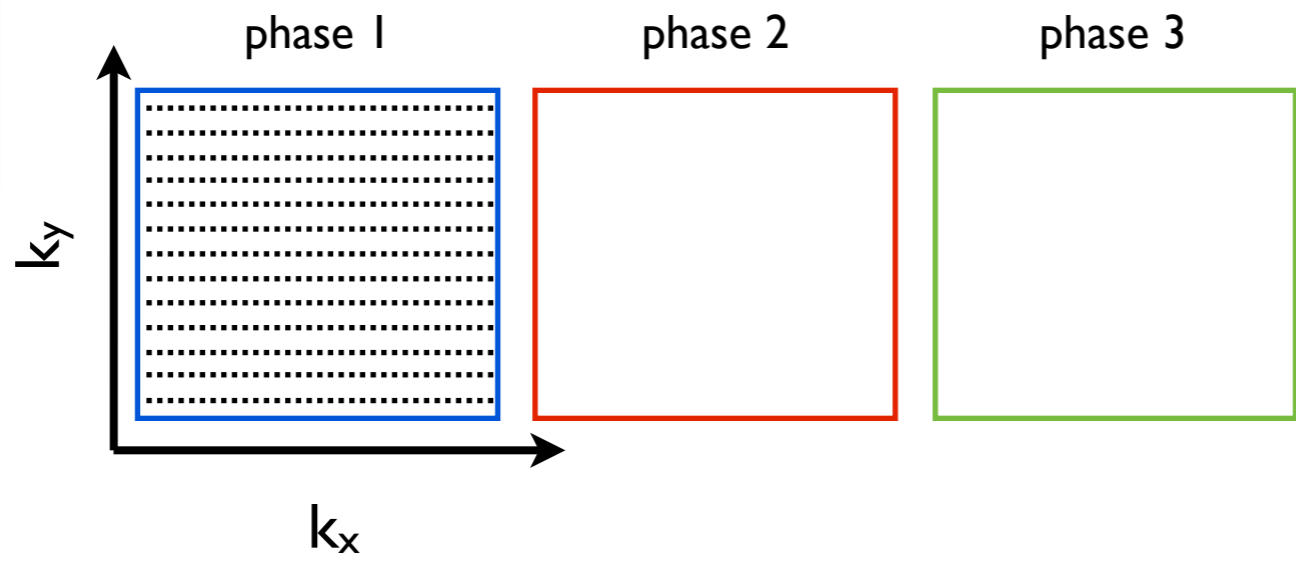
Basic cine



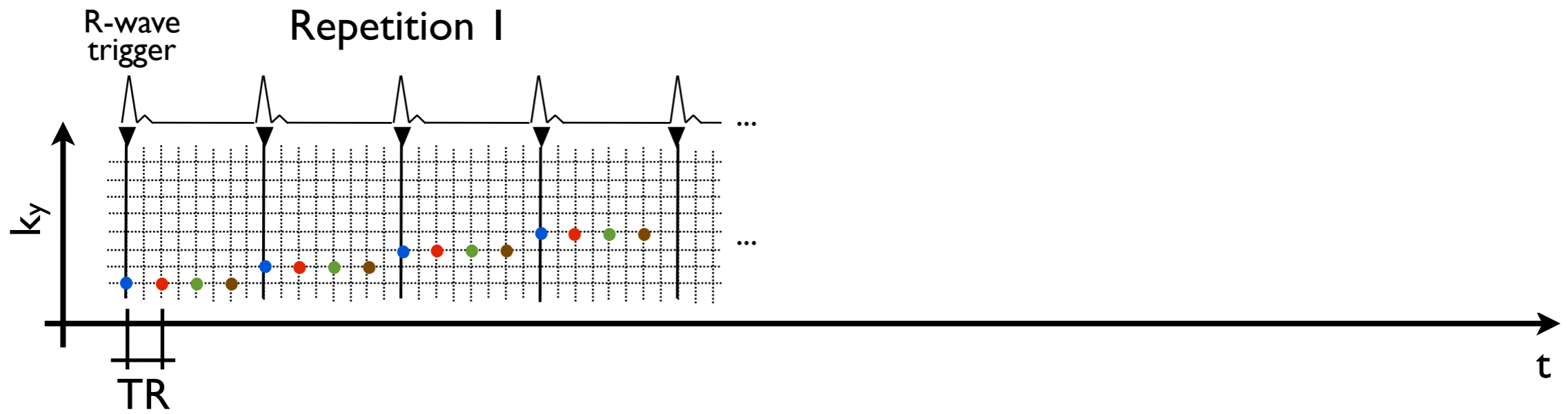
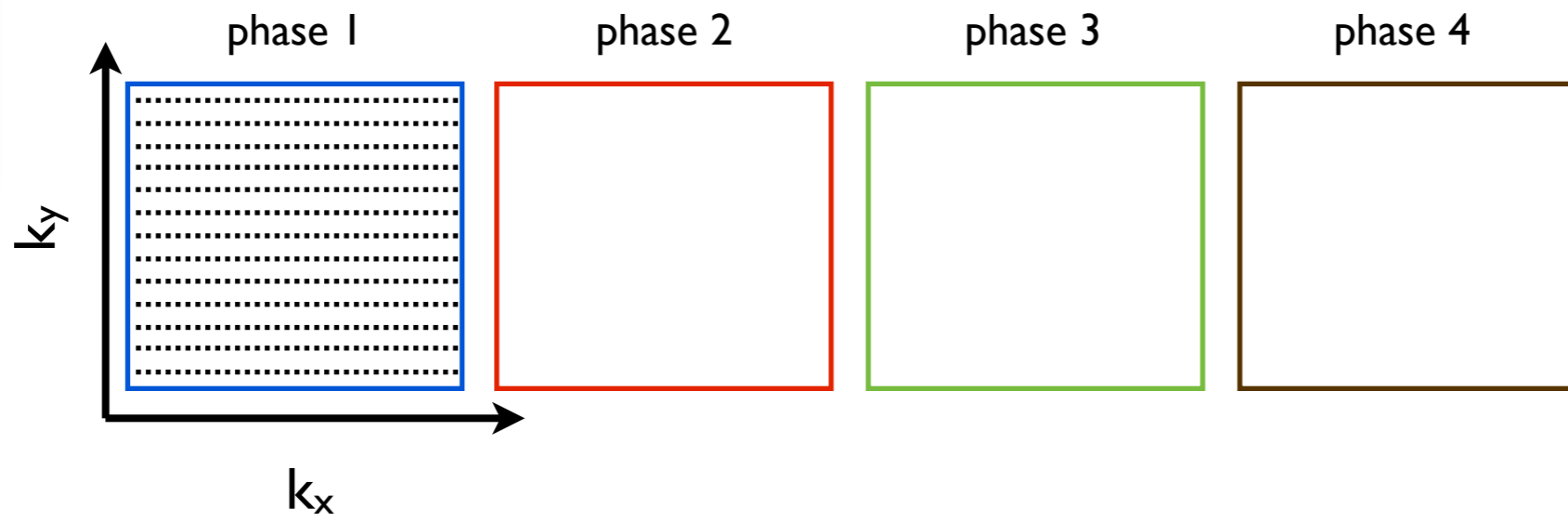
Basic cine



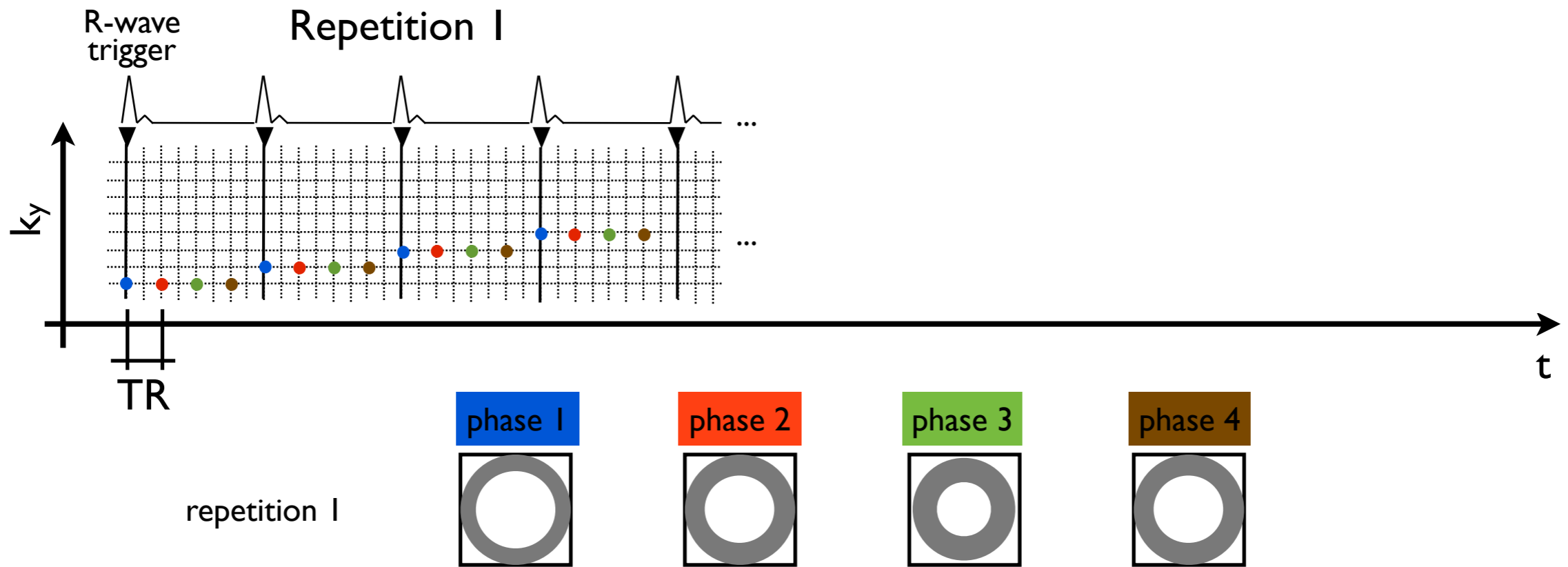
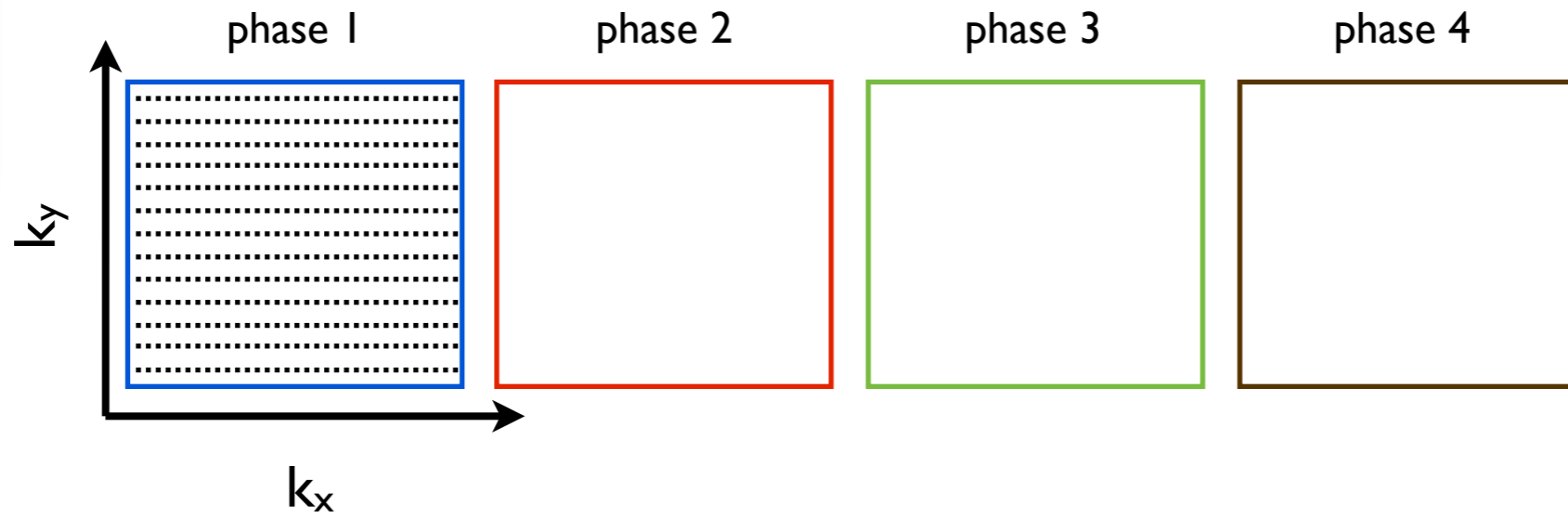
Basic cine



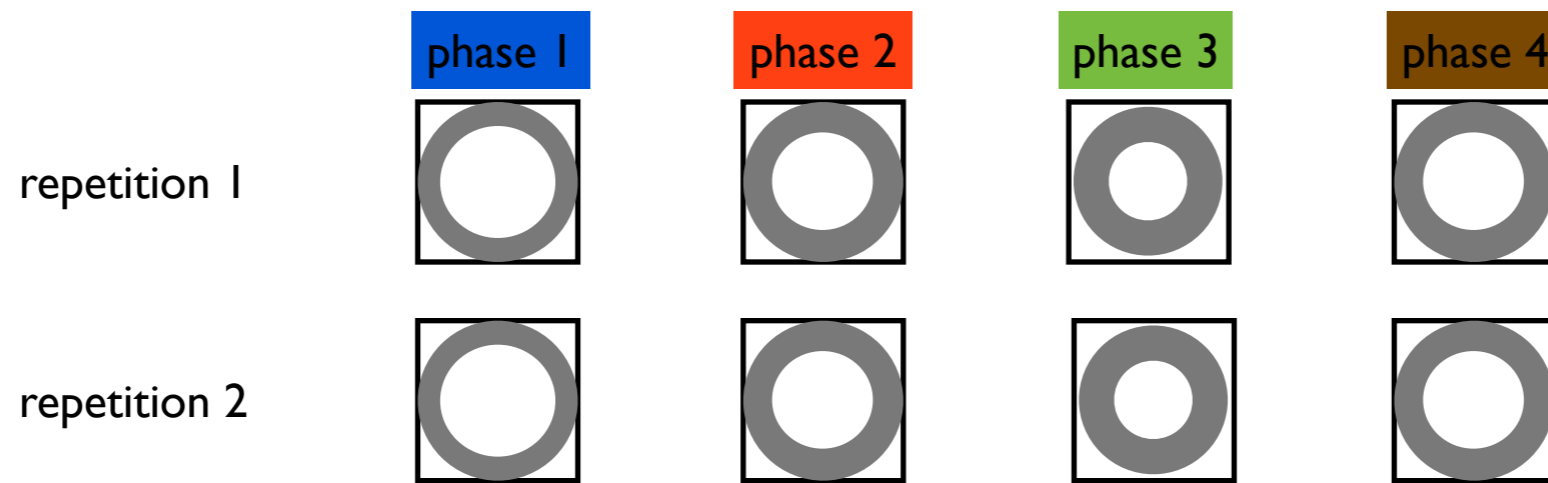
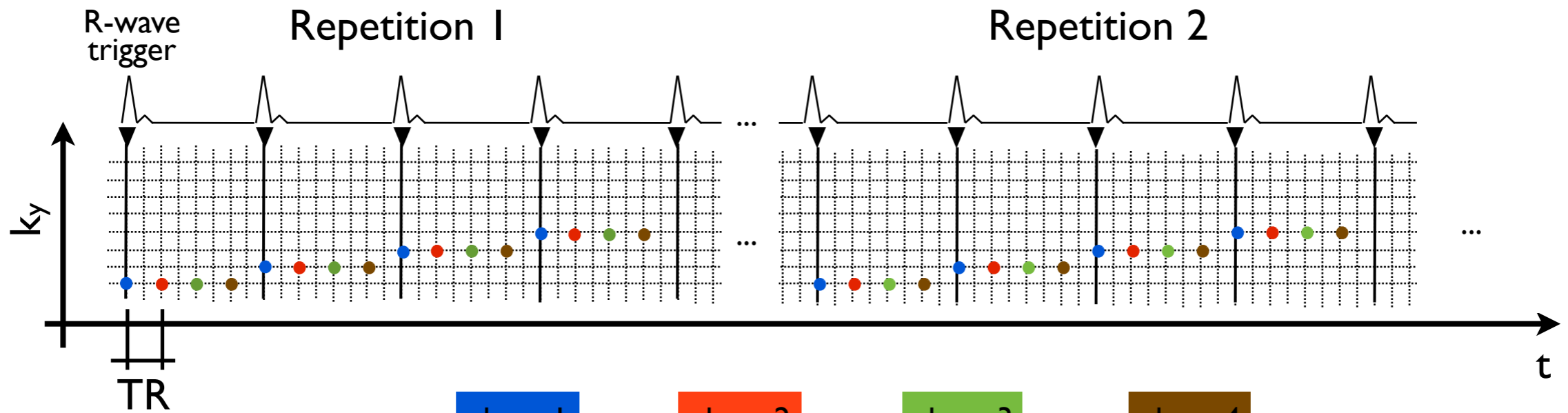
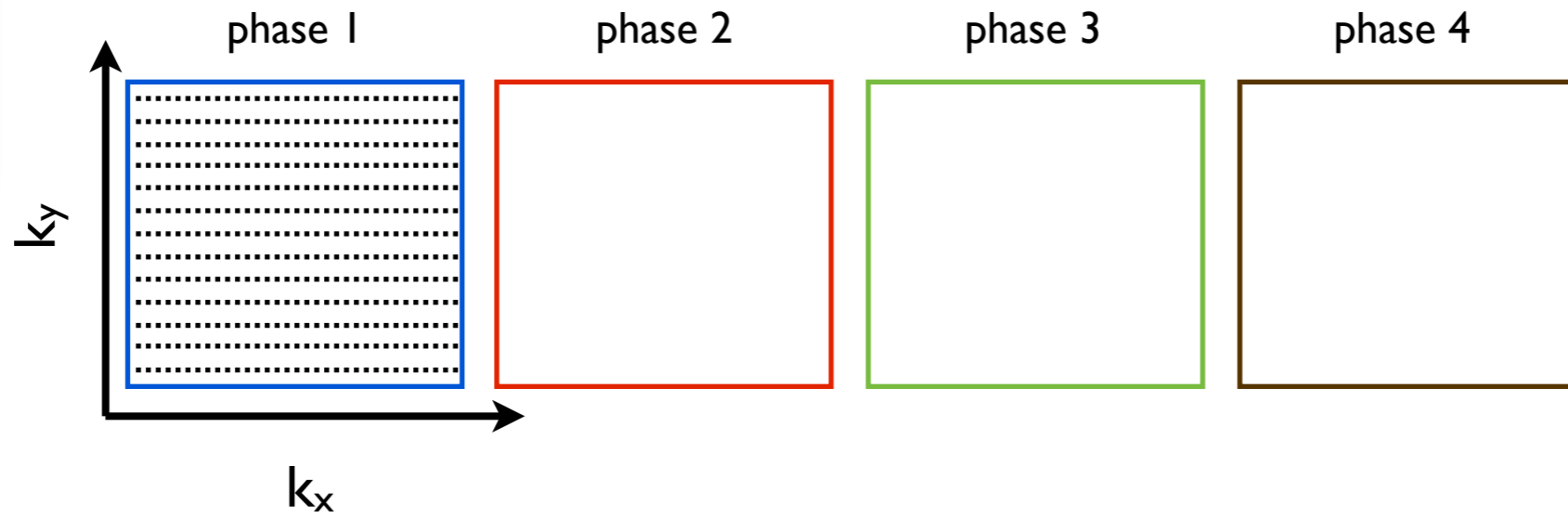
Basic cine



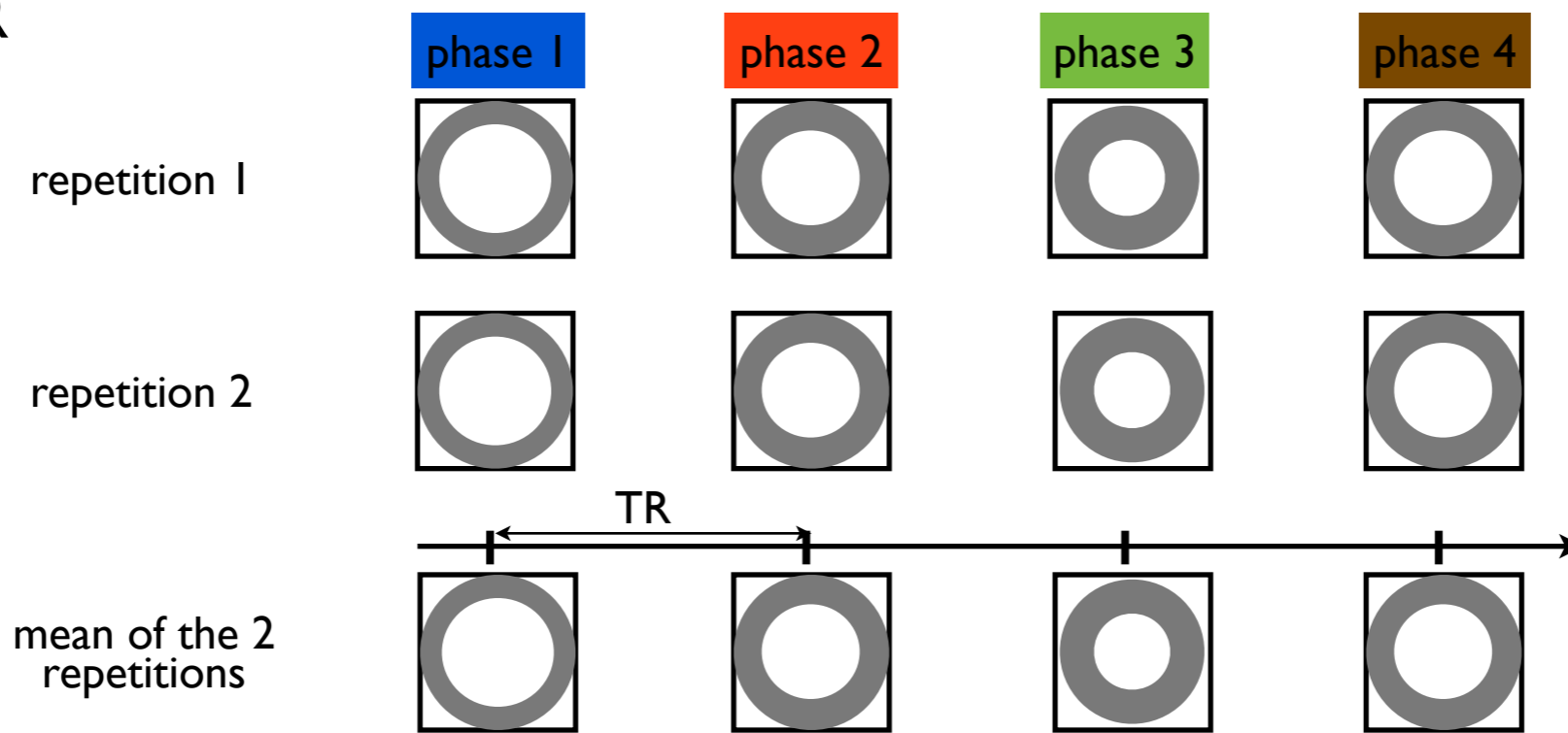
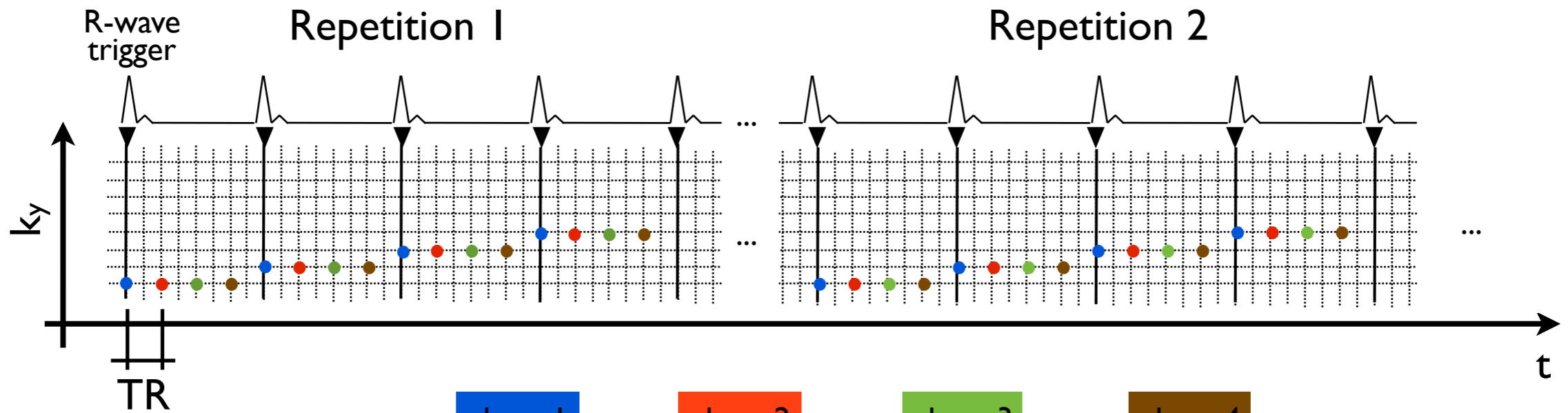
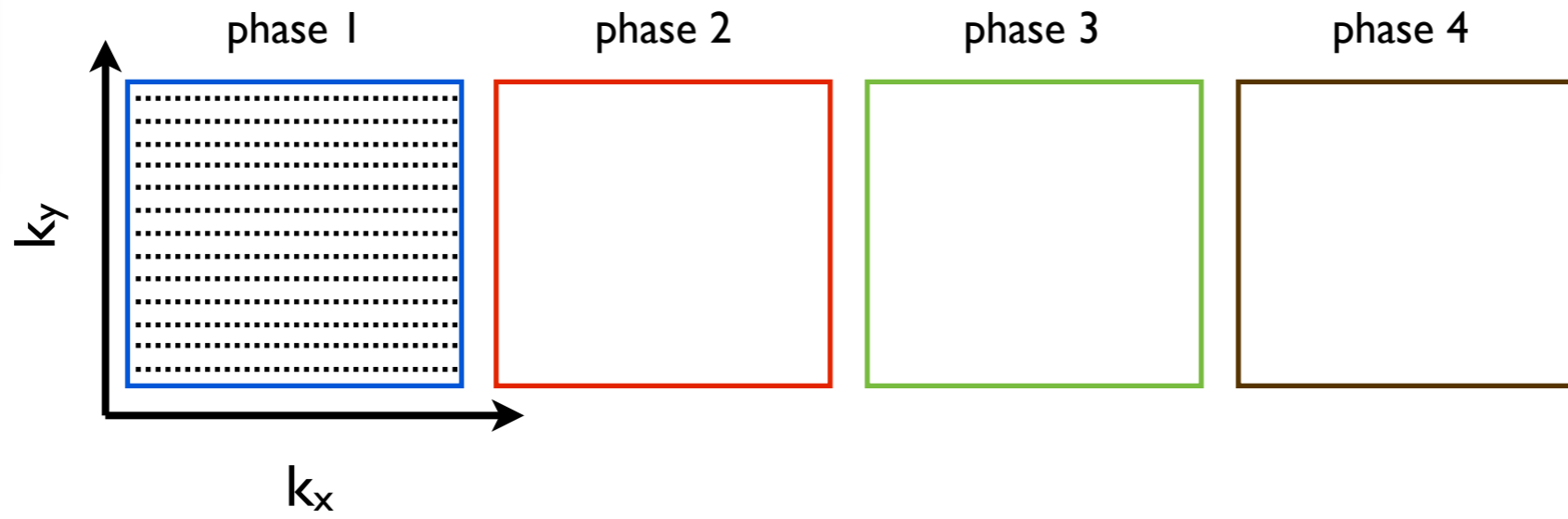
Basic cine



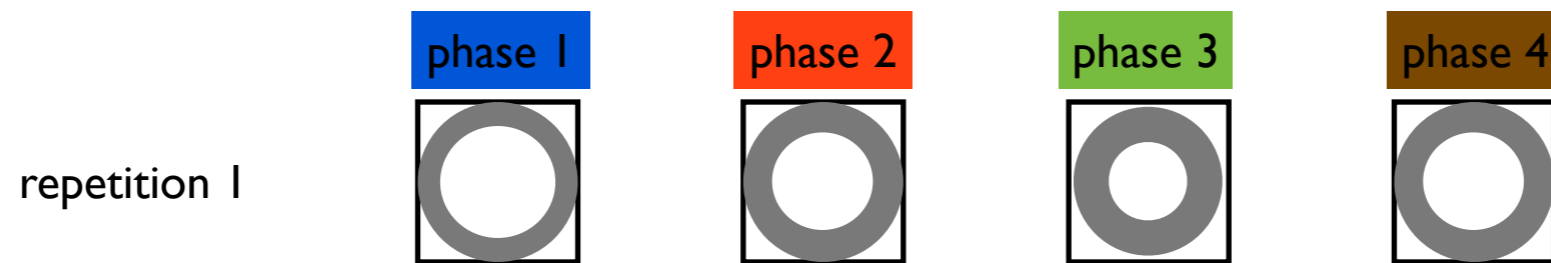
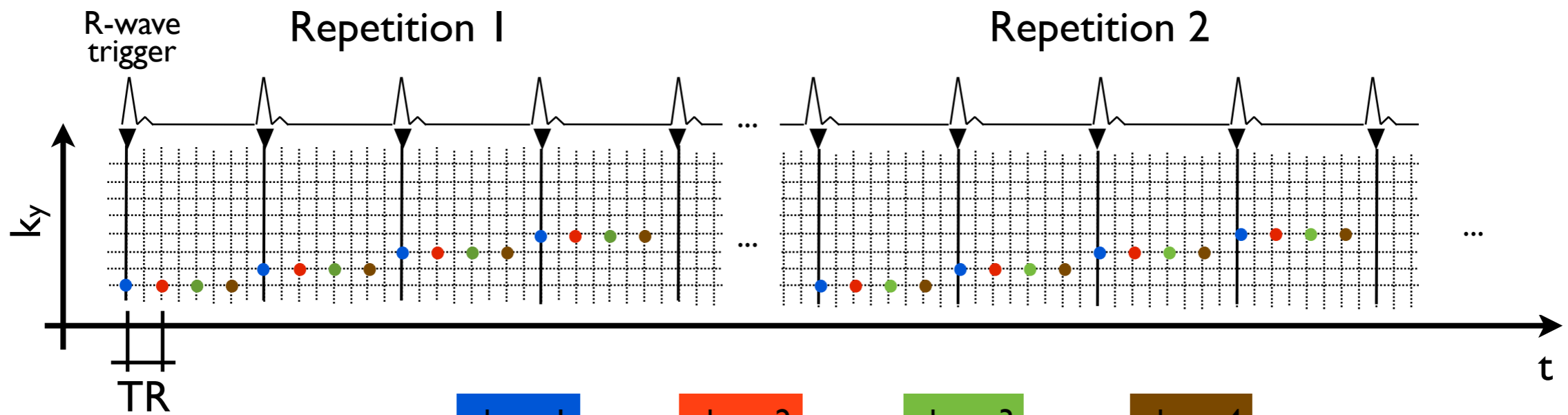
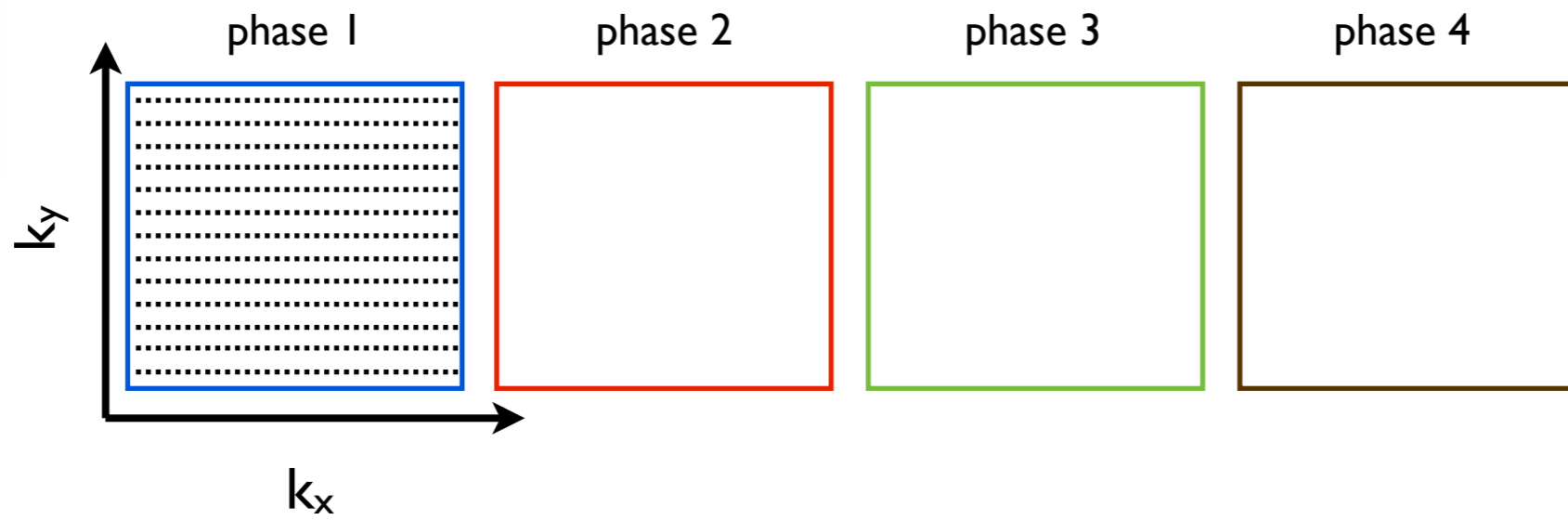
Basic cine



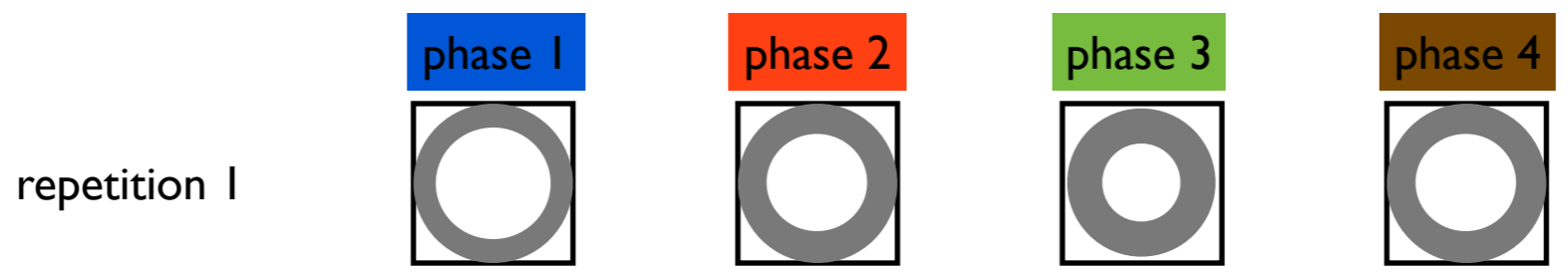
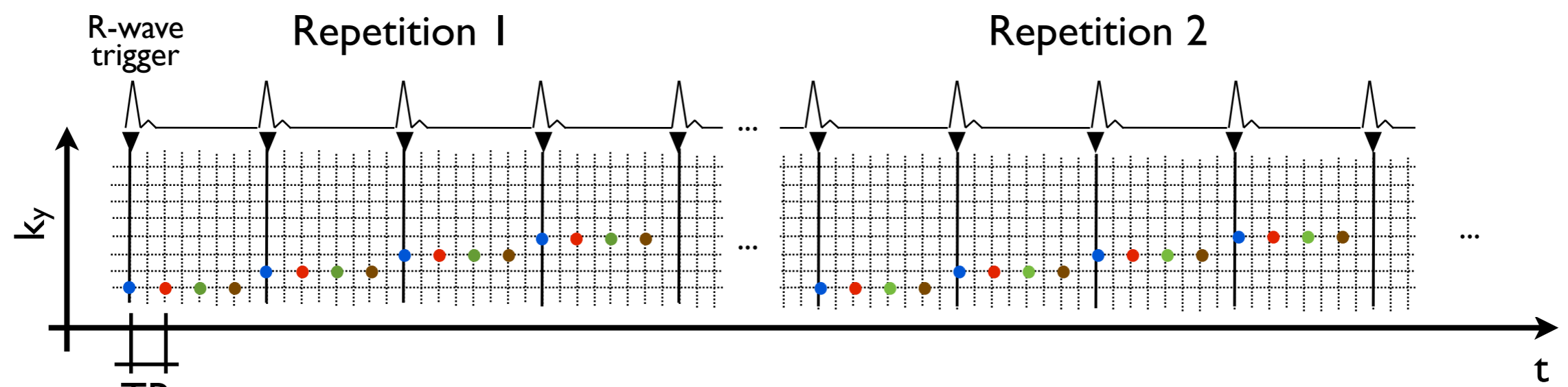
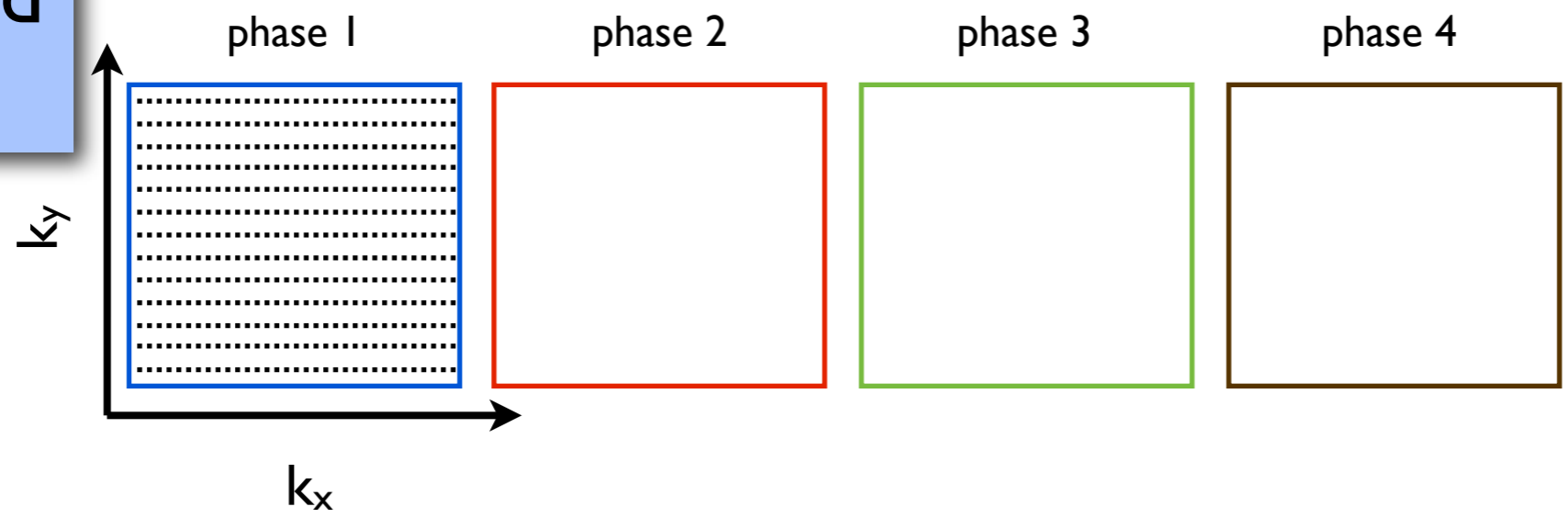
Basic cine



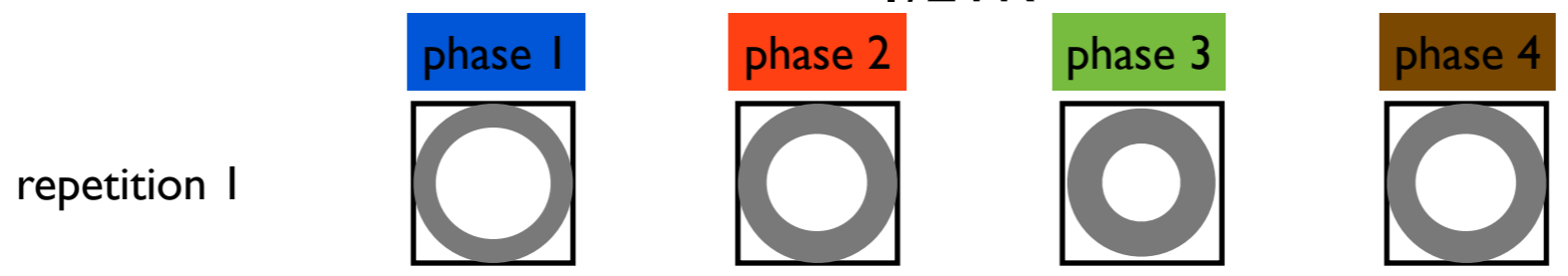
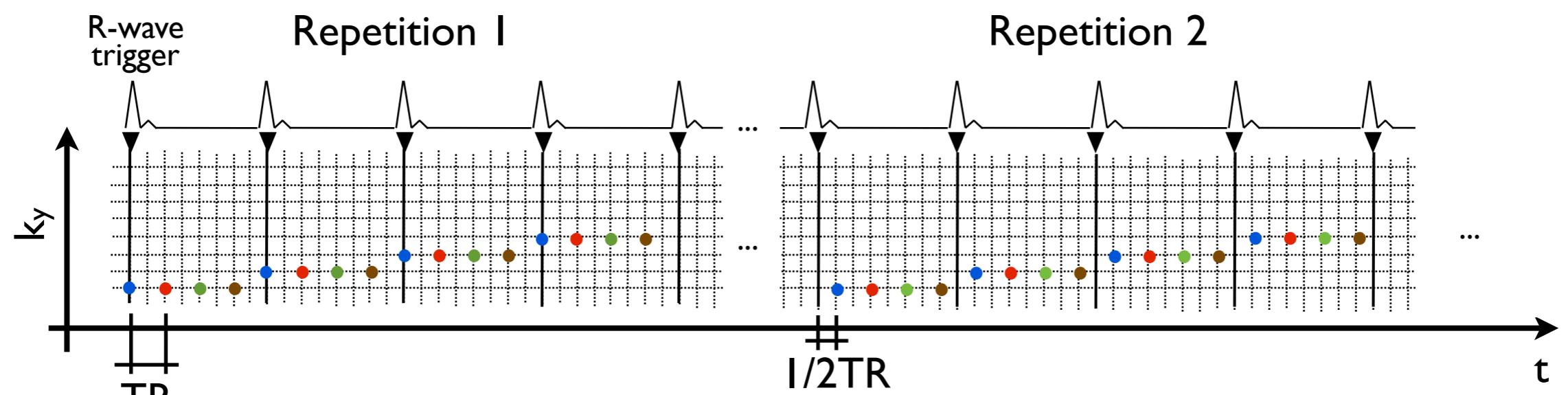
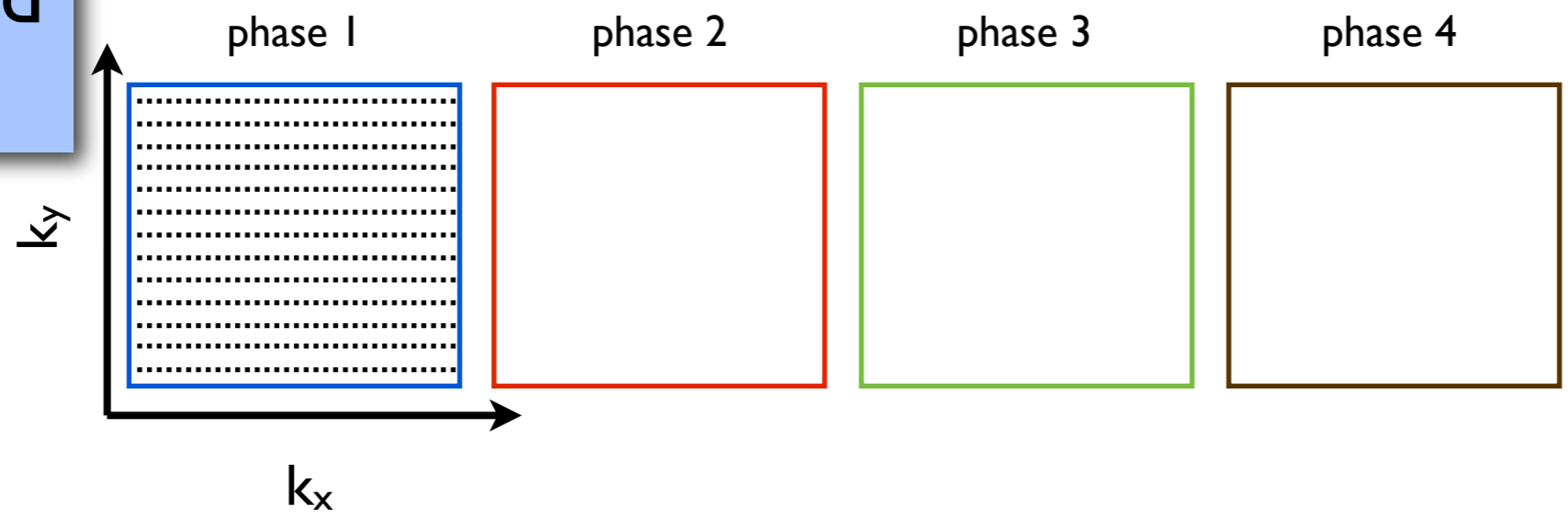
Basic cine



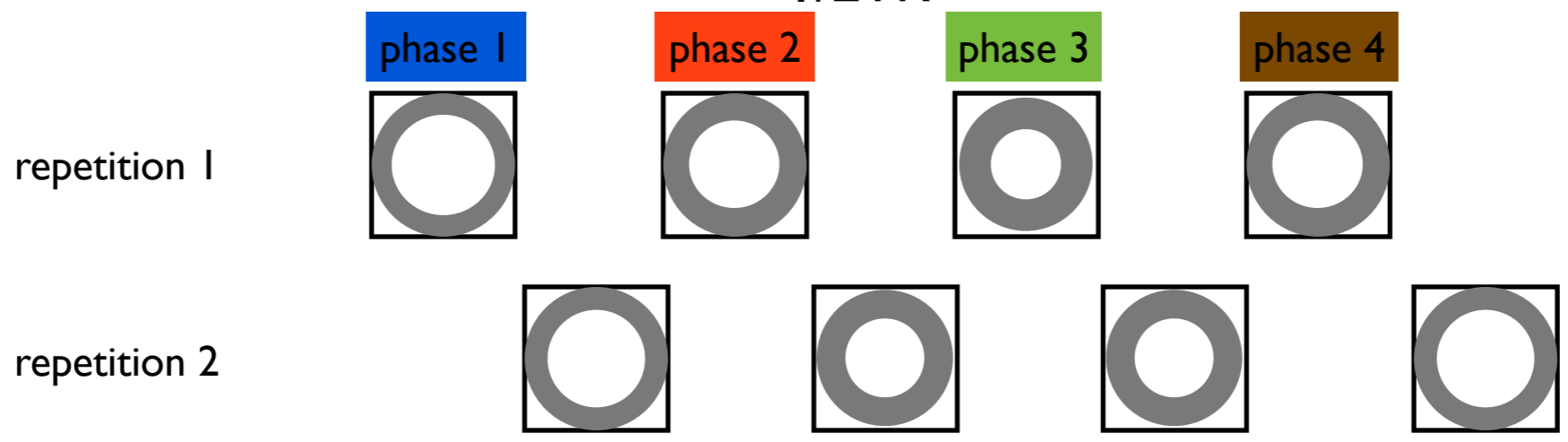
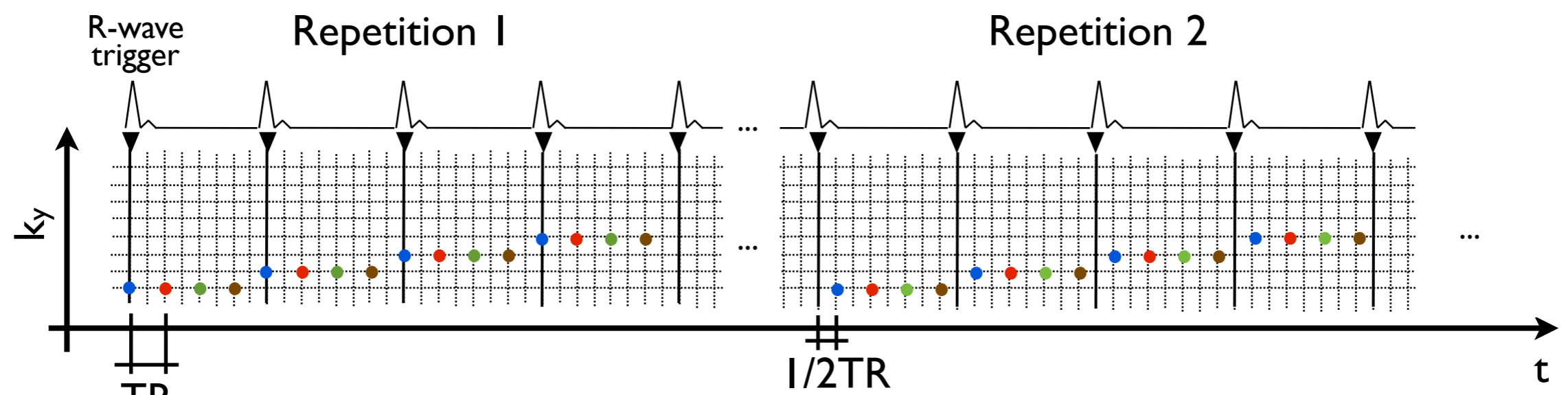
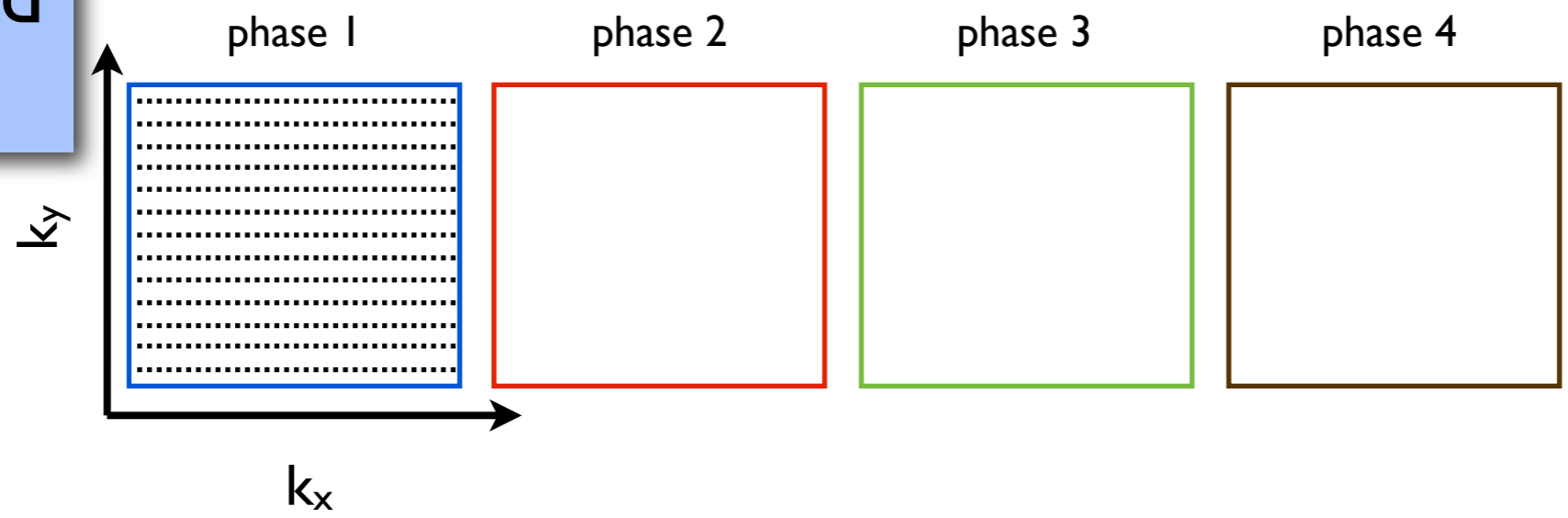
Interleaved cine



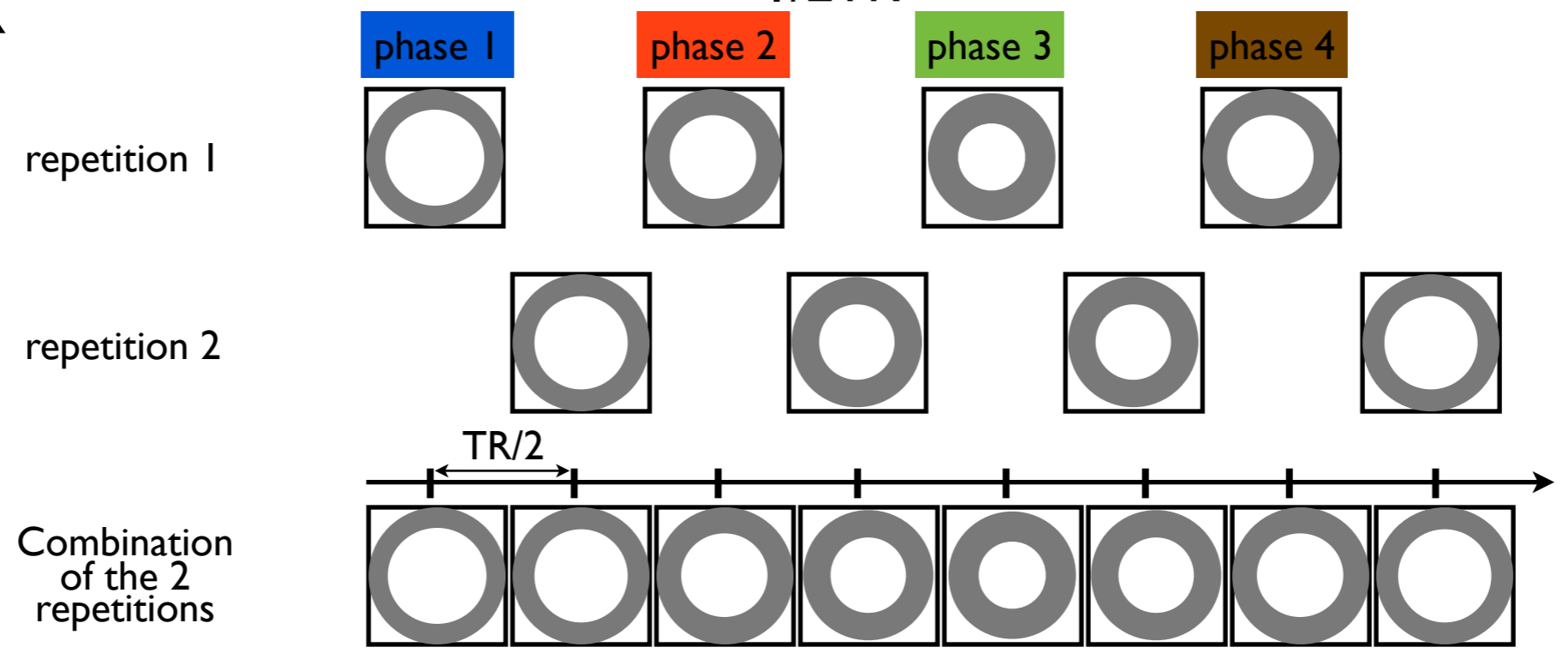
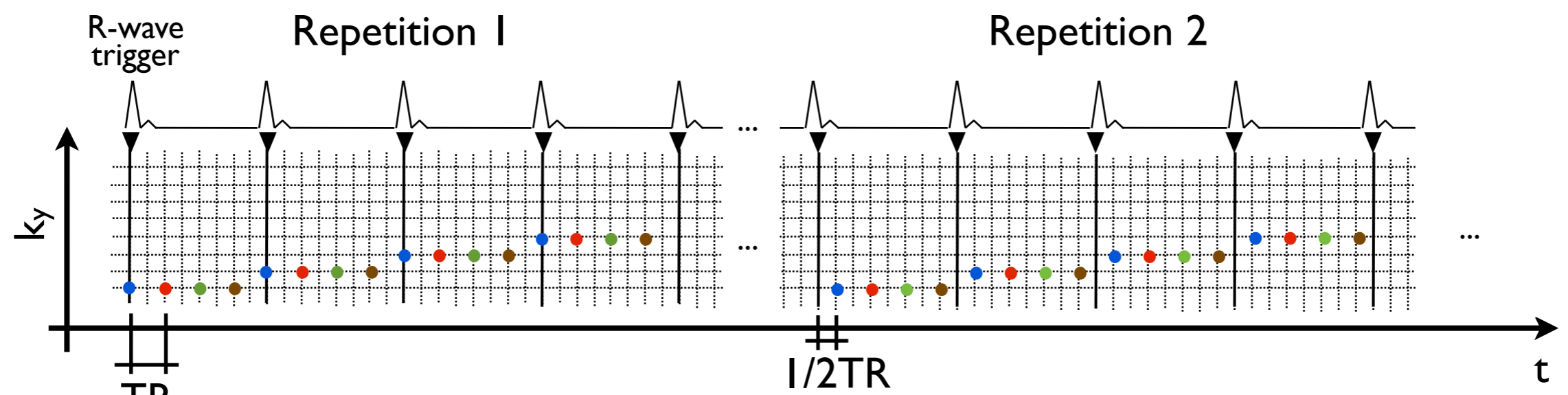
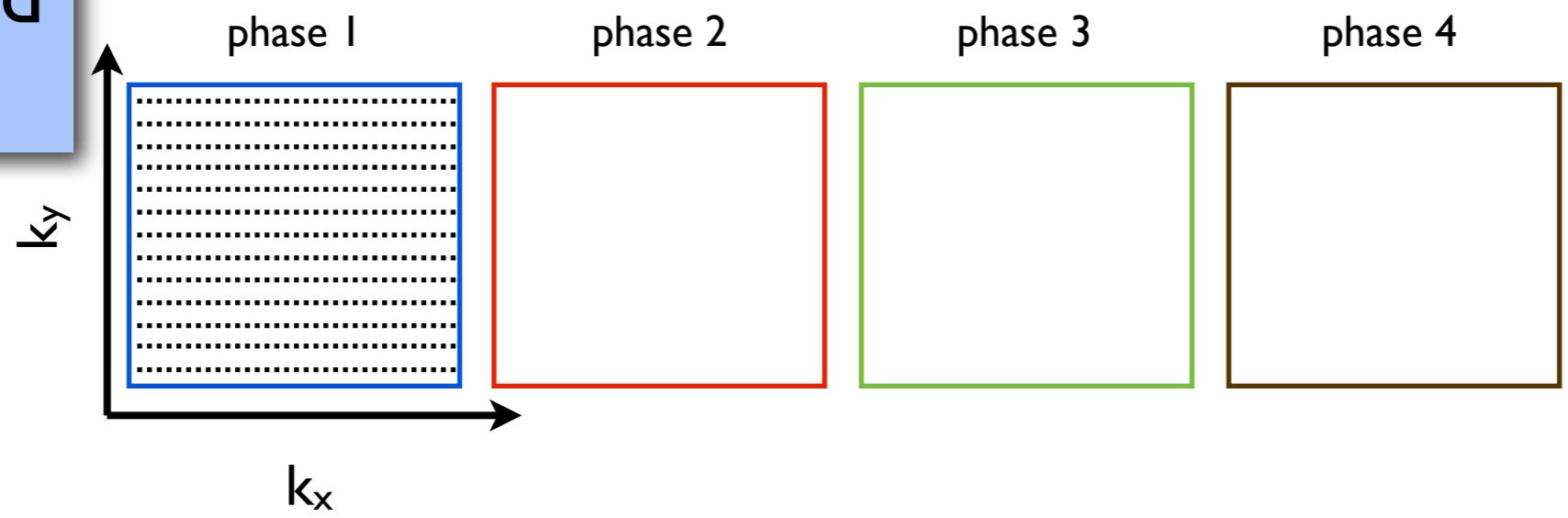
Interleaved cine



Interleaved cine



Interleaved cine



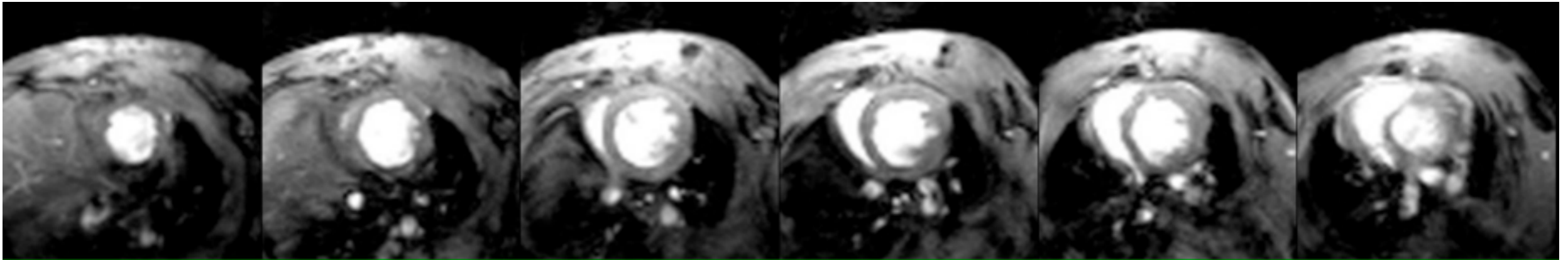
First results

Basic sequence (TR=13.5ms)

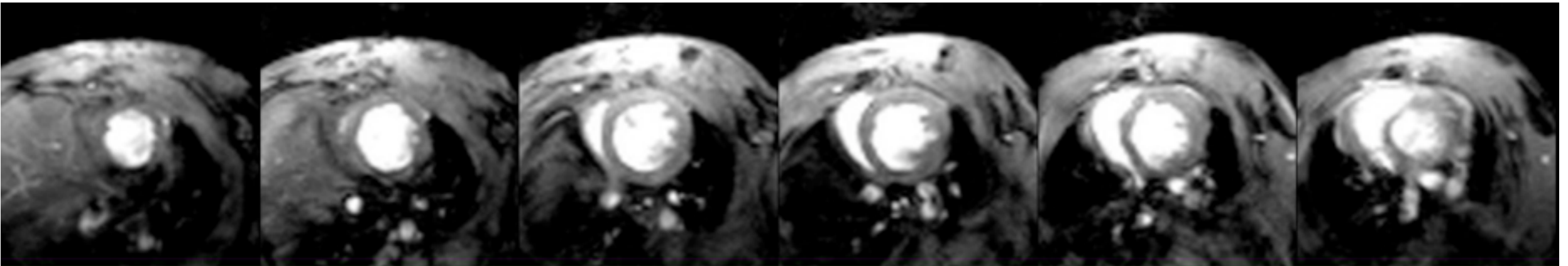
Interleaved cine (TR=6.8ms)

First results

Basic sequence (TR=13.5ms)

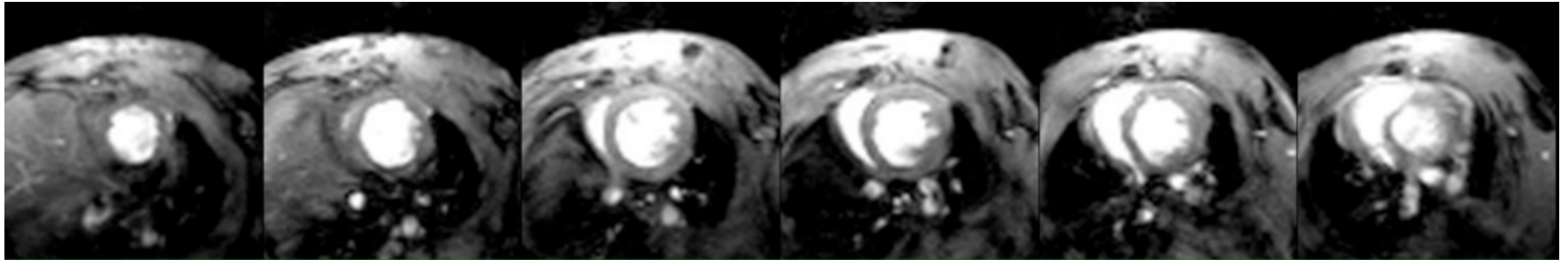


Interleaved cine (TR=6.8ms)

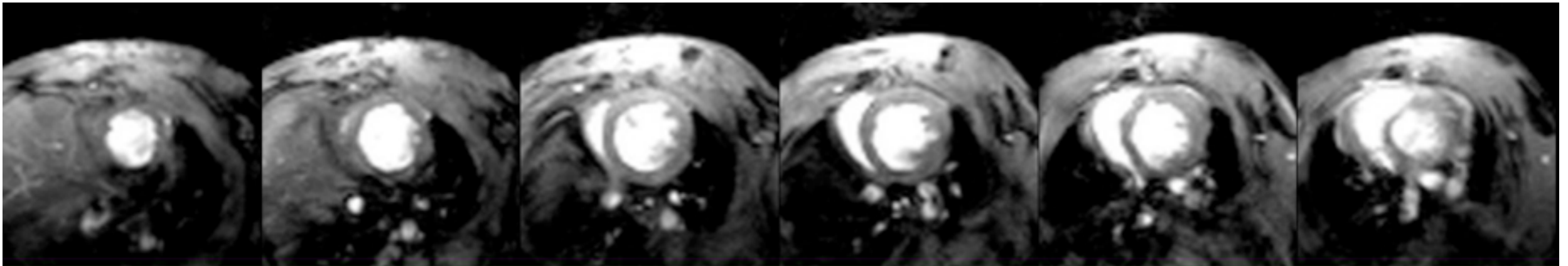


First results

Basic sequence (TR=13.5ms)



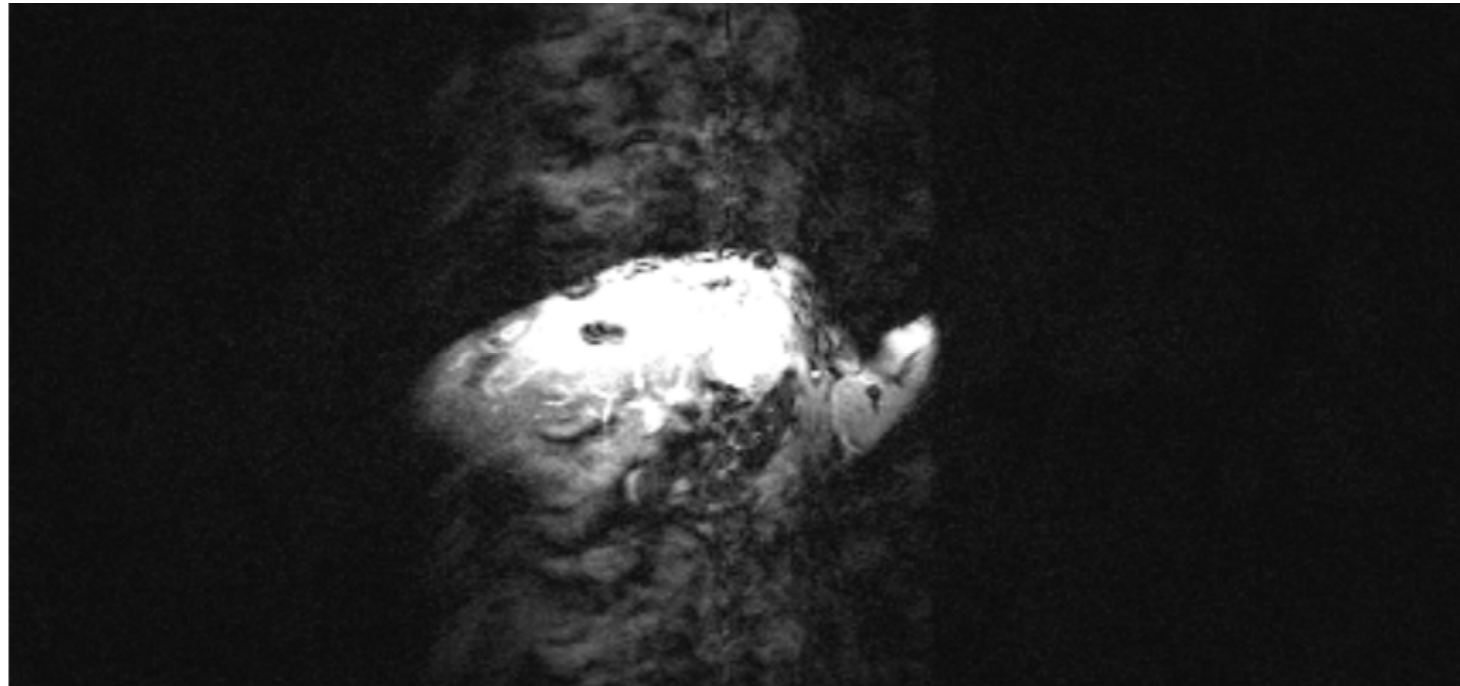
Interleaved cine (TR=6.8ms)



“Flickering” artifact due to combination of images with different artifacts (flow)

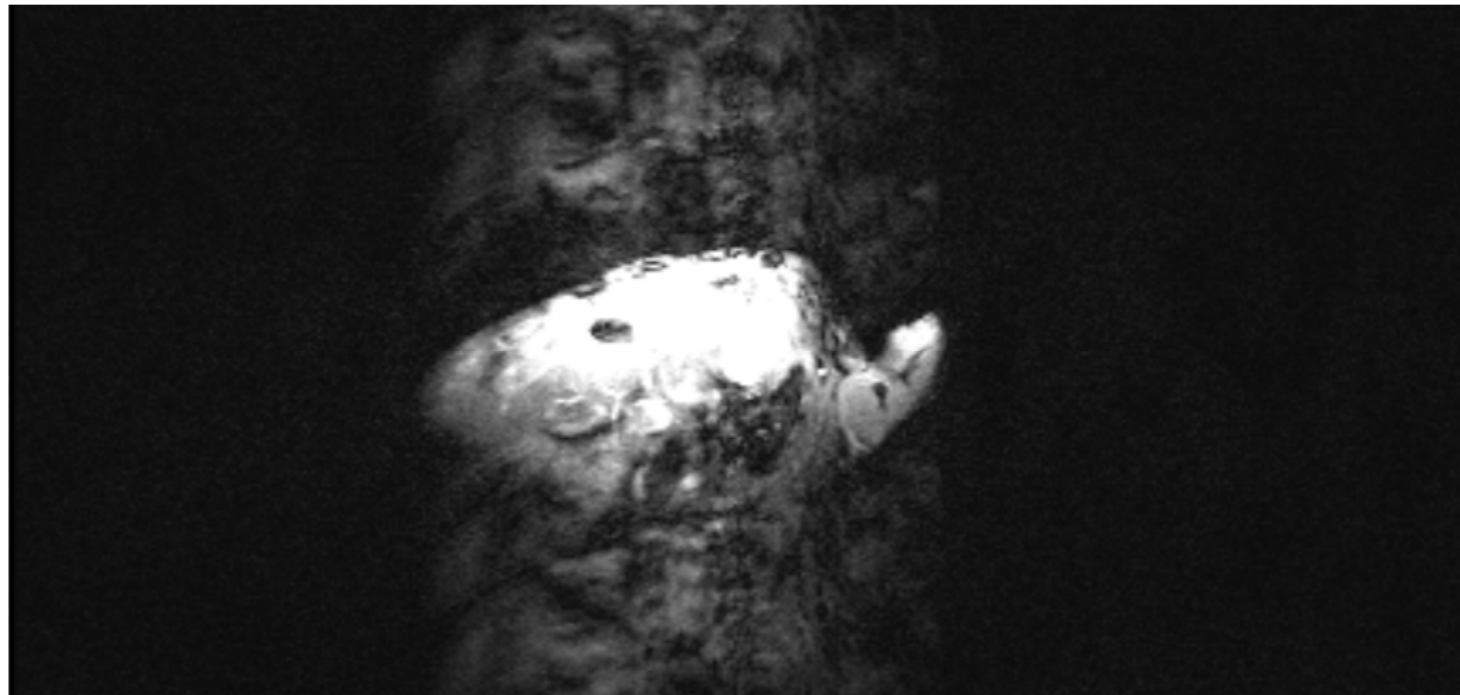
Flickering artifact

Phase I - rep 1



WW/WL set to see
ghosting artifacts

Phase I - rep 2



Flickering artifact

Phase I - rep 1



WW/WL set to see
ghosting artifacts

Phase I - rep 2



Image enhancement

Variational model

$$J(s) = \sum_n \|f_n - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

n: temporal indice
^: Fourier transform

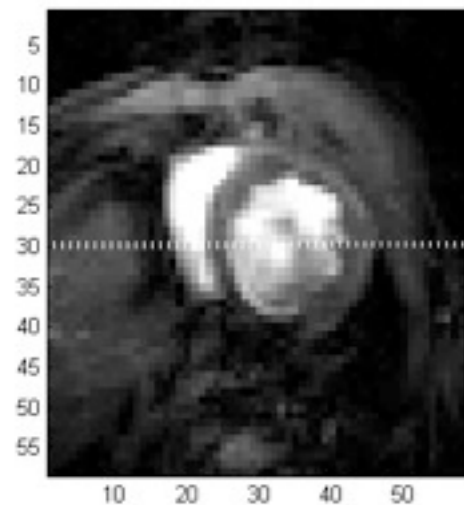
Image enhancement

Variational model

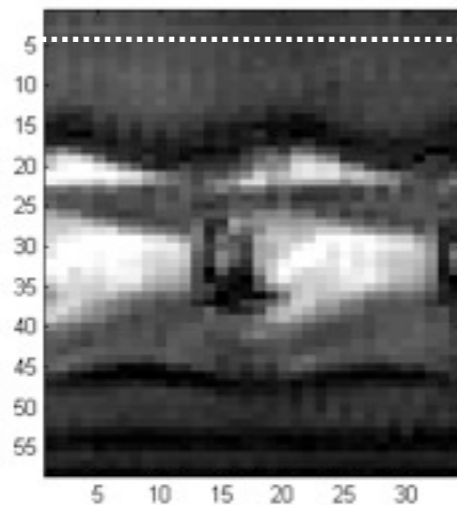
$$J(s) = \sum_n \|f_n - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

n: temporal indice
^: Fourier transform

Case I:



original



After soft thresh.

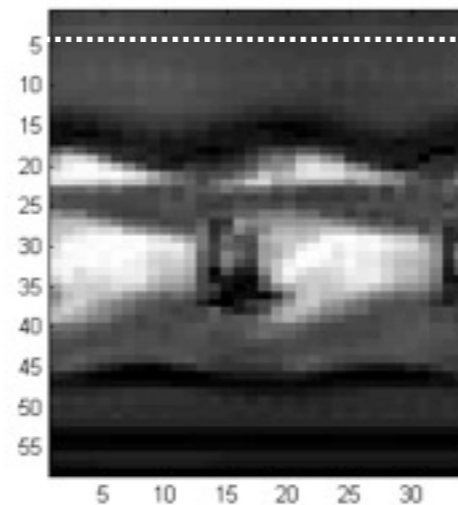


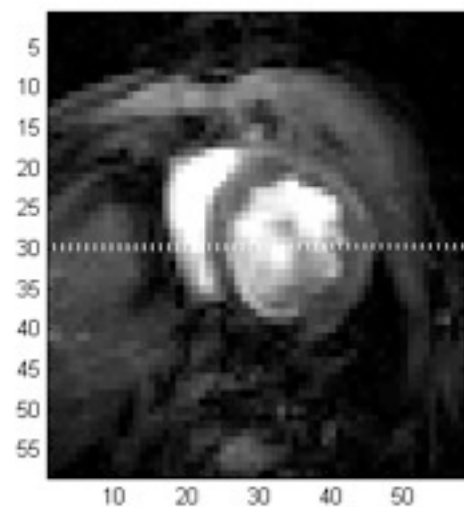
Image enhancement

Variational model

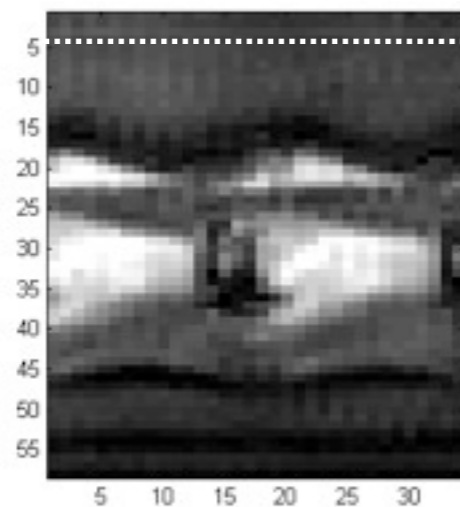
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Case I:



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After soft thresh.

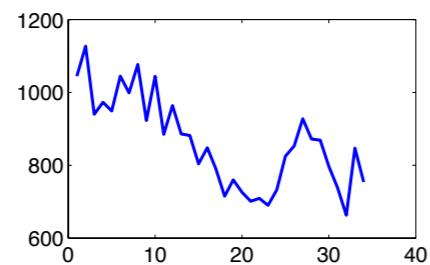
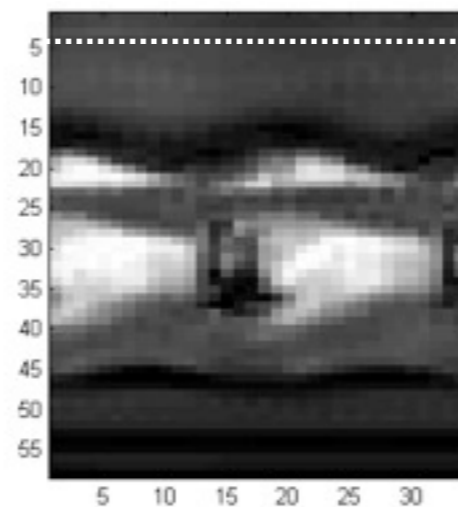


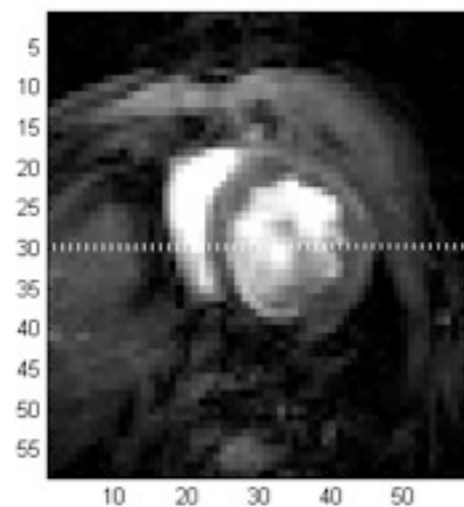
Image enhancement

Variational model

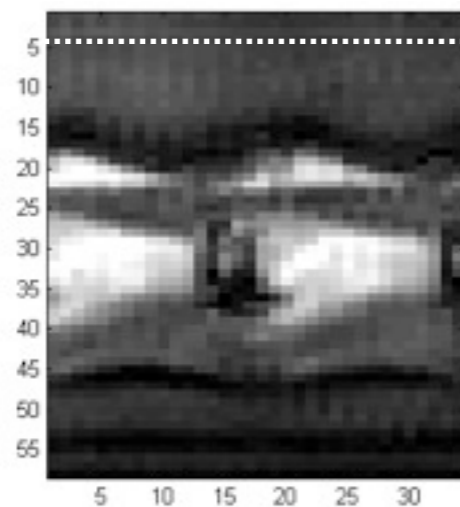
$$J(s) = \sum_n \|f_n - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

n: temporal indice
^: Fourier transform

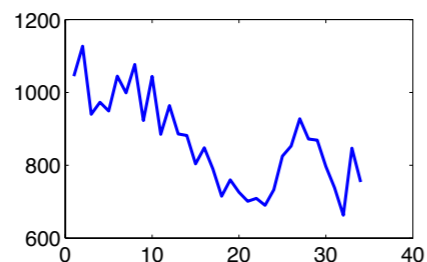
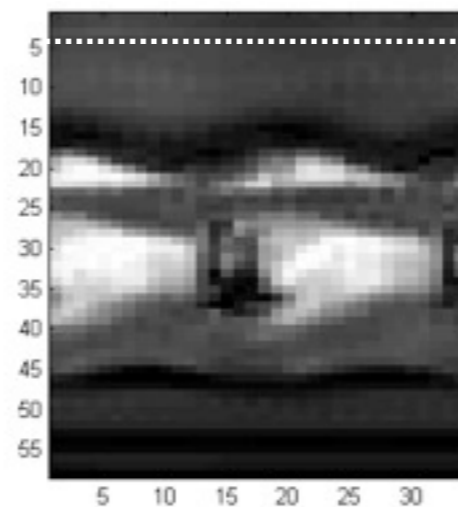
Case I:



original



After soft thresh.



FT

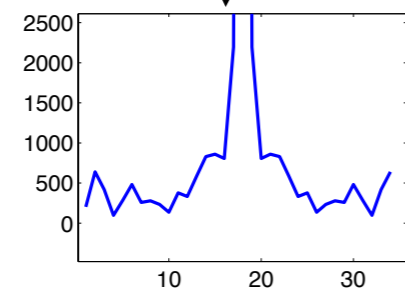


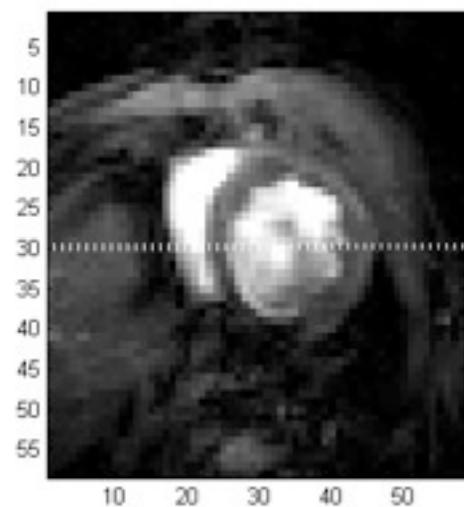
Image enhancement

Variational model

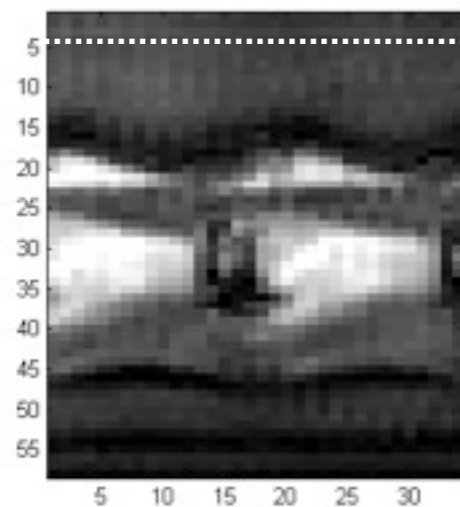
$$J(s) = \sum_n \|f_n - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

n : temporal indice
 $\hat{\cdot}$: Fourier transform

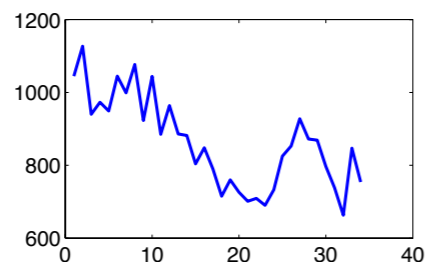
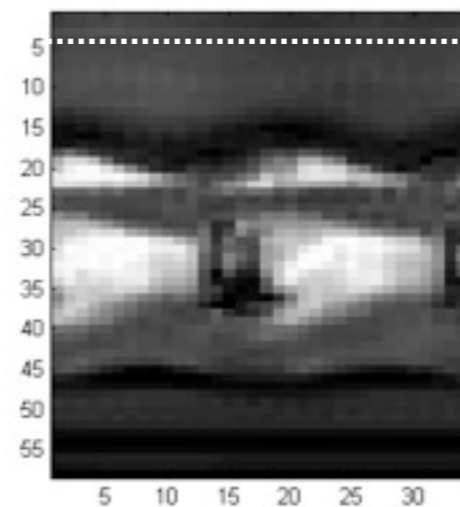
Case I:



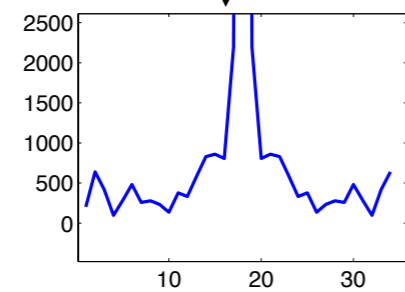
original



After soft thresh.



FT



soft thresh.

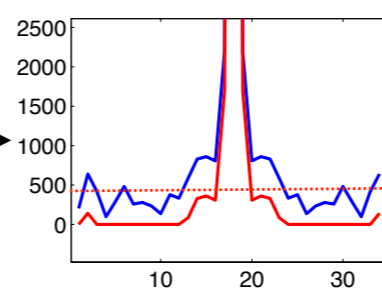


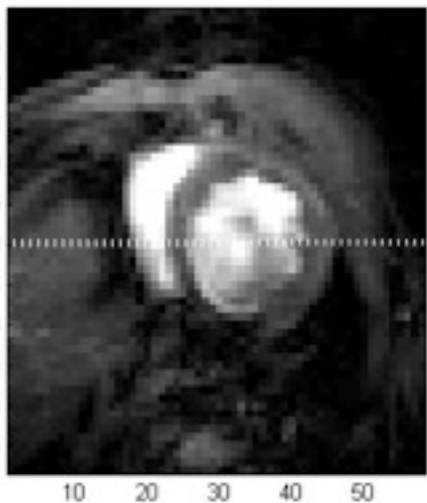
Image enhancement

Variational model

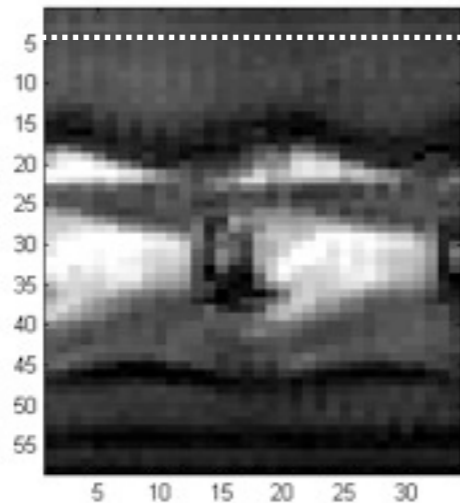
$$J(s) = \sum_n \|f_n - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

n: temporal indice
 $\hat{\cdot}$: Fourier transform

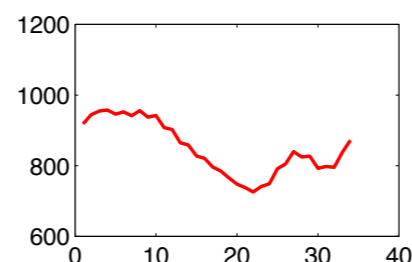
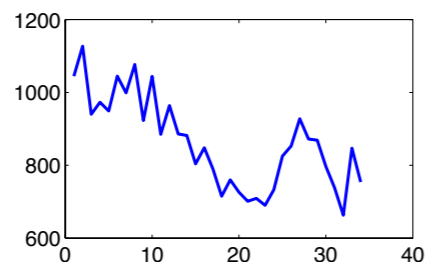
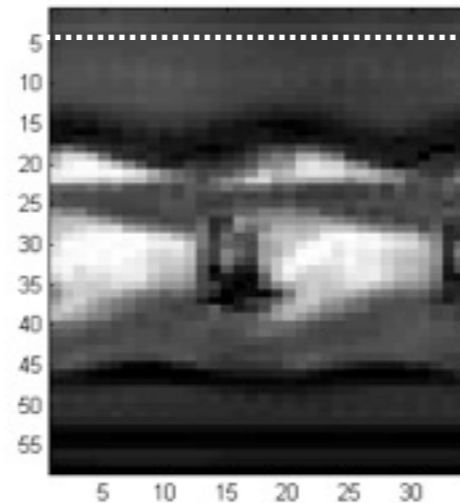
Case I:



original

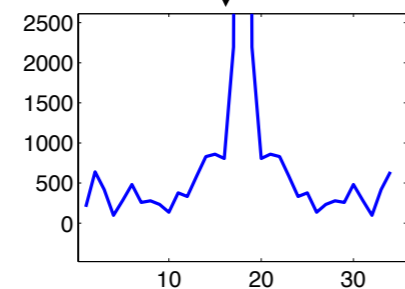


After soft thresh.



FT

FT⁻¹



soft thresh.

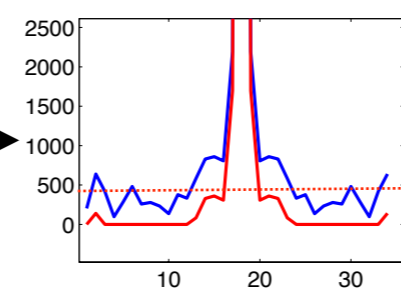


Image enhancement

Variational model

$$J(s) = \sum_n \|f_n - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

n: temporal indice
^: Fourier transform

Image enhancement

Variational model

$$J(s) = \sum_n \|f_n - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

n: temporal indice
^: Fourier transform

Case 2:

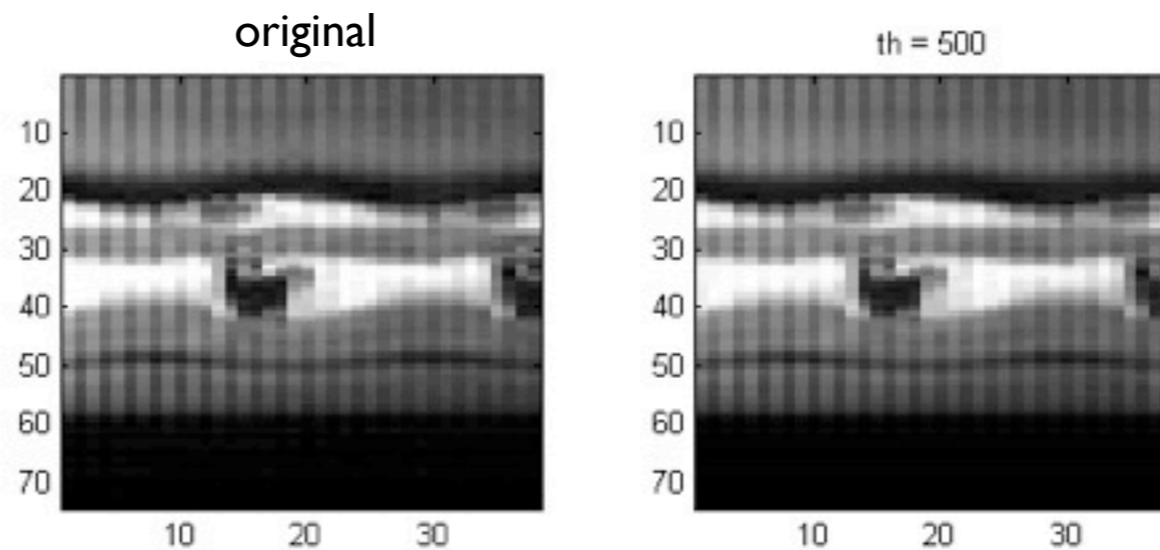


Image enhancement

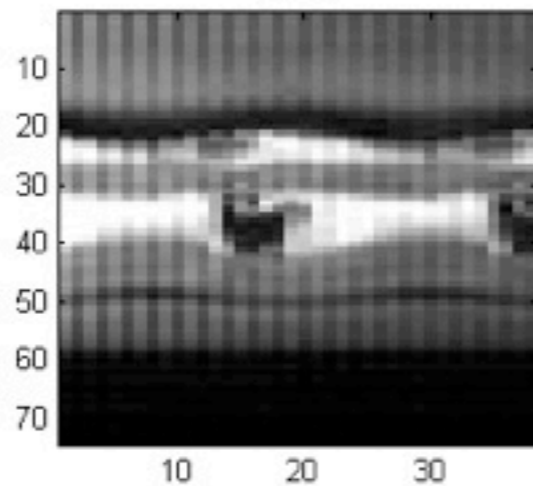
Variational model

$$J(s) = \sum_n \|f_n - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

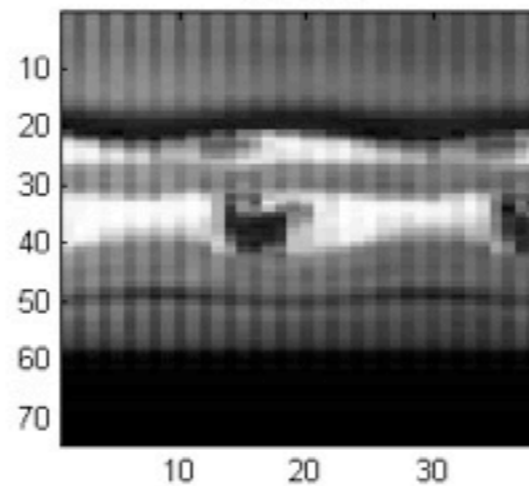
n: temporal indice
^: Fourier transform

Case 2:

original



th = 500



th = 5000

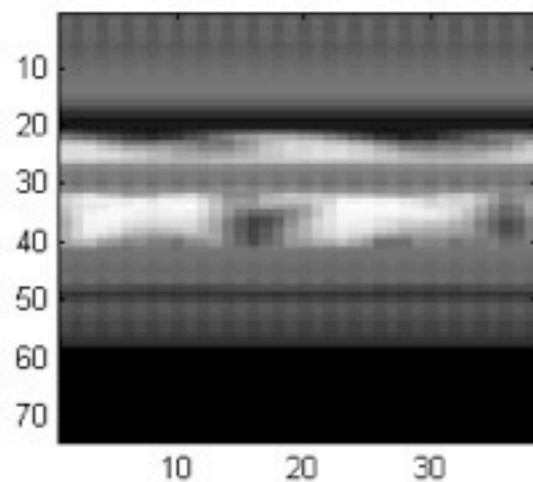


Image enhancement

Variational model

$$J(s) = \sum_n \|f_n - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

n: temporal indice
^: Fourier transform

Case 2:

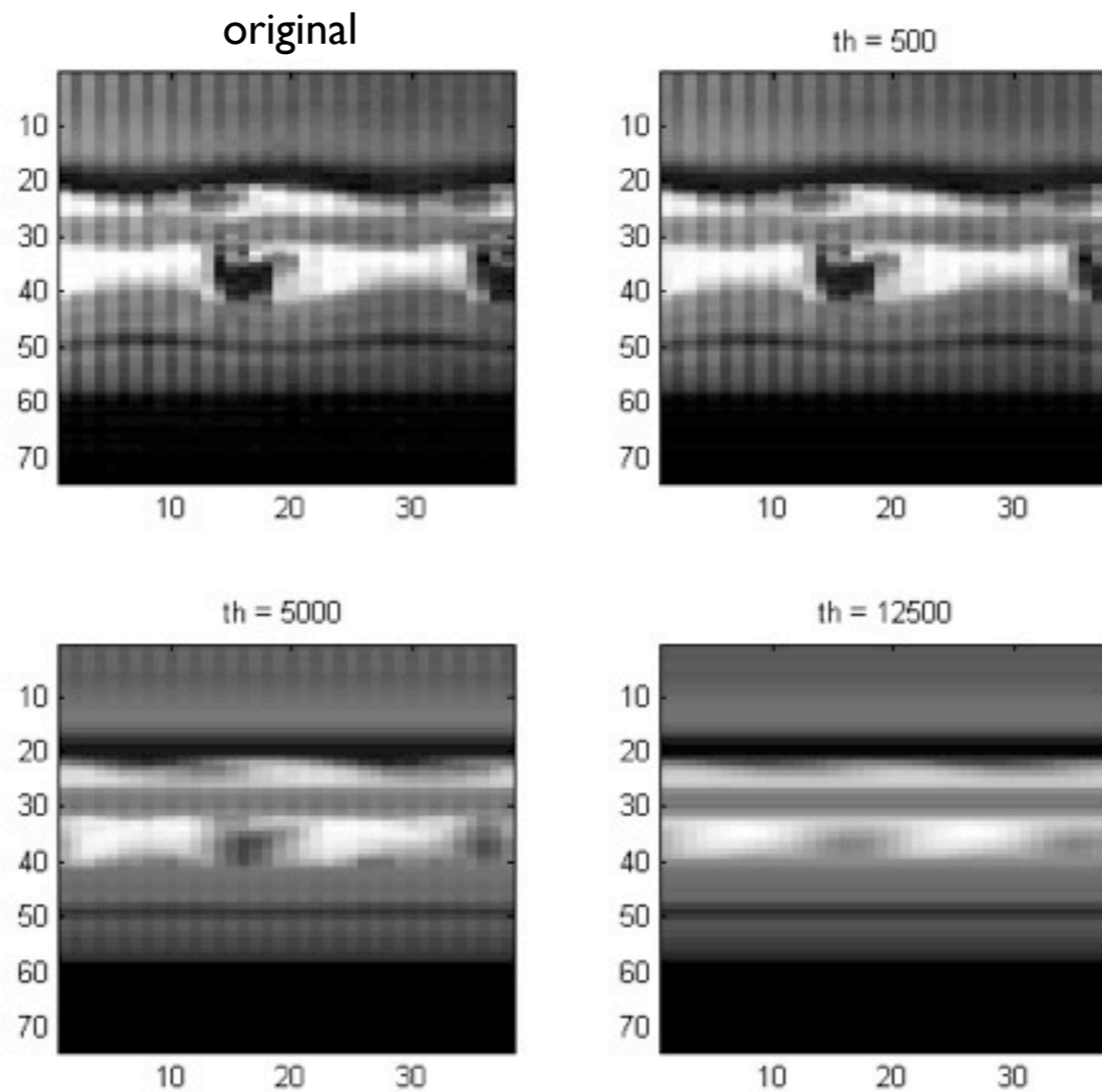


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

n: temporal indice
^: Fourier transform

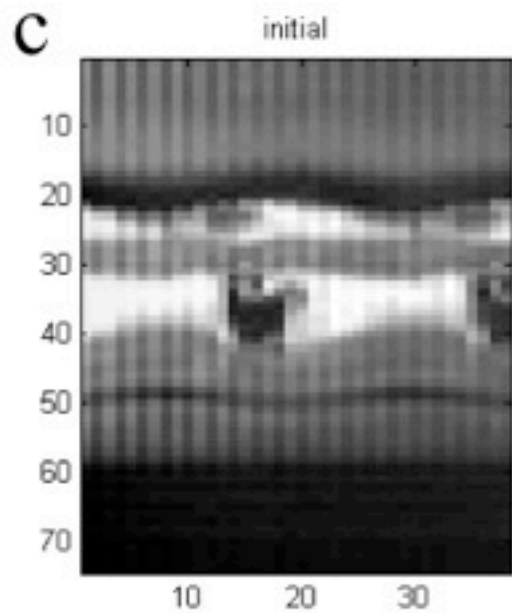
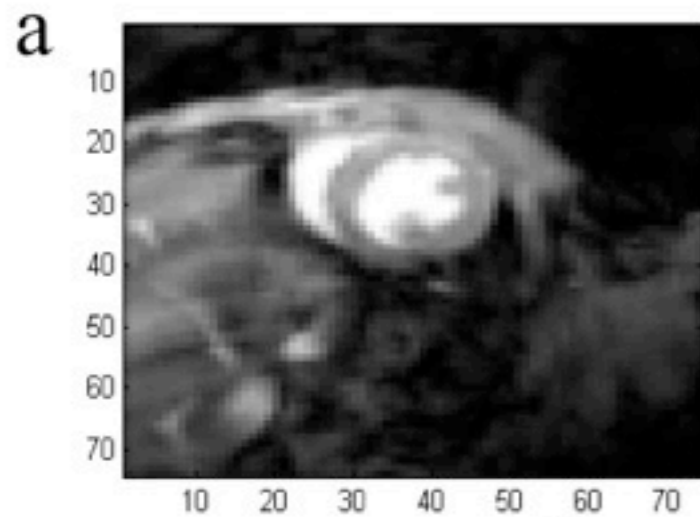


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

n: temporal indice
^: Fourier transform

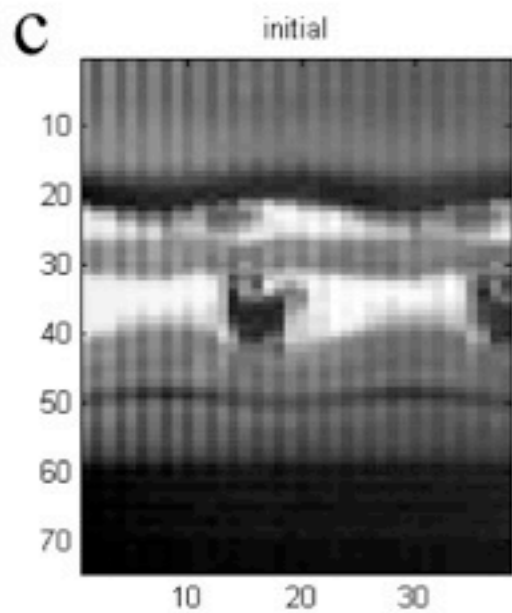
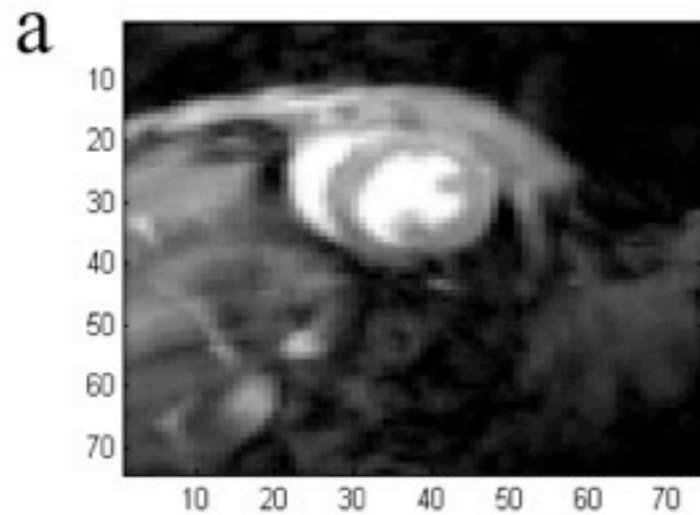
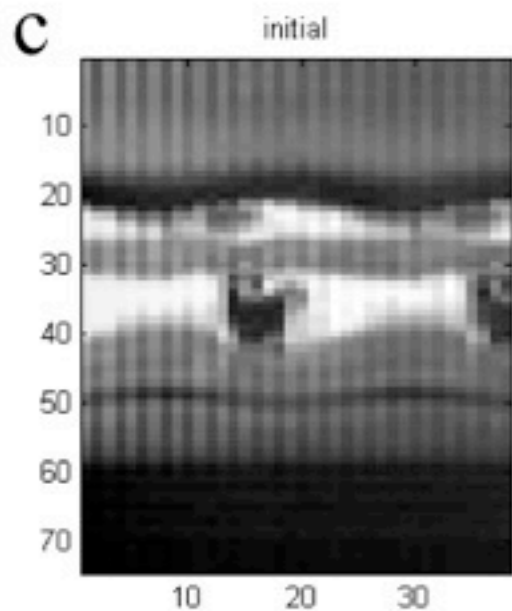
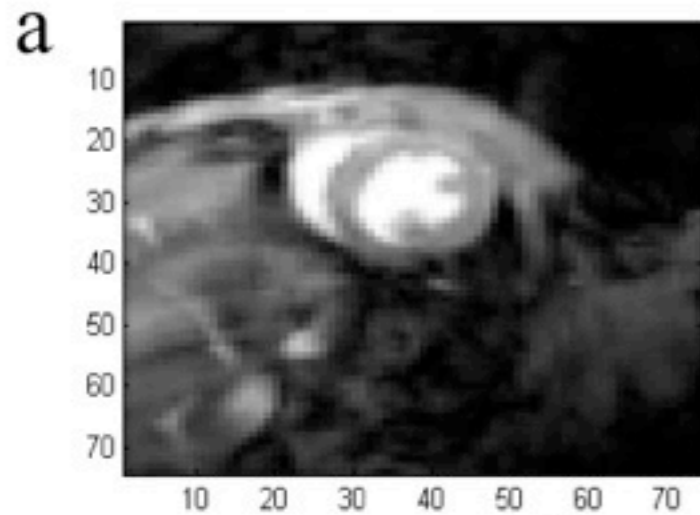


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$ n: temporal indice
^: Fourier transform



Step I $\mu_n = \frac{1}{N_x} \sum_x \frac{1}{N_y} \sum_y f_n(x, y)$

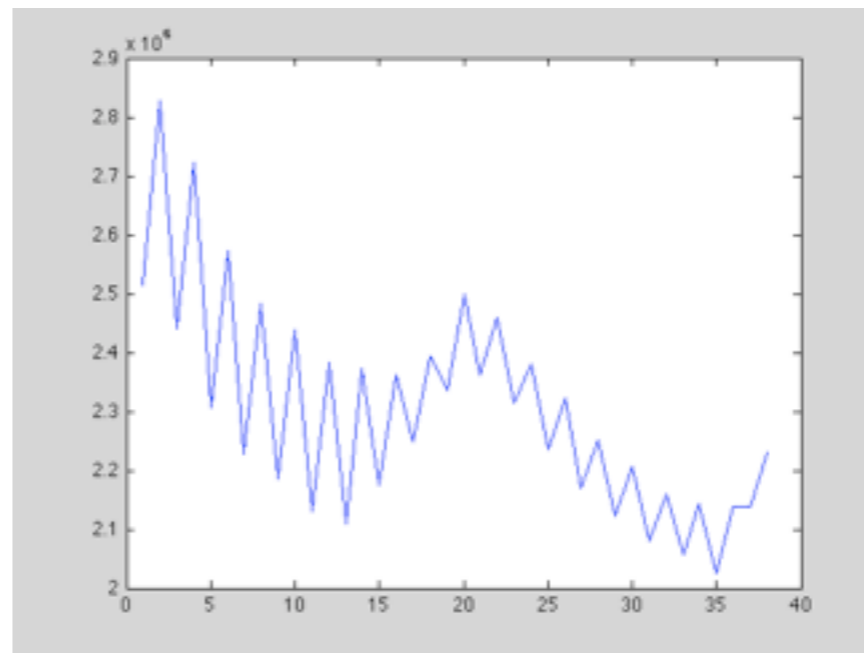
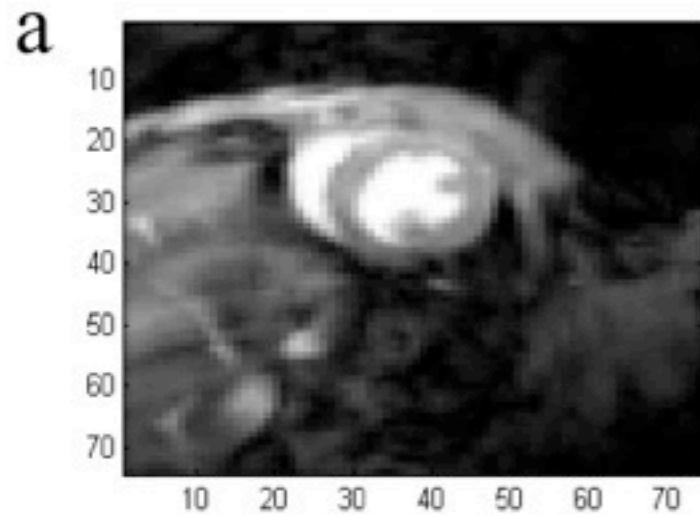


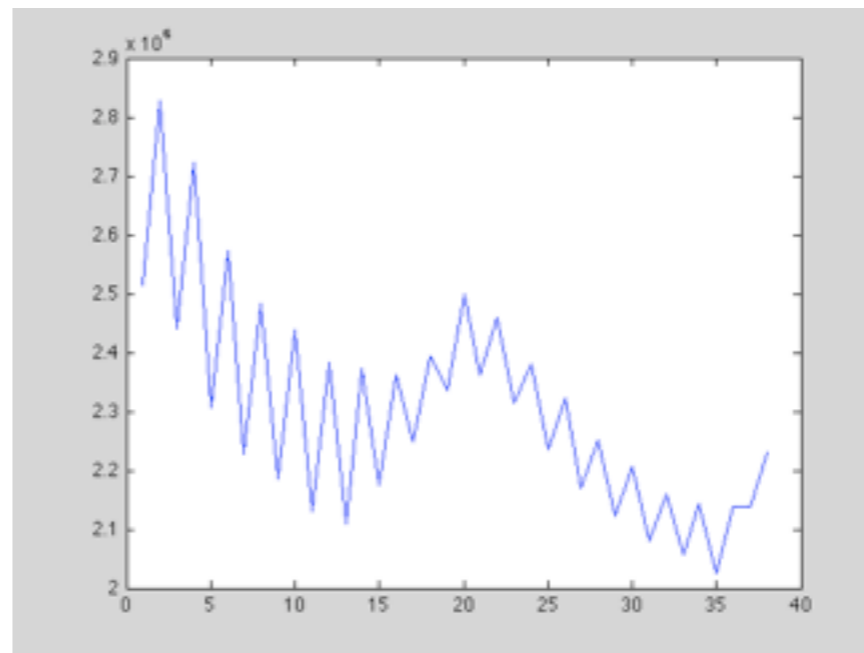
Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

n: temporal indice
 \wedge : Fourier transform



Step I $\mu_n = \frac{1}{N_x} \sum_x \frac{1}{N_y} \sum_y f_n(x, y)$



fit ω, ϕ

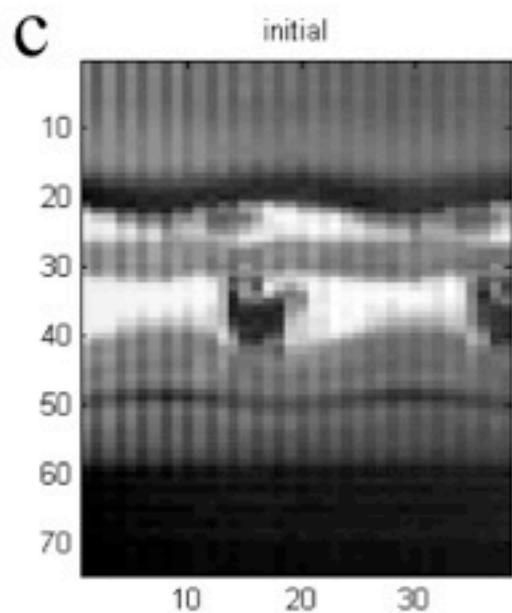


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

n: temporal indice
^: Fourier transform

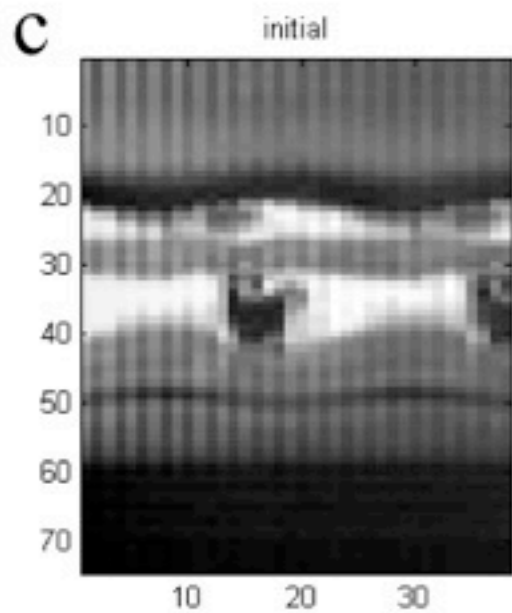
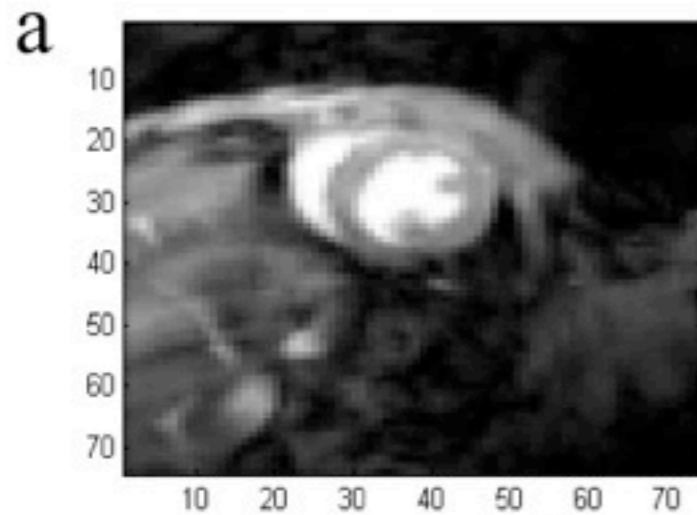


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

n: temporal indice
^: Fourier transform

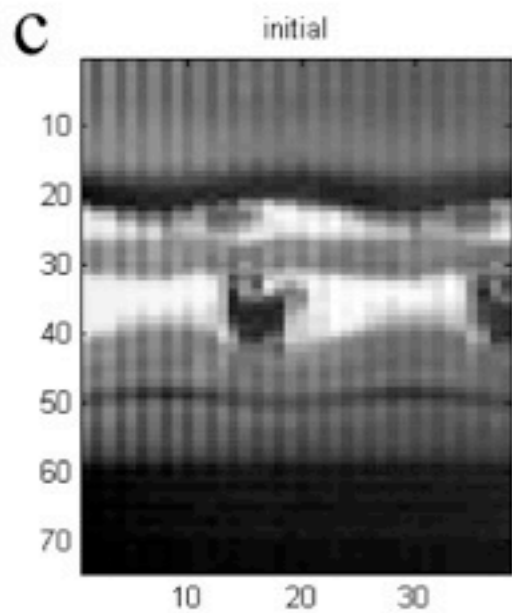
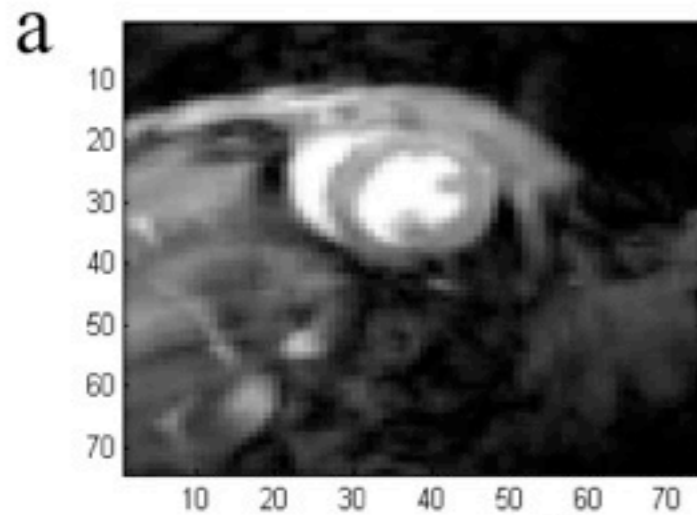
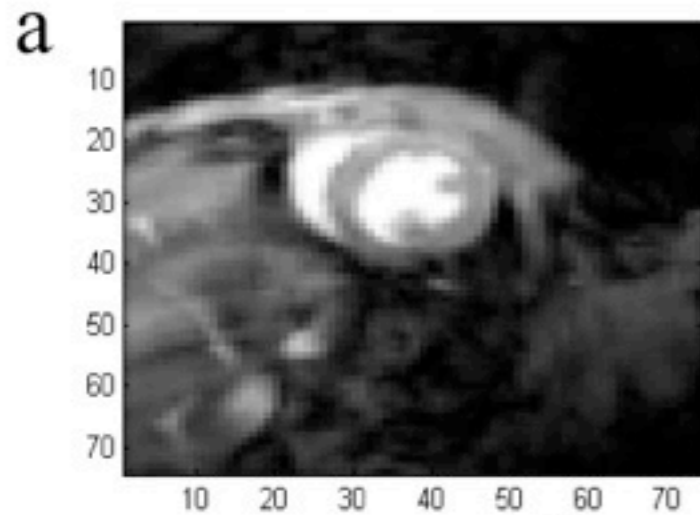


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

n: temporal indice
^: Fourier transform



Step 2: fit $A(x,y)$

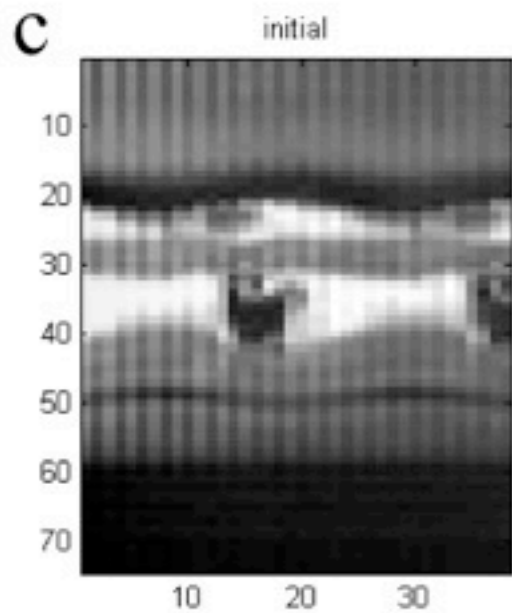
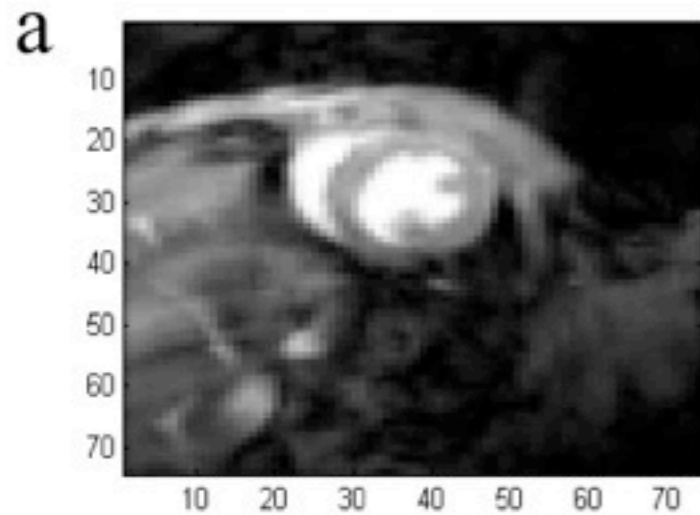


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$ n: temporal indice
 \wedge : Fourier transform



Step 2: fit $A(x,y)$

$$\frac{\partial J}{\partial A} = 0$$

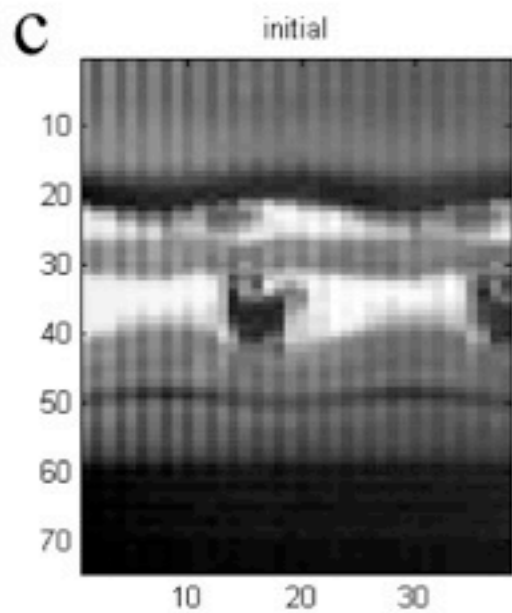
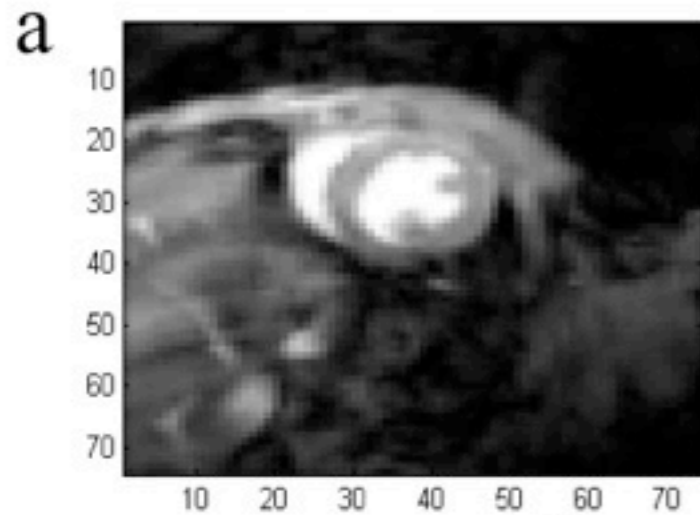


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$ n: temporal indice
 \wedge : Fourier transform



Step 2: fit $A(x,y)$

$$\frac{\partial J}{\partial A} = 0$$

$$A(x, y) = \frac{\sum_n f_n(x, y) \sin(\omega n + \phi)}{\sum_n \sin^2(\omega n + \phi)}$$

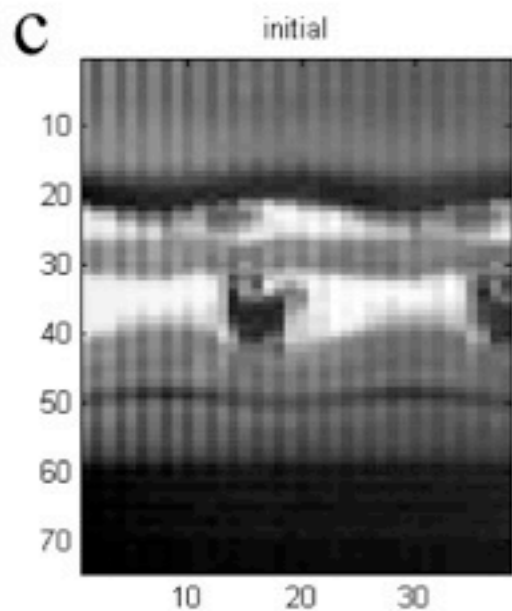


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

n: temporal indice
^: Fourier transform

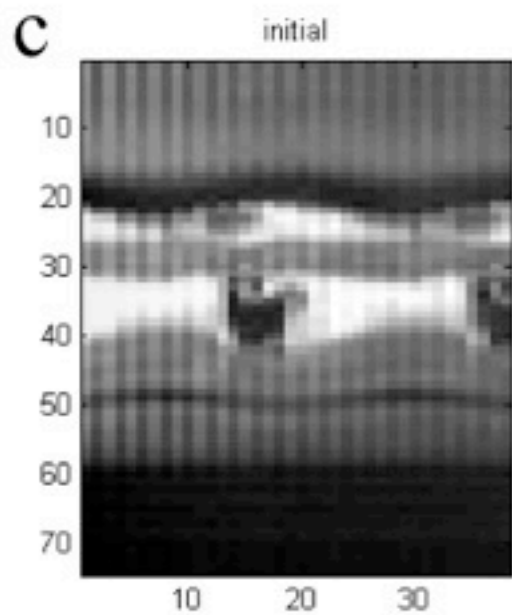
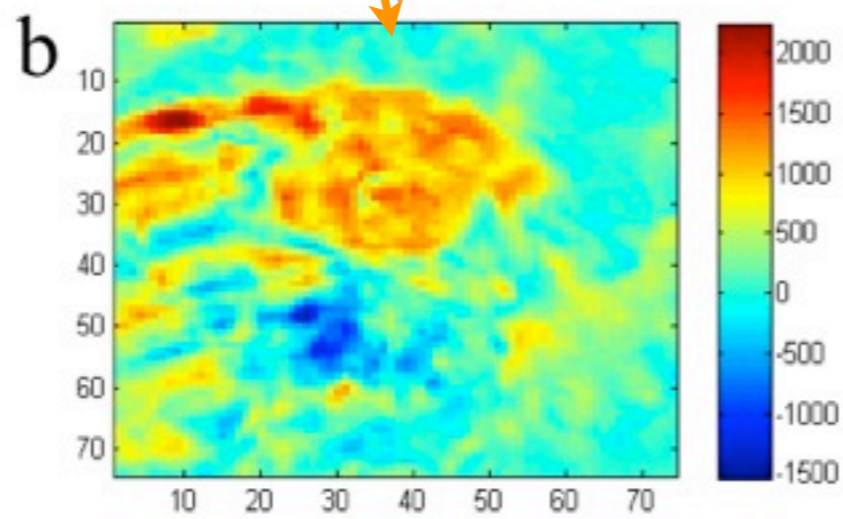
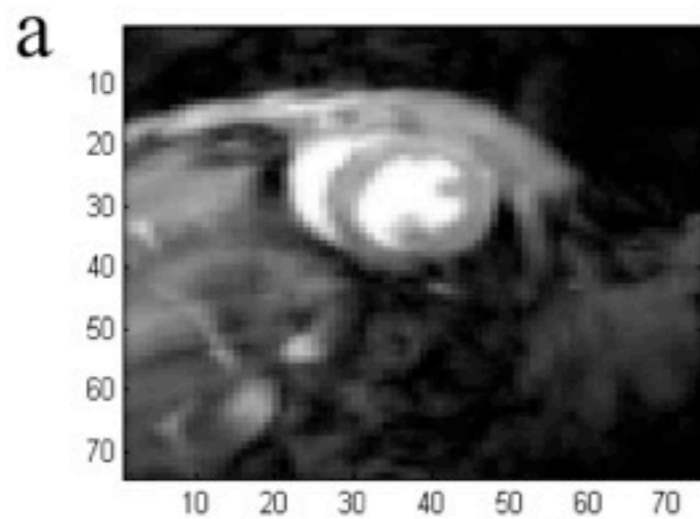


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

n: temporal indice
^: Fourier transform

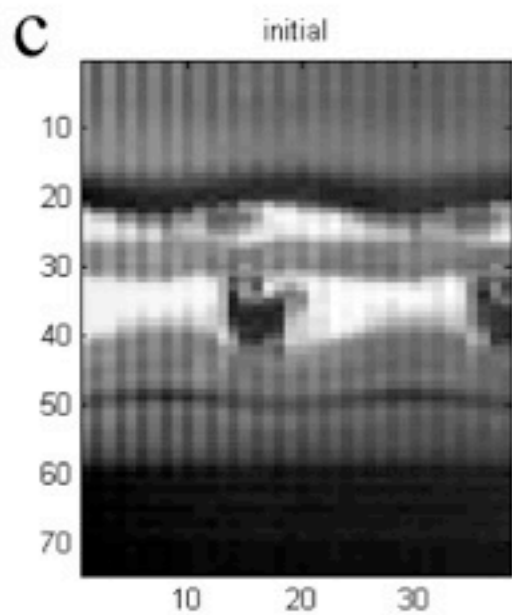
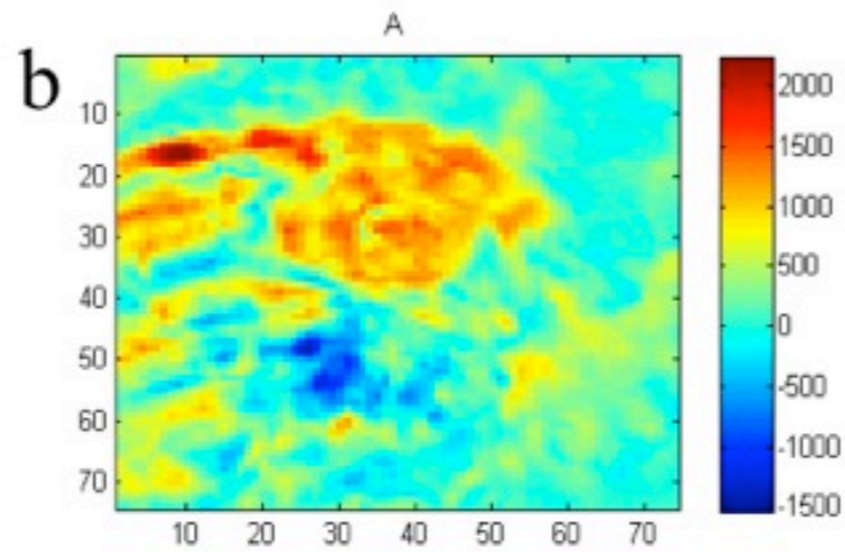
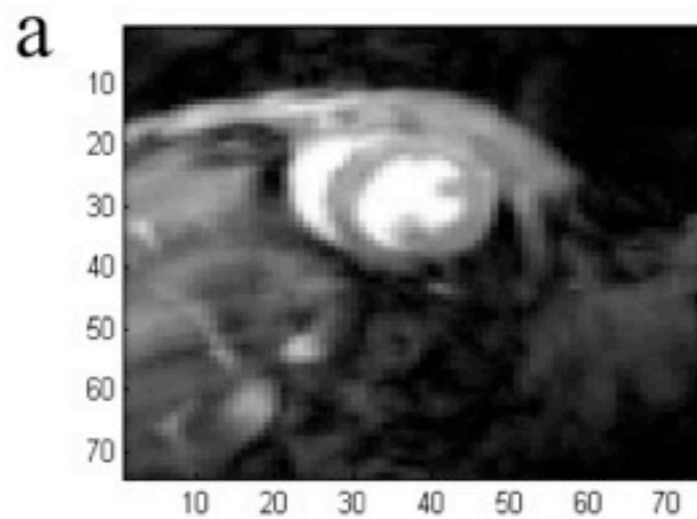
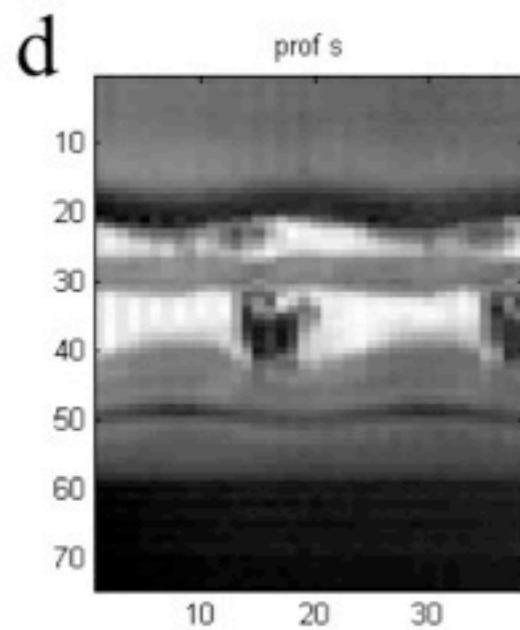
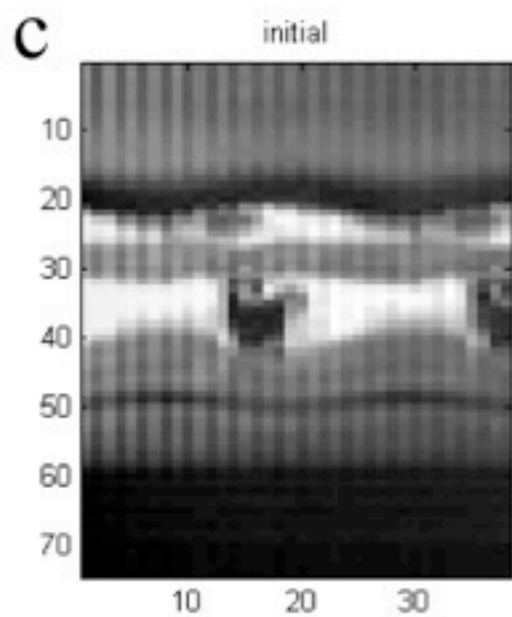
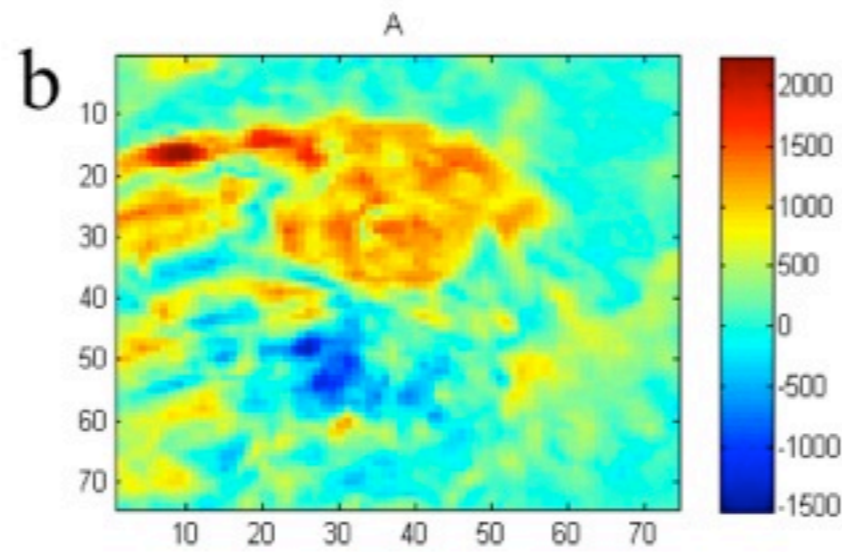
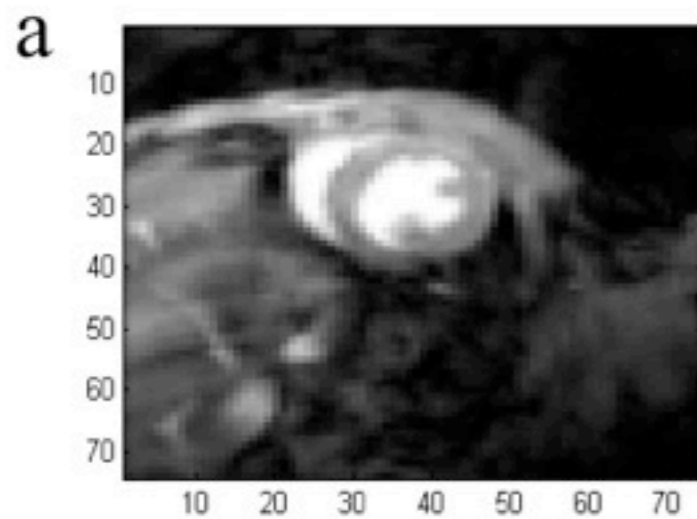


Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

n: temporal indice
^: Fourier transform



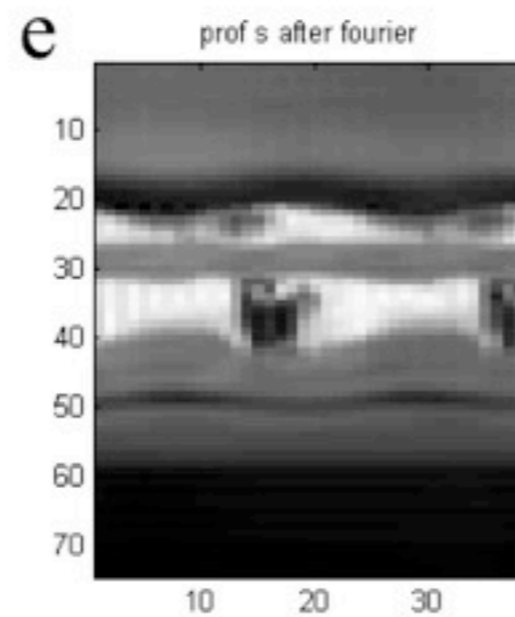
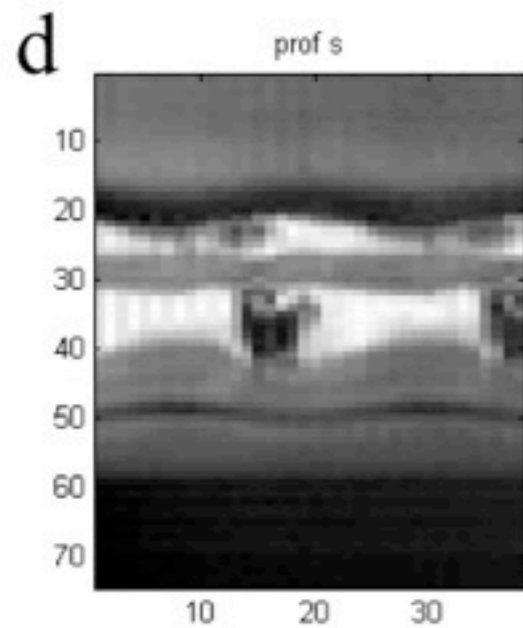
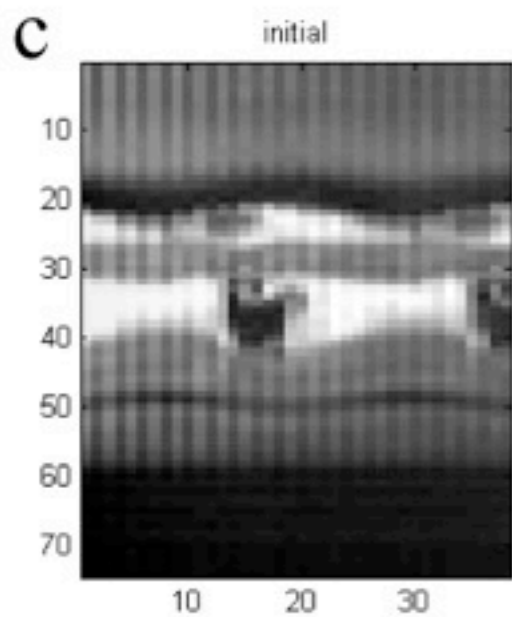
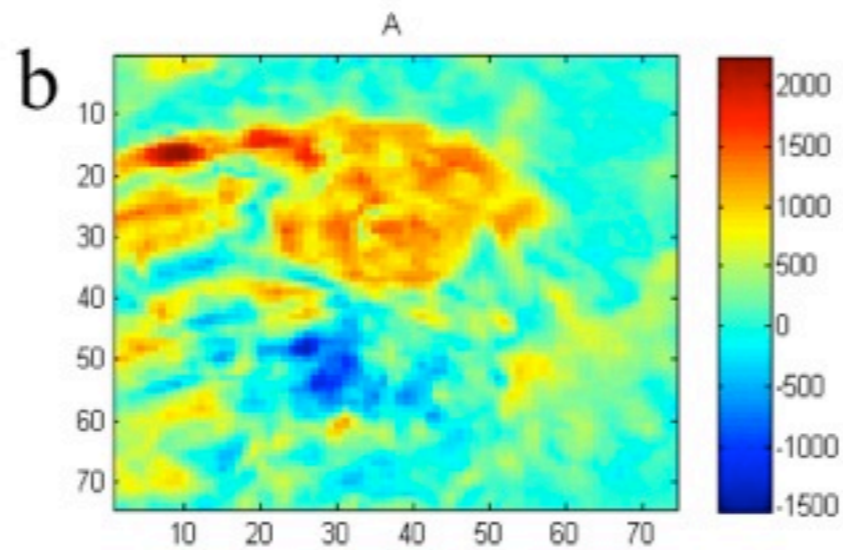
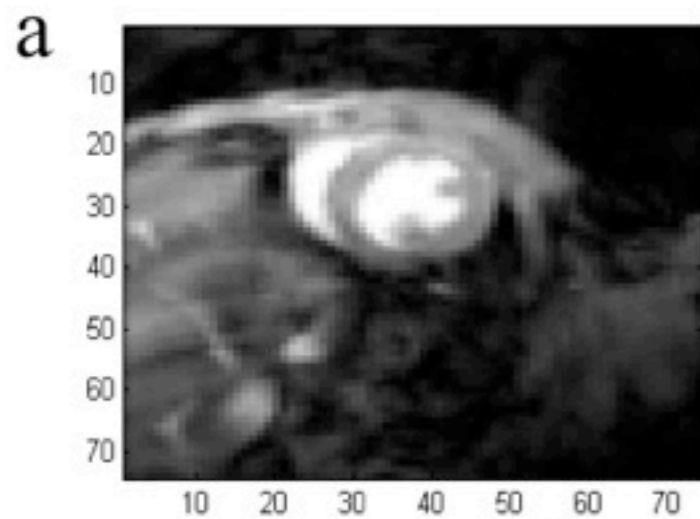
e

prof s after fourier

Image enhancement

Variational model $J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$

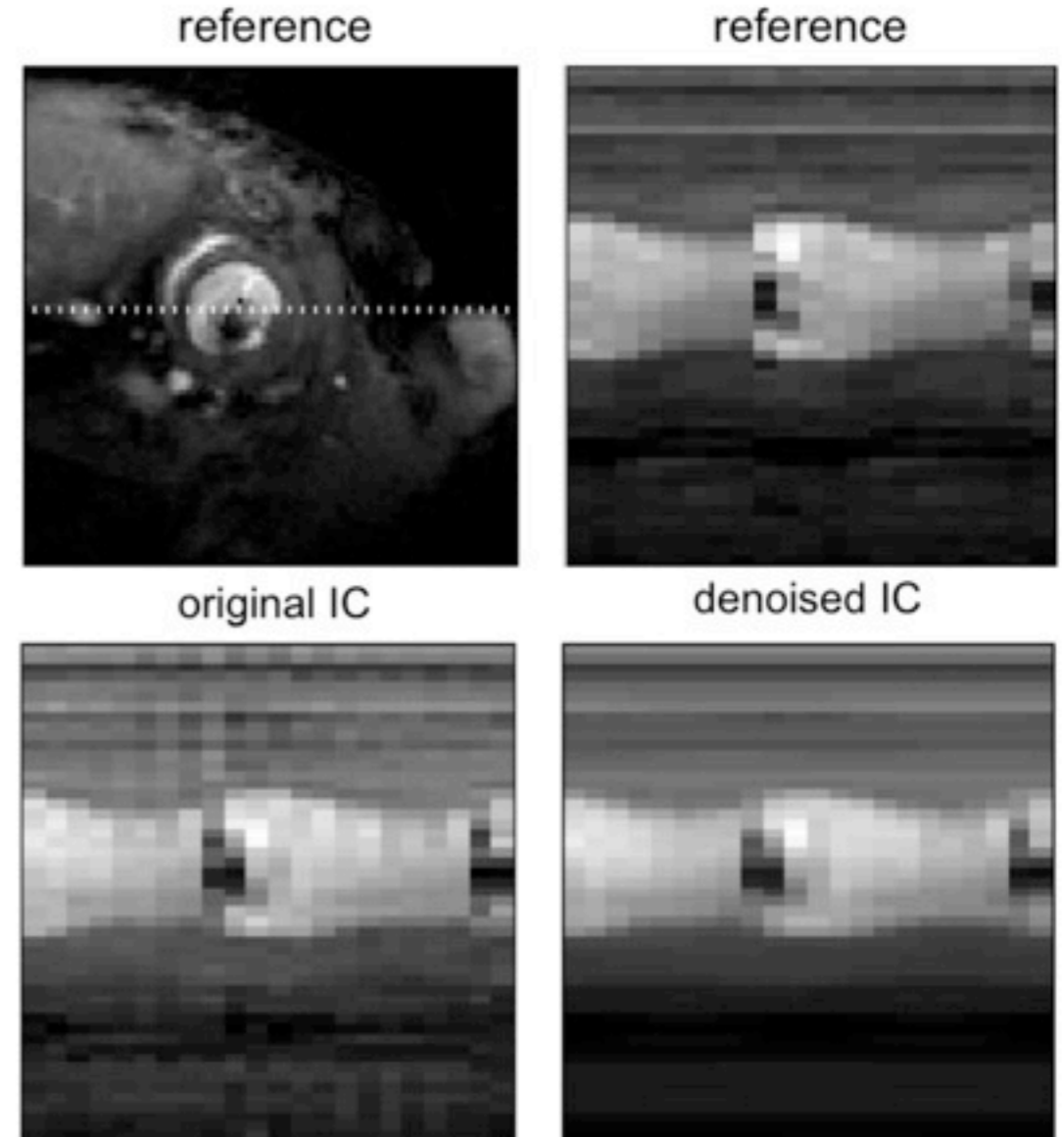
n: temporal indice
^: Fourier transform



Denoising evaluation

Reference:
Basic cine TR 13.5 ms - 2 averages

Interleaved cine:
TR 27 ms - 2 repetitions (no average)
final TR = 13.5 ms



2 evaluation indices :
Energy E

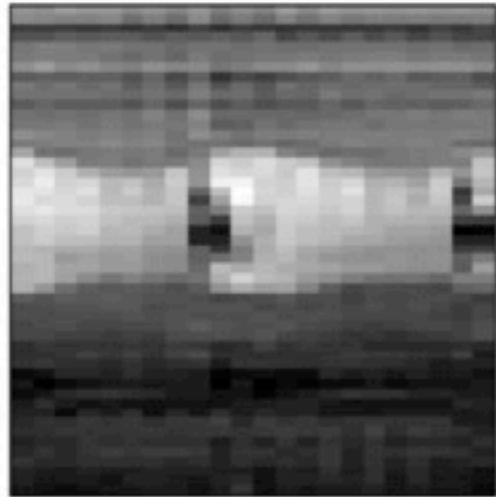
$$E = \sum \left| \frac{\partial}{\partial t} \|I - I_{ref}\|^2 \right|$$

Noise reduction

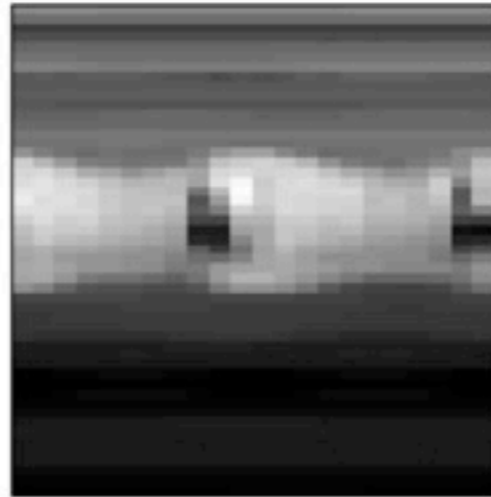
$$10 \log \frac{E_{denoised}}{E_{original}}$$

Denoising evaluation

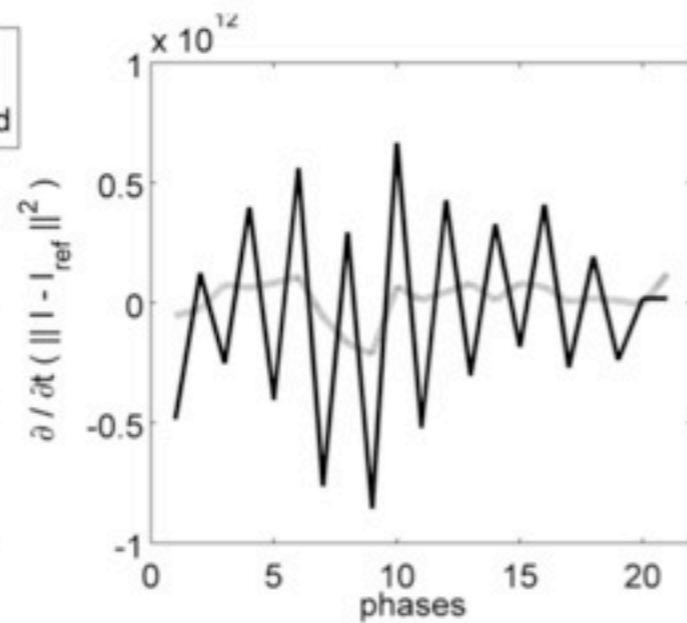
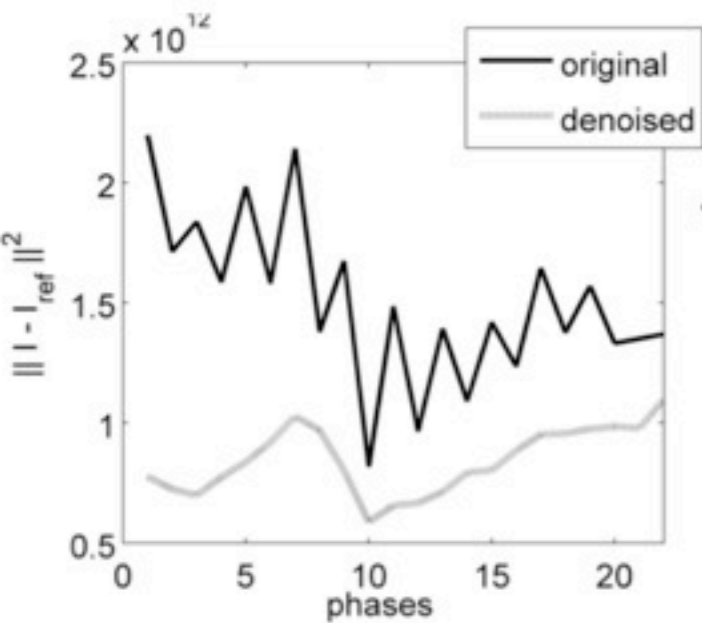
original IC



denoised IC

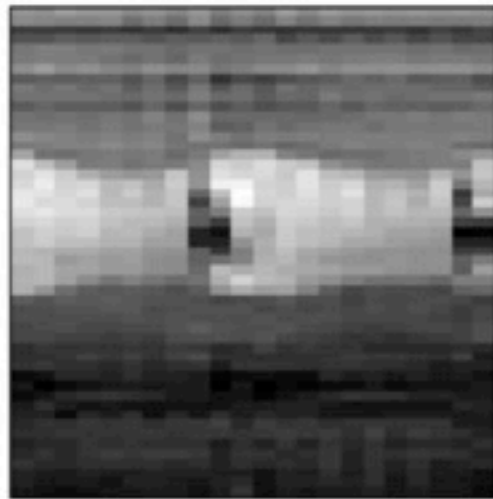


$$J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$

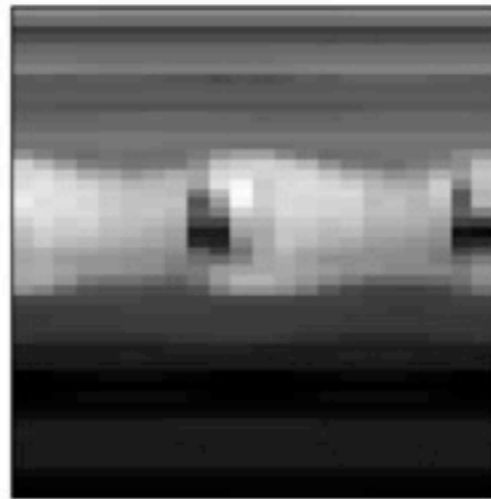


Denoising evaluation

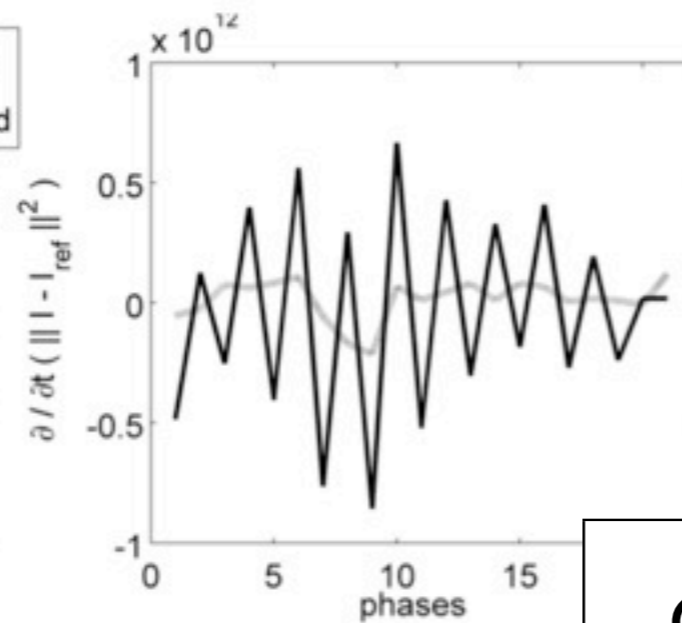
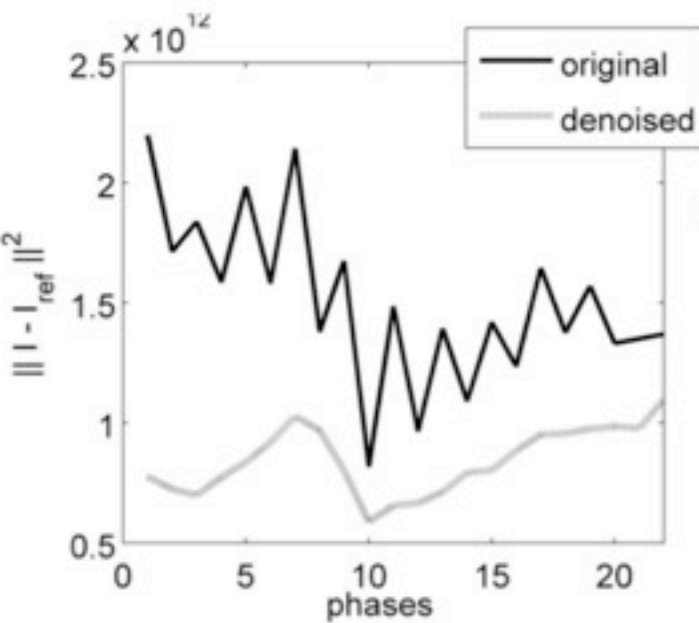
original IC



denoised IC



$$J(s, A, \omega, \phi) = \sum_n \|f_n - A \sin(\omega n + \phi) - s_n\|_2^2 + \lambda \sum_n \|\hat{s}_n\|_1$$



Noise reduction (n=8):

E

Noise reduction

	E	Noise reduction
Original IC	$2.11 \pm 2.88 * 10^{24}$	-
Soft threshold only	$4.37 \pm 6.12 * 10^{23}$	-6.0 ± 2.2 dB
Complete model	$1.24 \pm 1.58 * 10^{23}$	-10.4 ± 3.8 dB

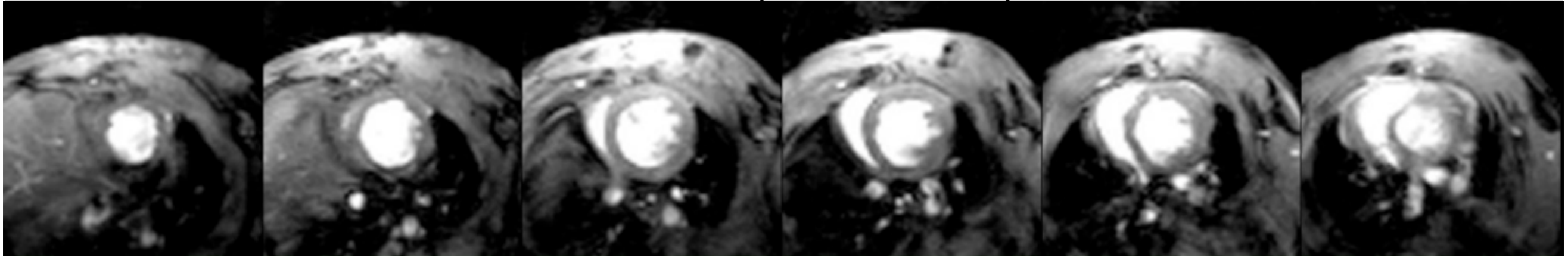
Final result

Basic cine (TR=13.5 ms)

Original images

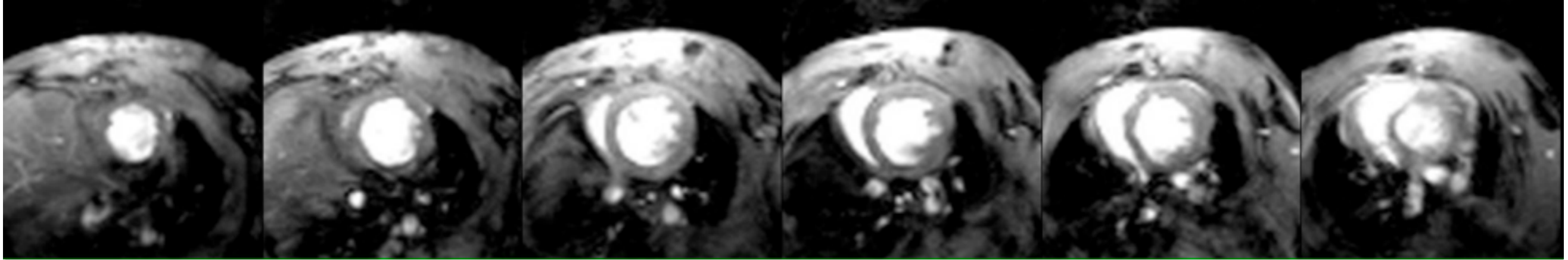
Final result

Basic cine (TR=13.5 ms)

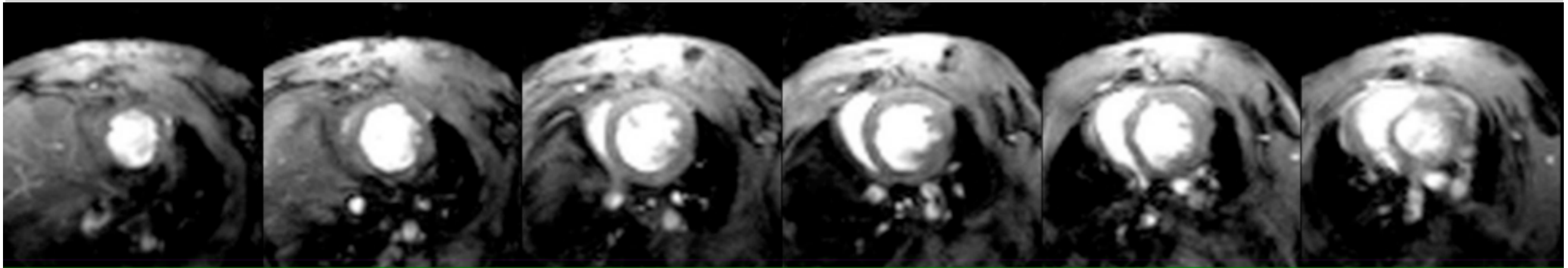


Final result

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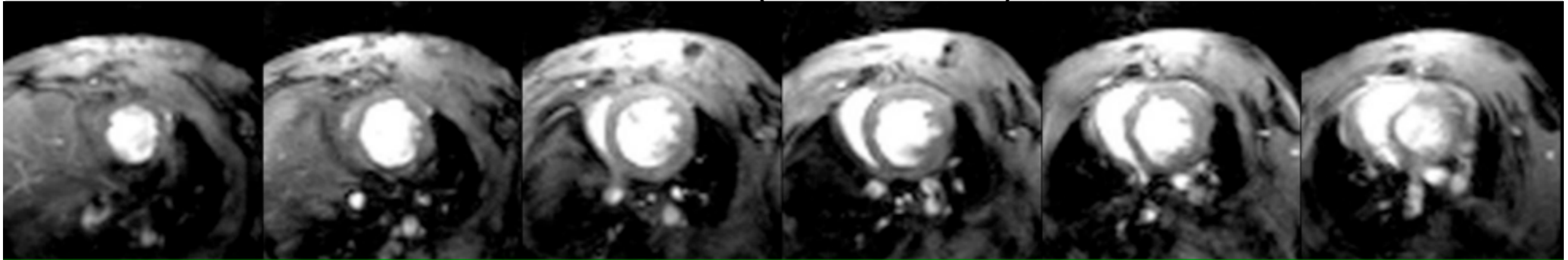


Interleaved cine (TR=6.8 ms)

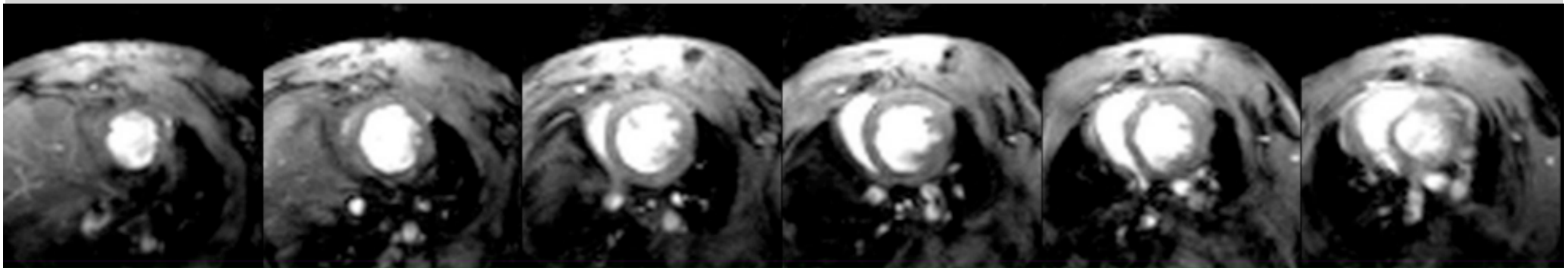


Final result

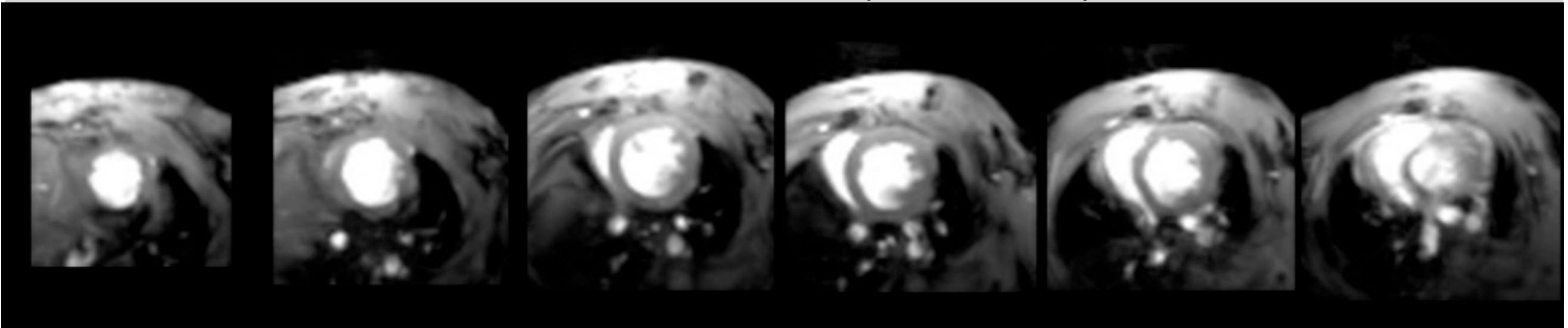
Basic cine (TR=13.5 ms)



Interleaved cine (TR=6.8 ms)

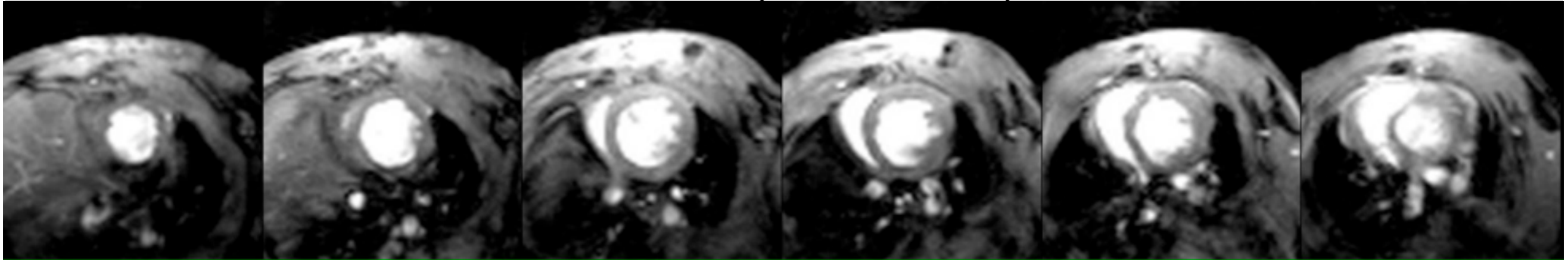


After enhancement (TR=6.8 ms)

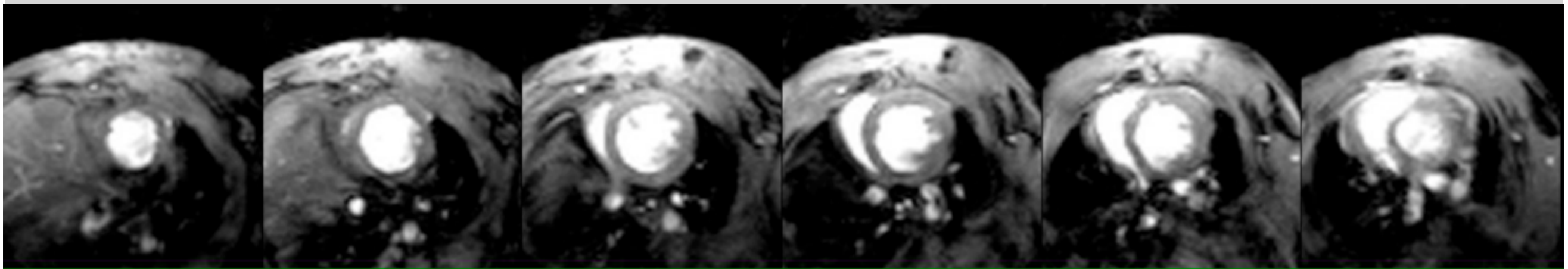


Final result

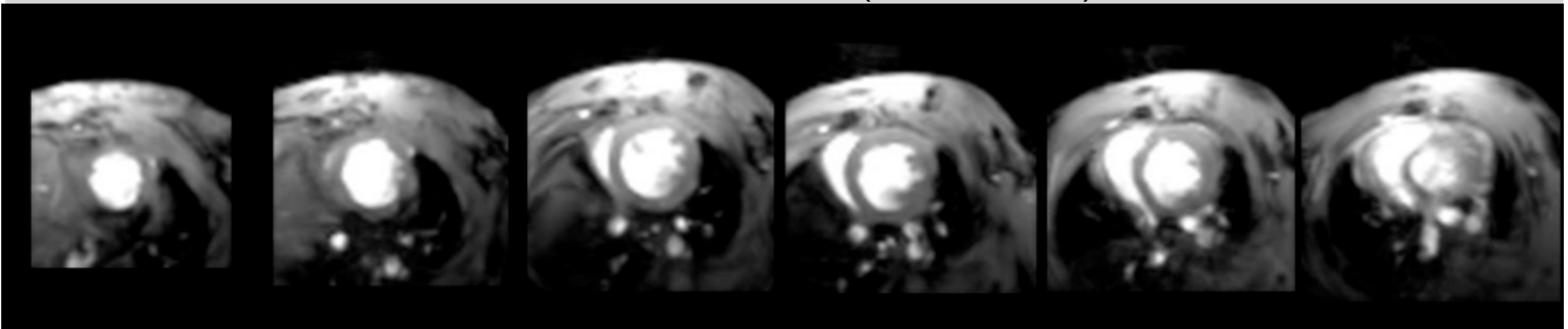
Basic cine (TR=13.5 ms)



Interleaved cine (TR=6.8 ms)



After enhancement (TR=6.8 ms)



Tool to achieve sufficient temporal resolution for mice imaging (< 9 ms)

Applications in cardiac research

Multi-slices short axis cine to evaluate:

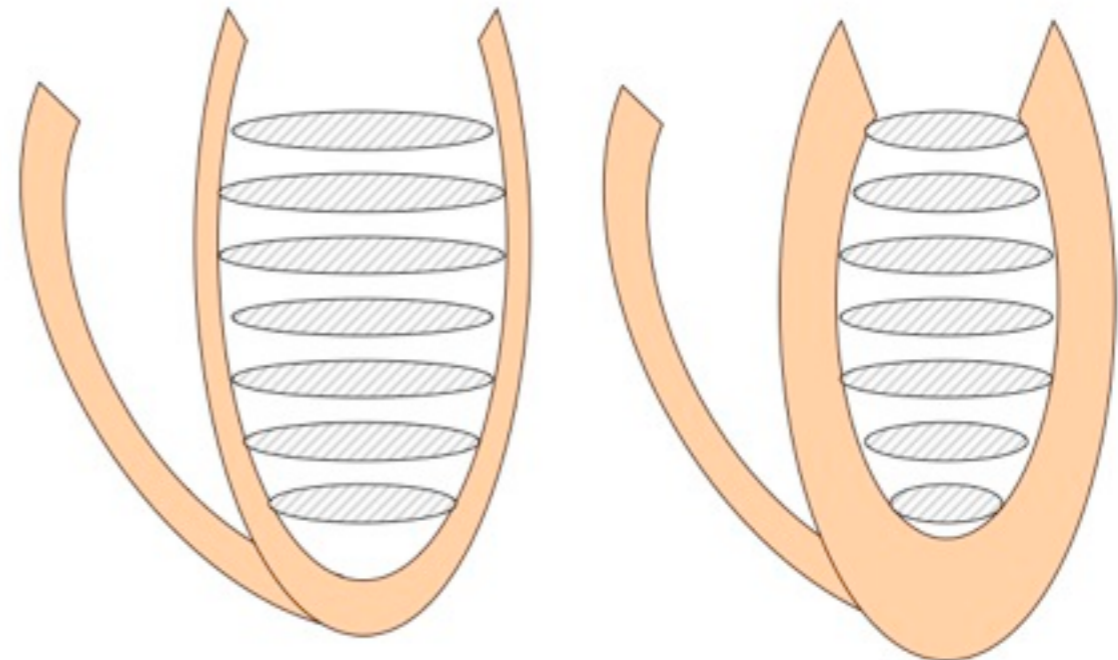
Heart mass



$$(LV + RV)_{mass} = \gamma \cdot \text{Slice thickness} \cdot \sum_{\text{all slices}} [\text{epi area} - (\text{LV endo area} + \text{RV endo area})]$$

γ is the specific gravity of the myocardium, $\gamma = 1.055 \text{ g/cm}^3$

Cardiac function



Diastole

Systole

End-diastolic volume
EDV

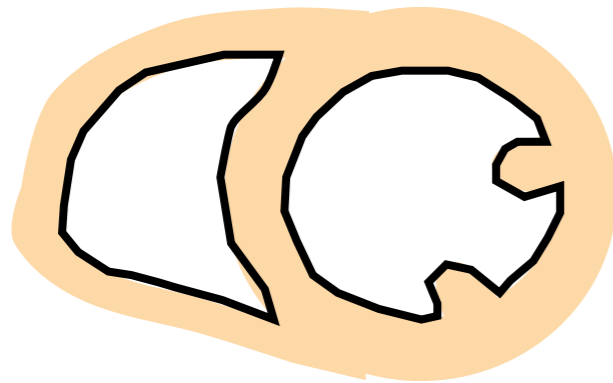
End-systolic volume
ESV

$$EF = \frac{EDV - ESV}{EDV}$$

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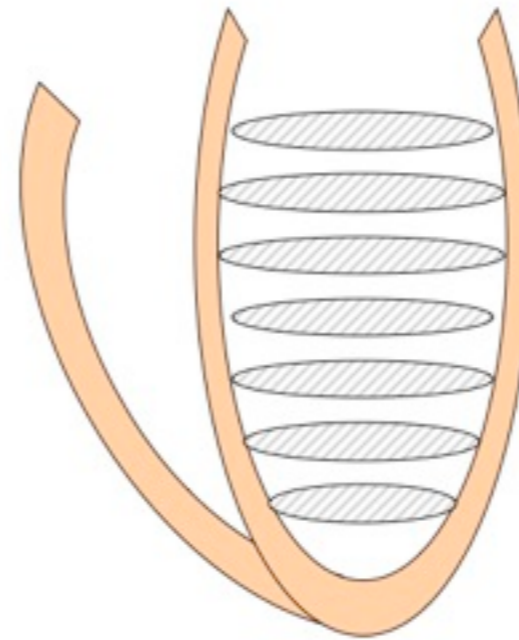
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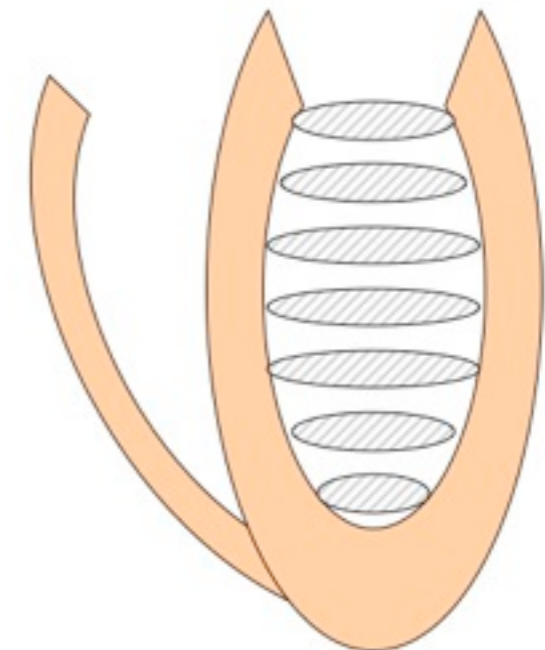
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Cardiac function



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End-diastolic volume
EDV



Systole

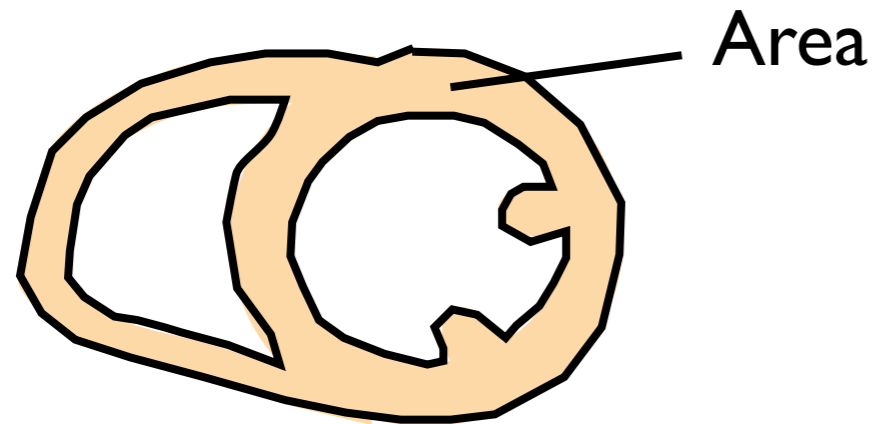
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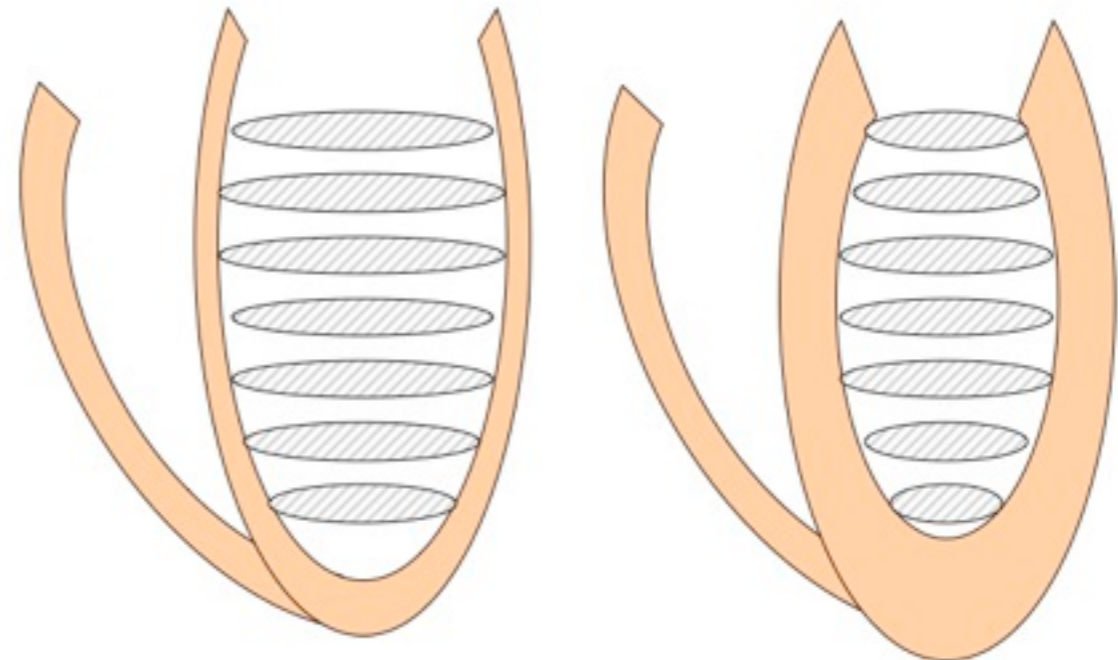
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Diastole

Systole

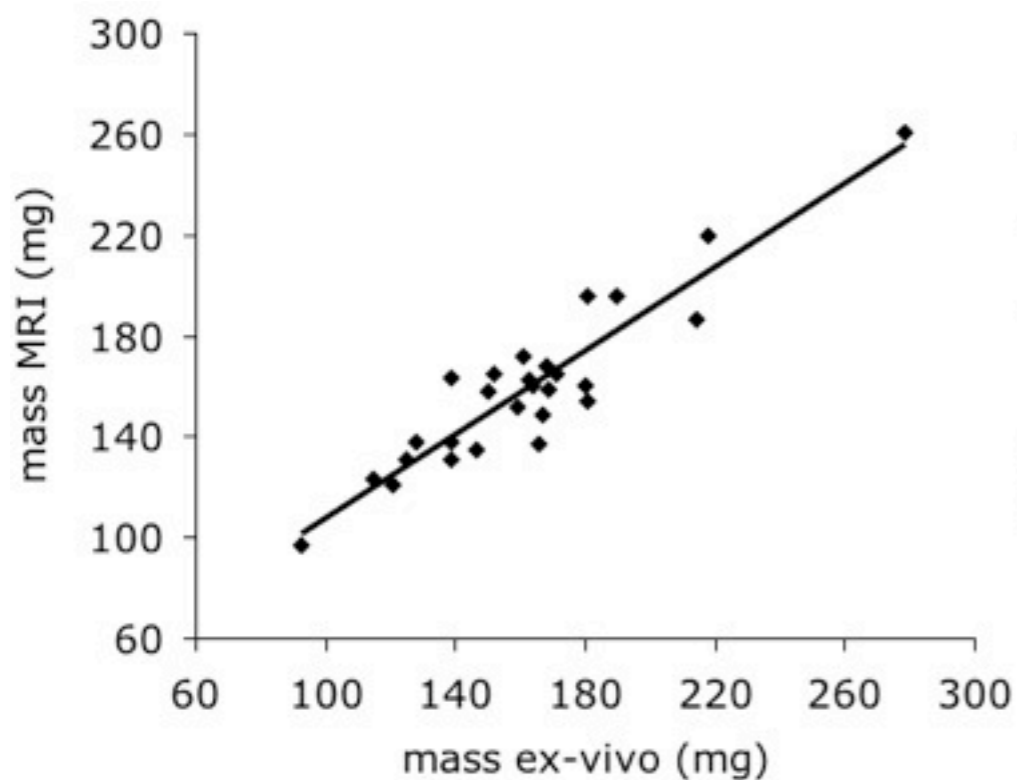
End-diastolic volume
EDV

End-systolic volume
ESV

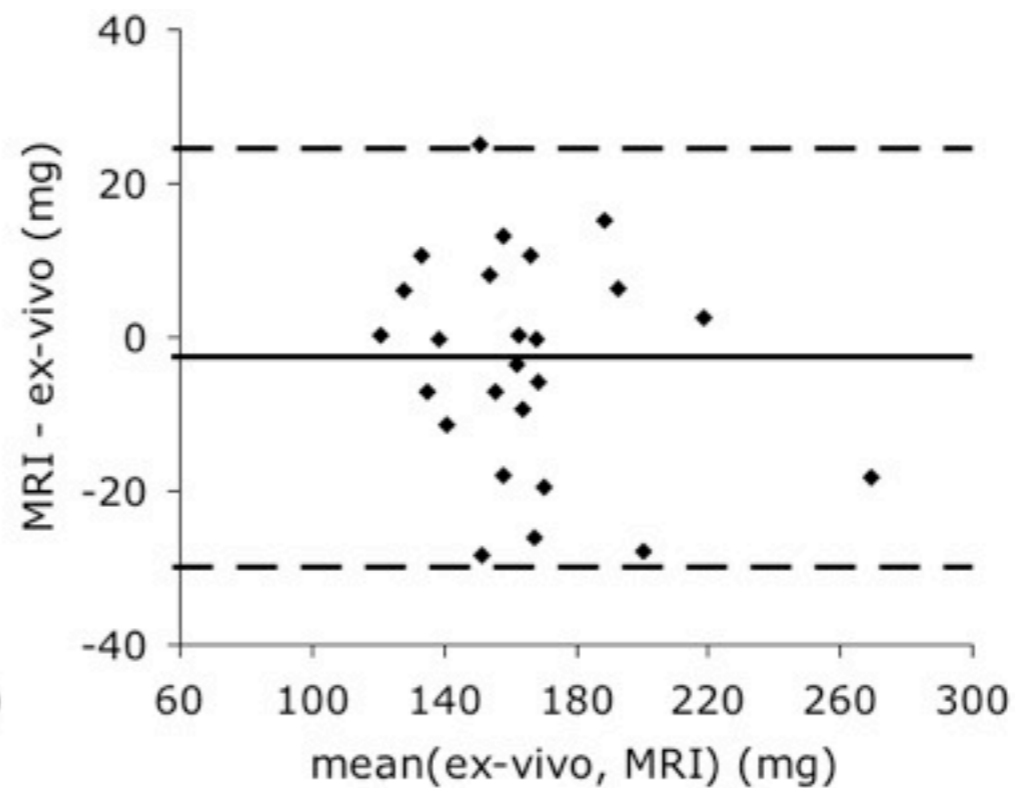
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Diabetes - Cardiac hypertrophy

- Transgenic mice developing cardiac hypertrophy (n=23)
- Wild type mice (n=4)



$$y = 0.829x + 25.1, R^2 = 0.89 \text{ (n=27)}$$



$$\text{bias} = 2.62 \text{ mg}$$
$$1.96 \text{ SD} = 27.15 \text{ mg}$$

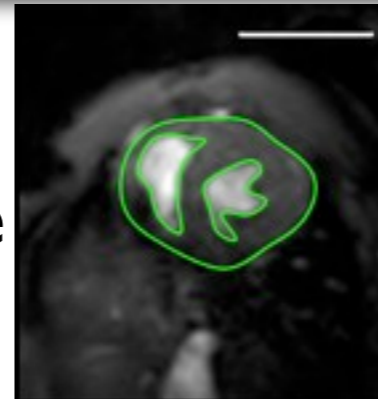
Yang et al.¹
obtained
1.96 SD = 15 mg
on a dedicated
scanner (4.7 T)

¹ Yang, Circulation 109(9):1161, 2004

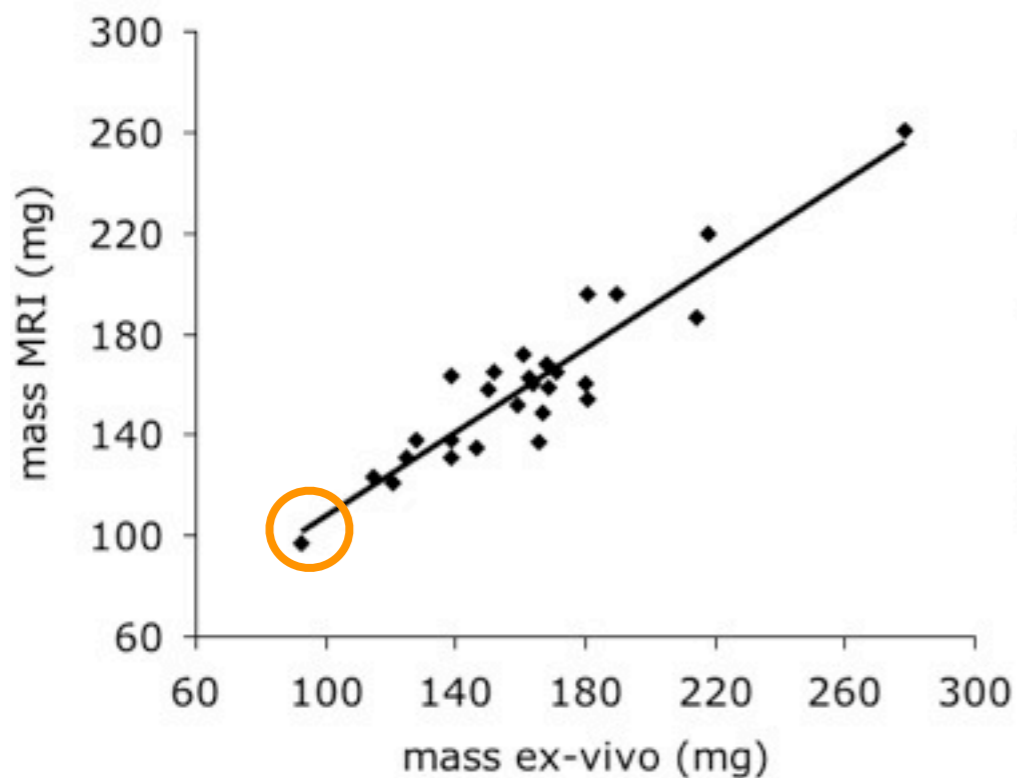
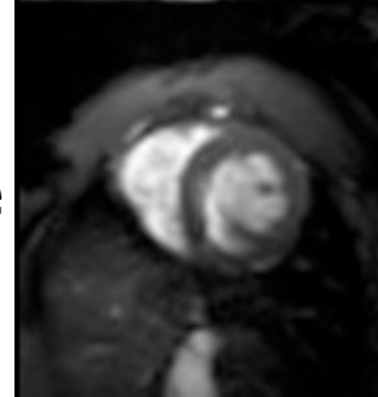
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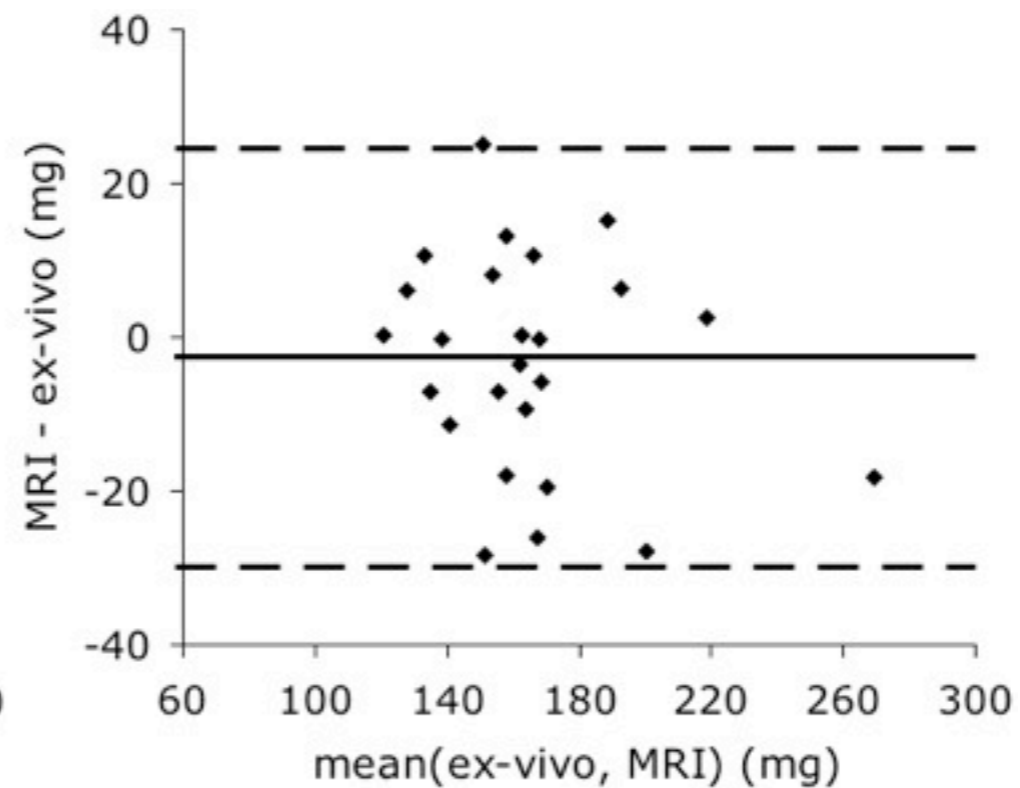
Systole



Diastole



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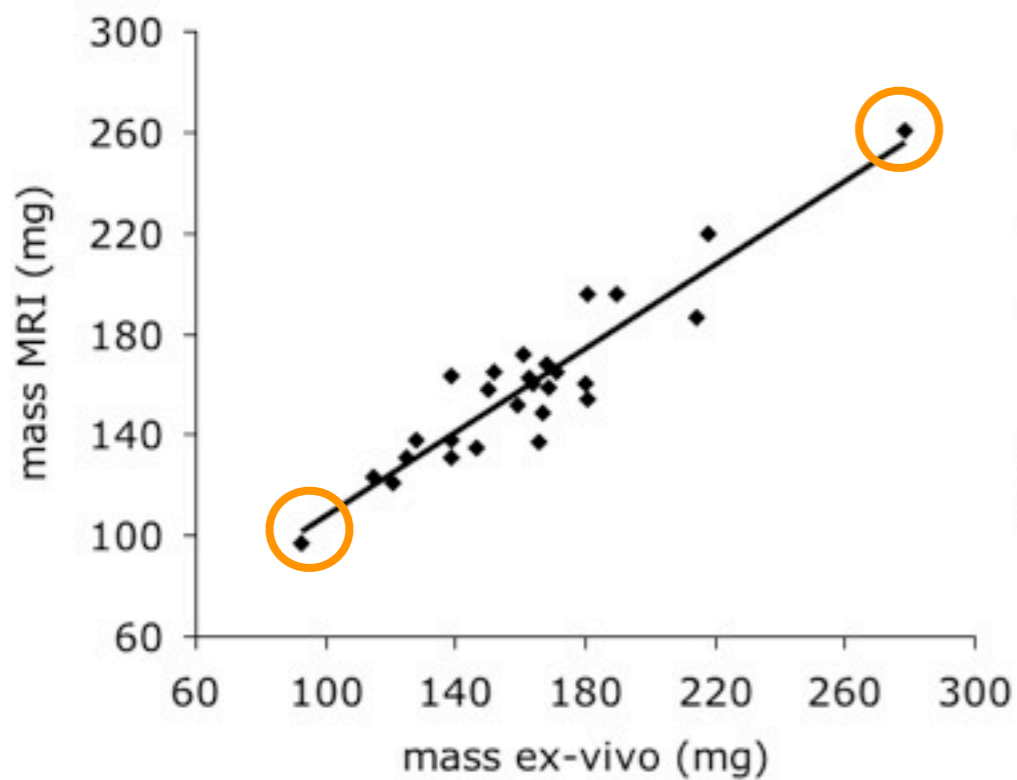
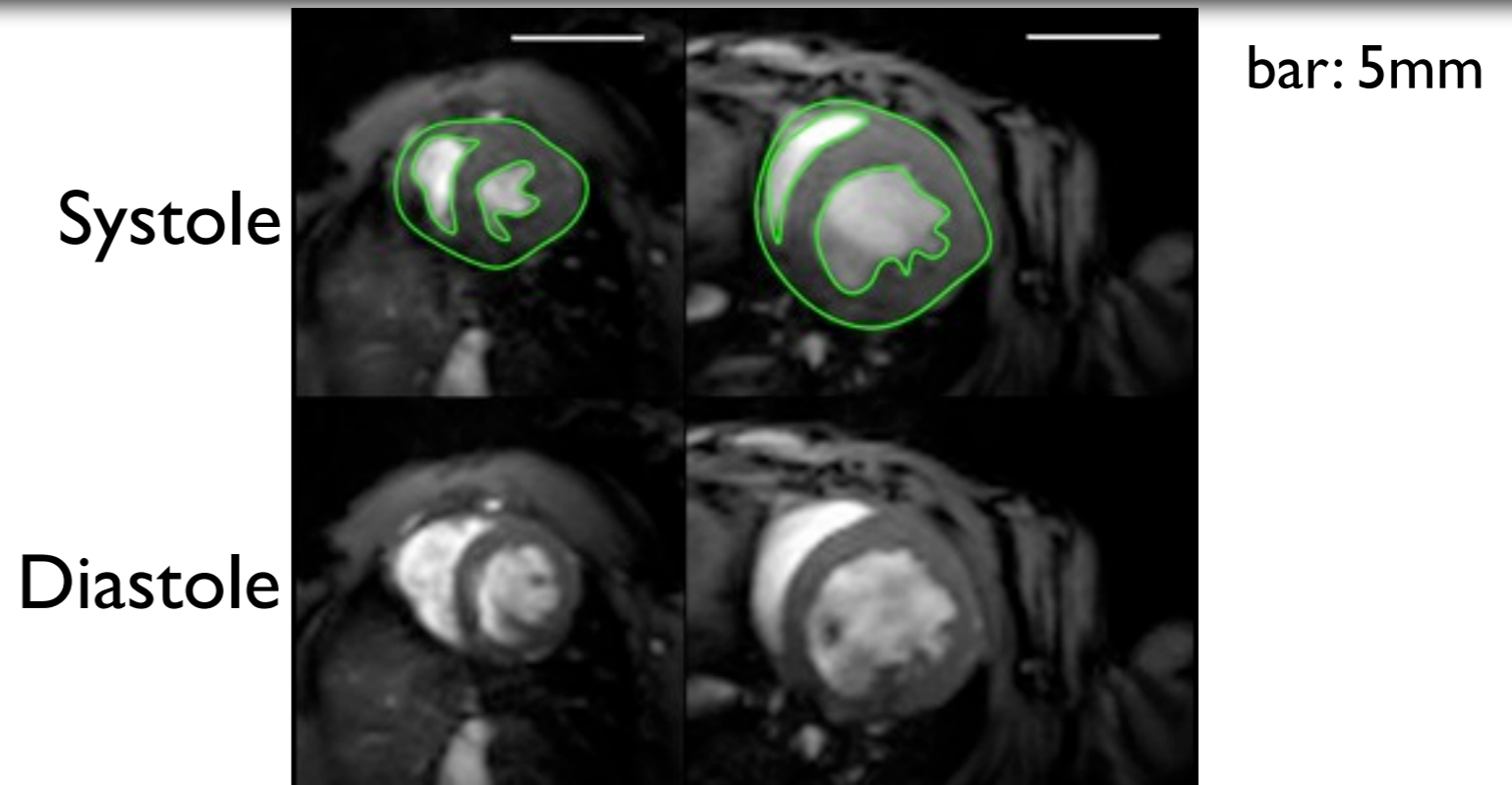
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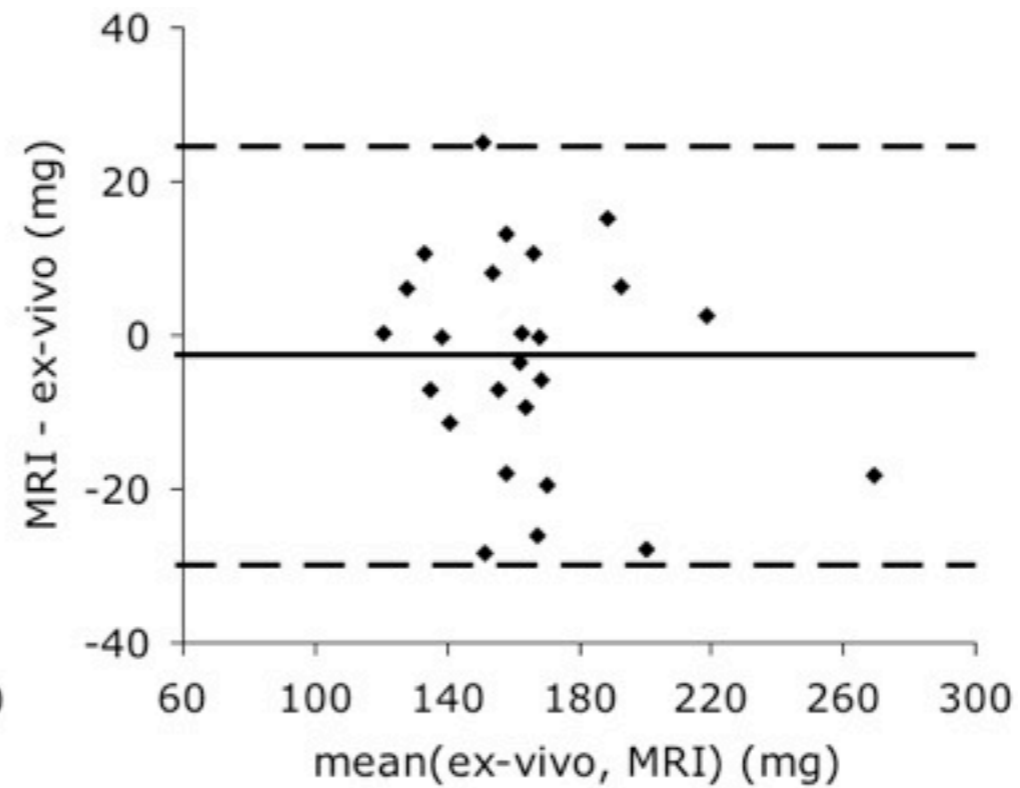
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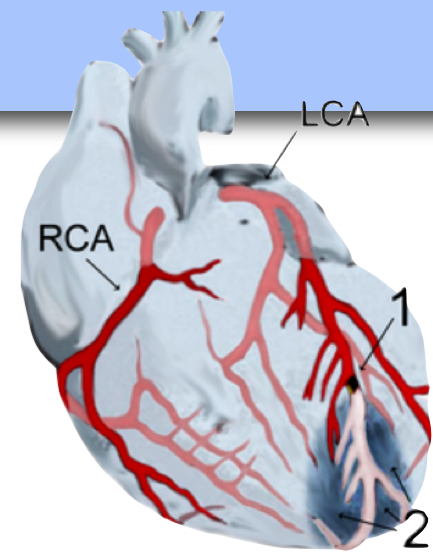
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Infarction - Function impairment

WT: Control mice C57/BL6 WT n=4

MI: C57/BL6 with complete ligation of LAD, 24h after surgery (n=3)

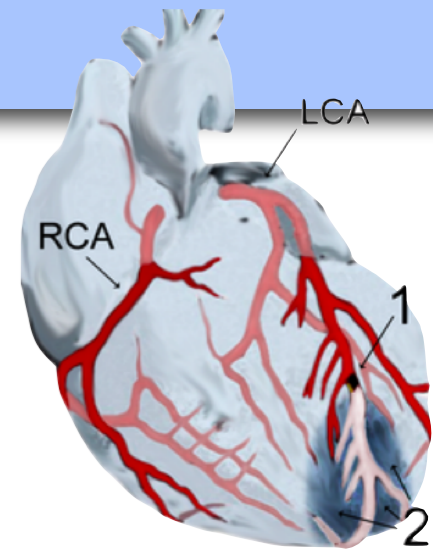
Evaluate effect of drug on infarct size and heart failure



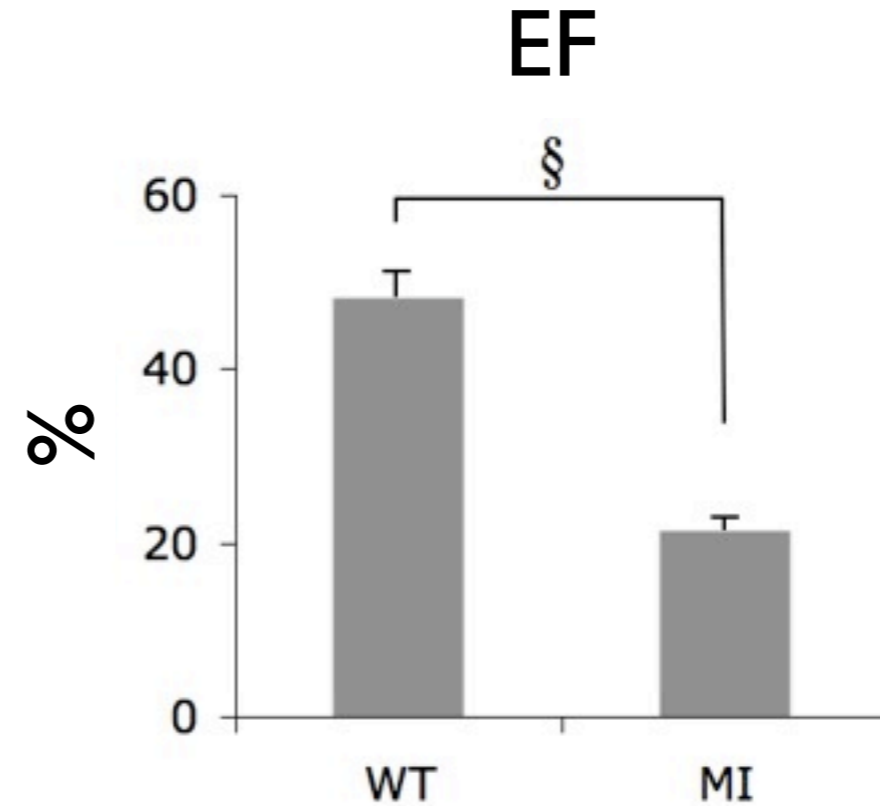
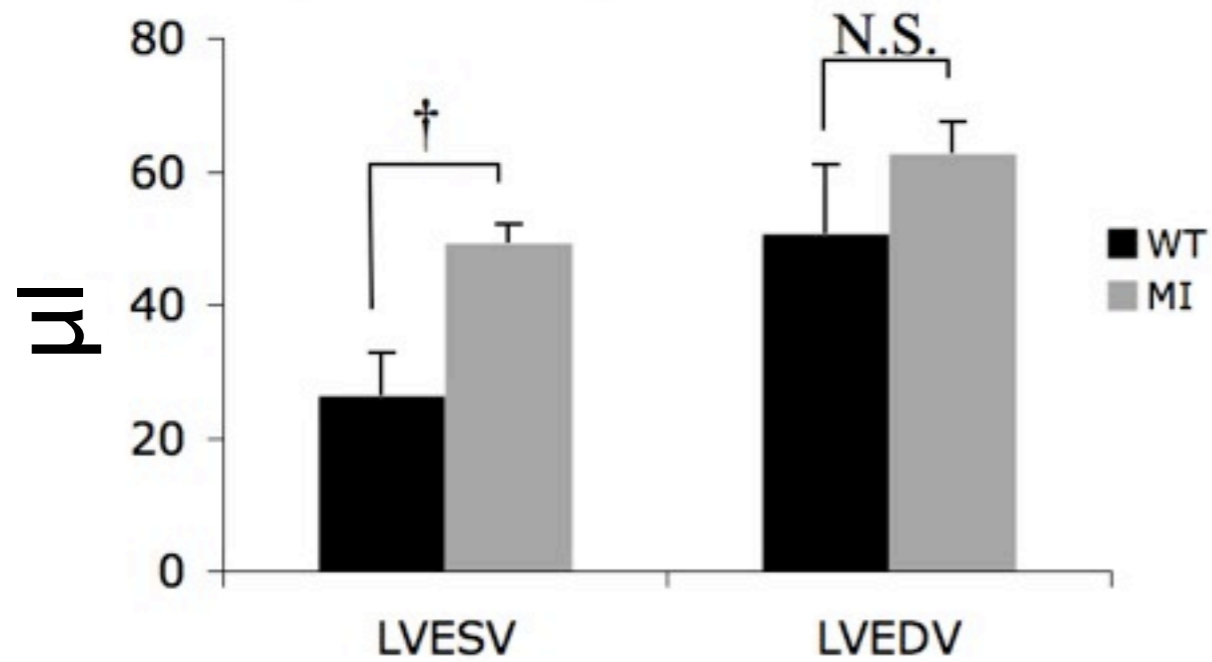
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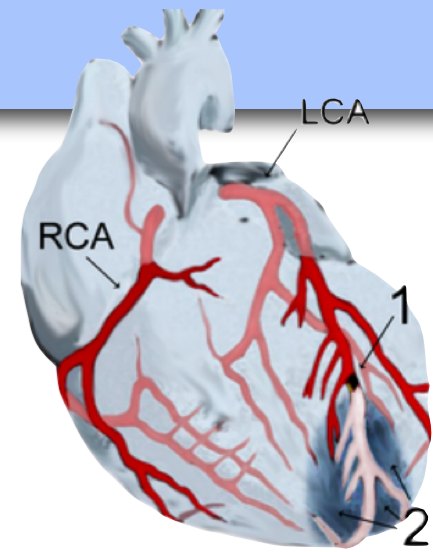


Basic cine (TR=13.5ms)



* p<0.05,
§ p<0.01,
† p<0.001

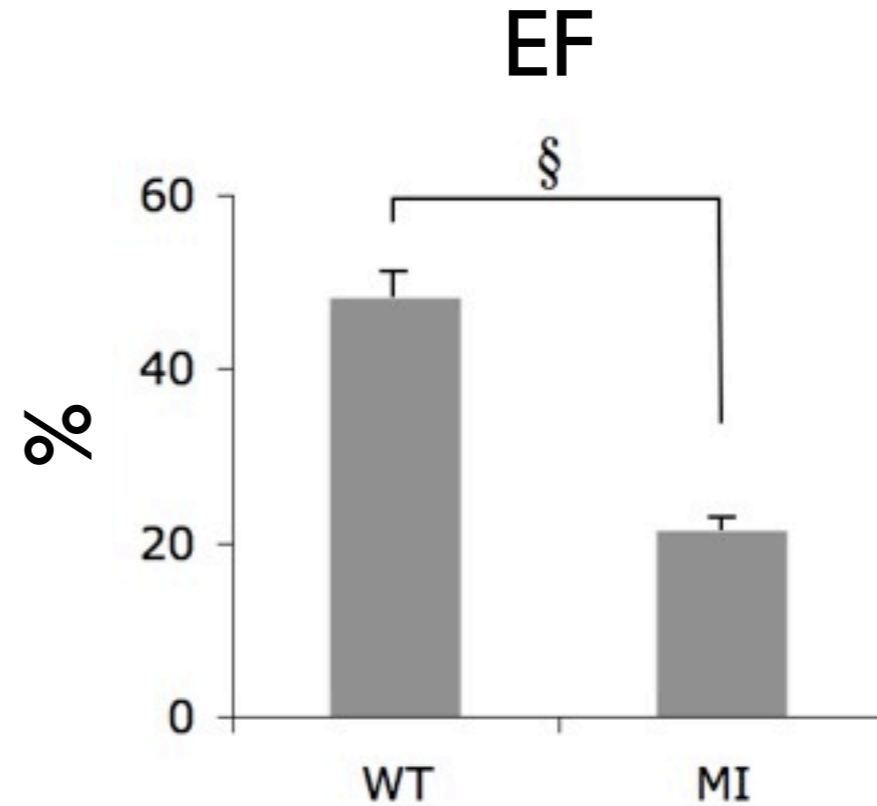
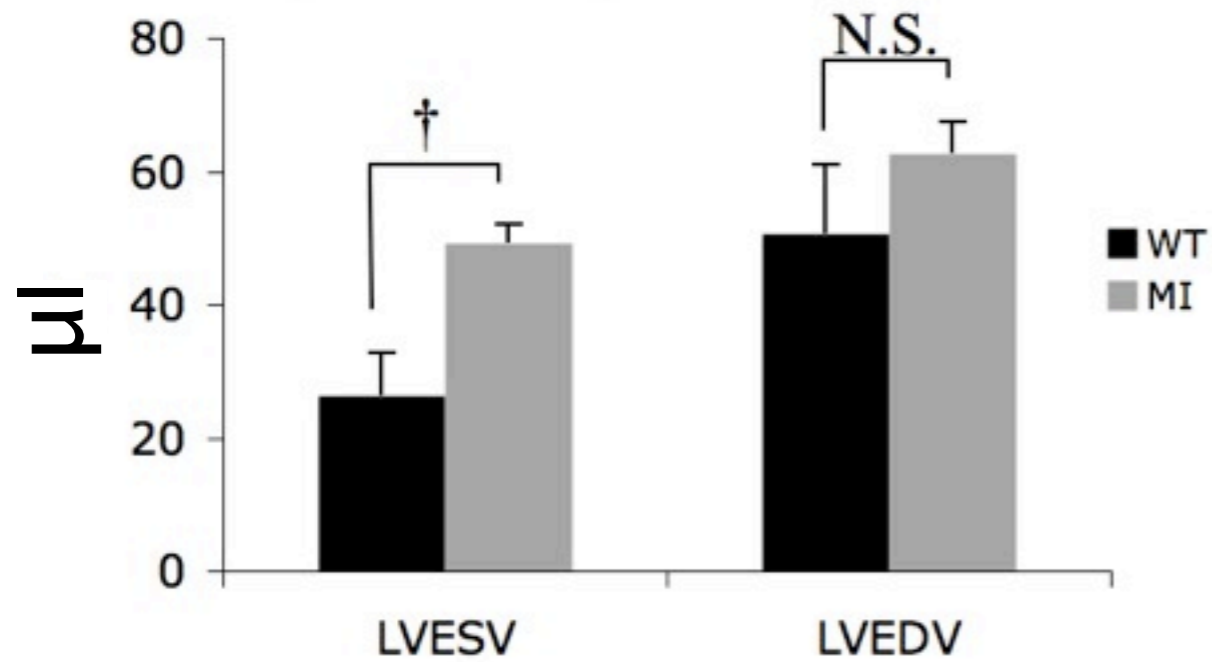
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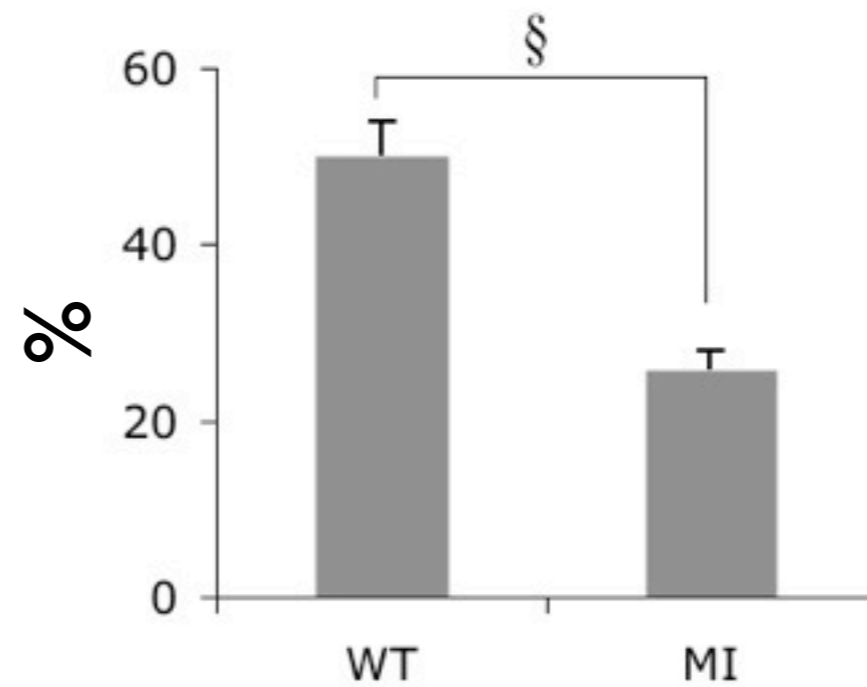
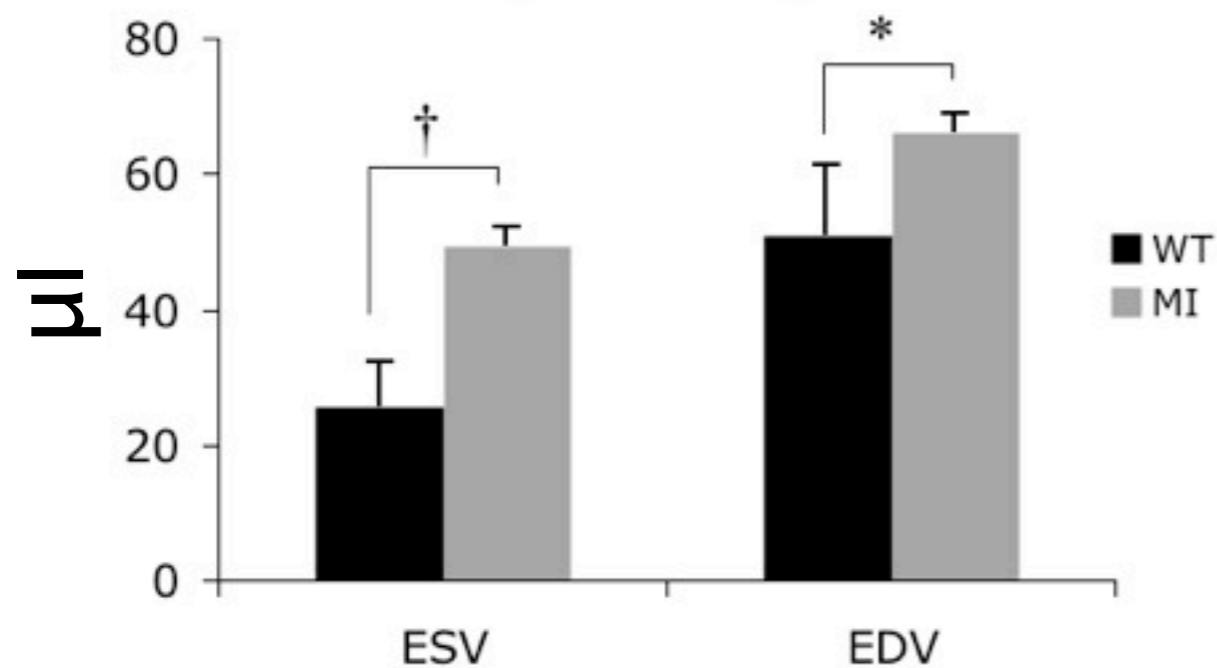
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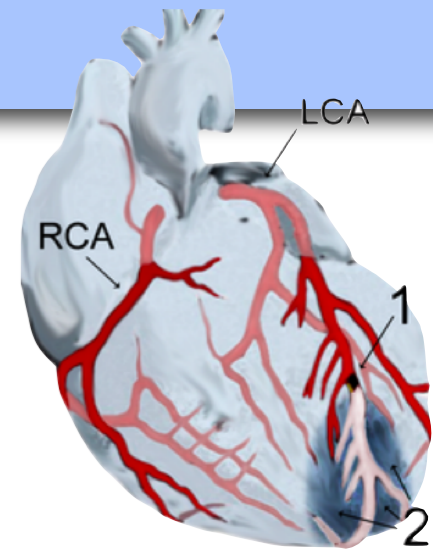


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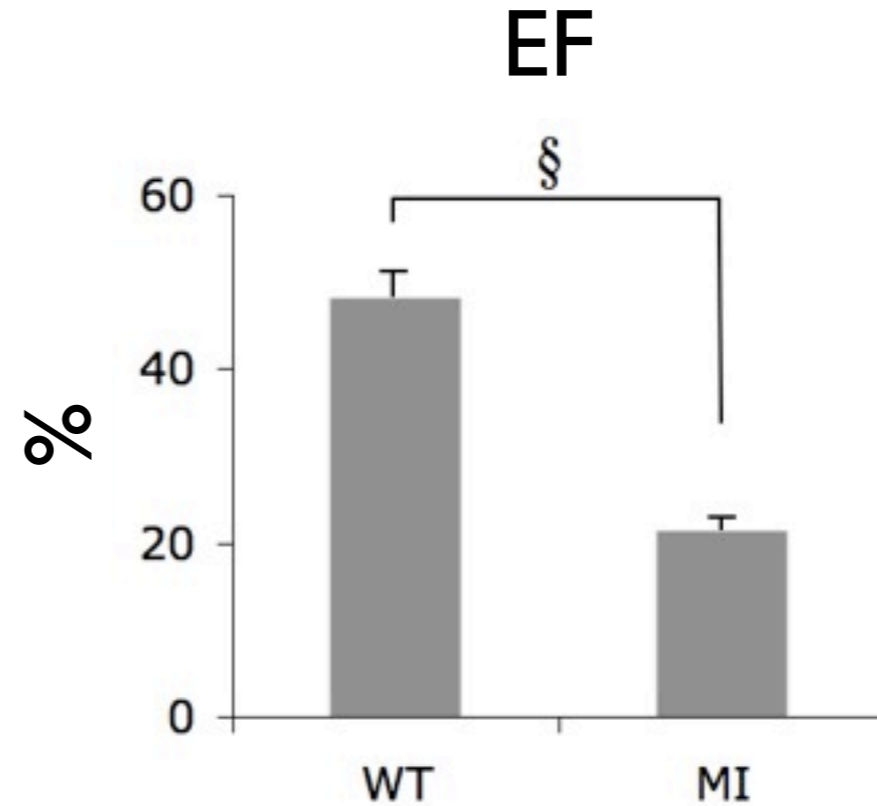
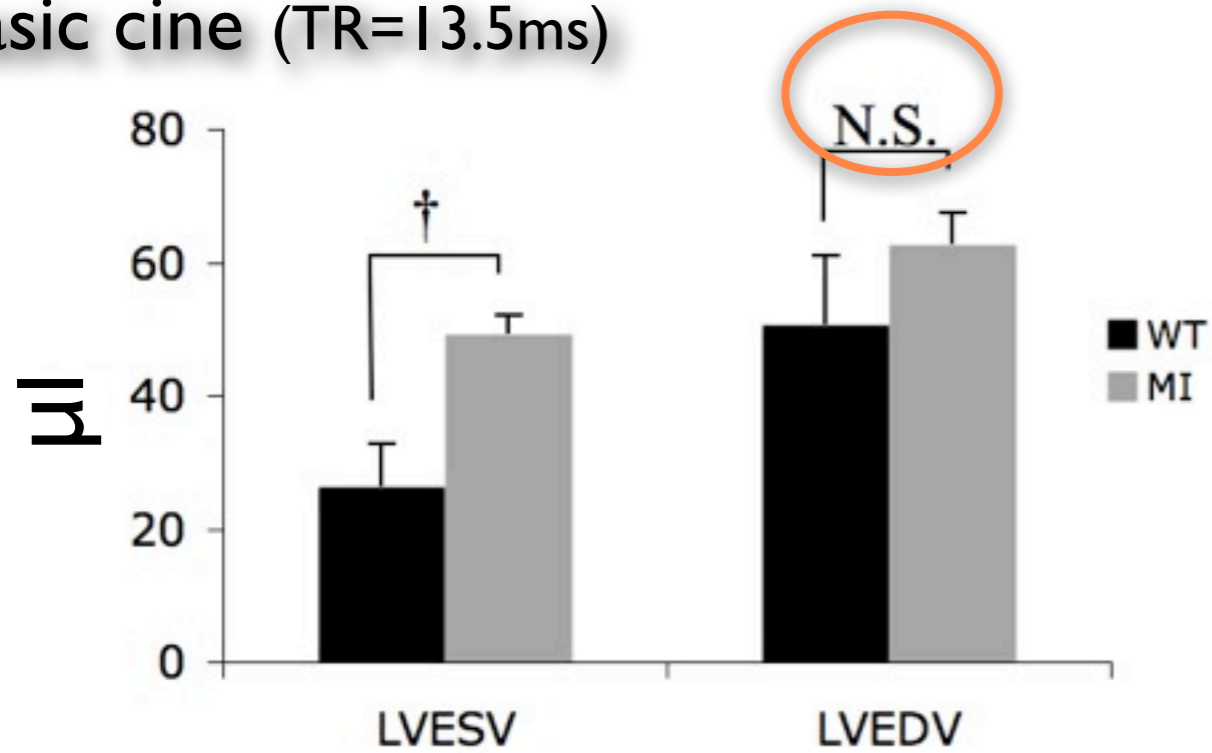
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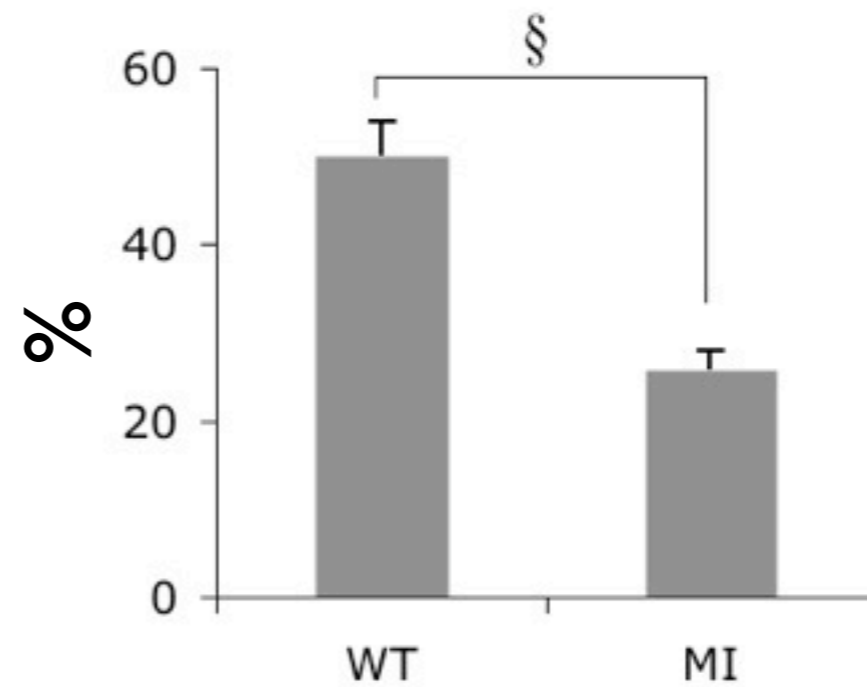
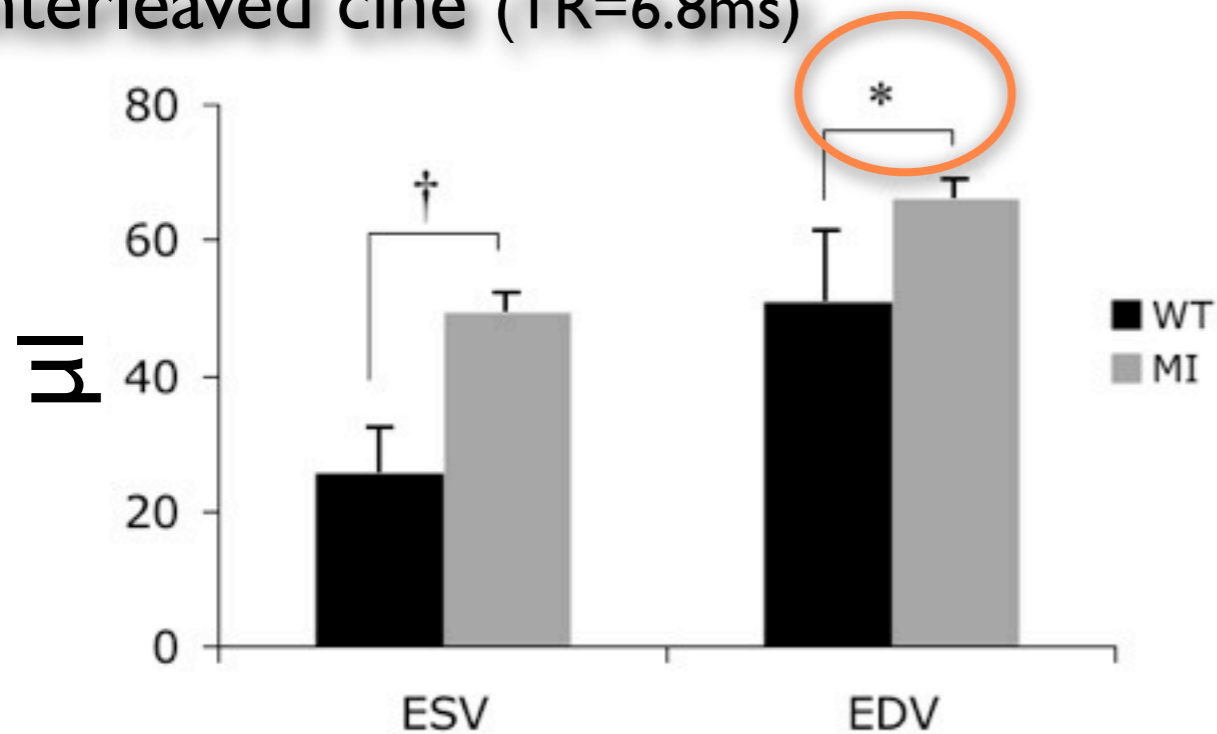
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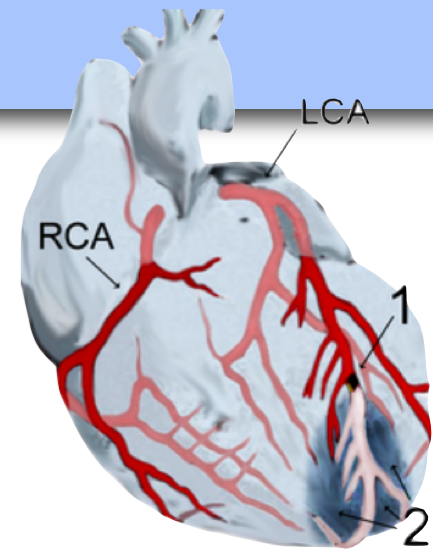


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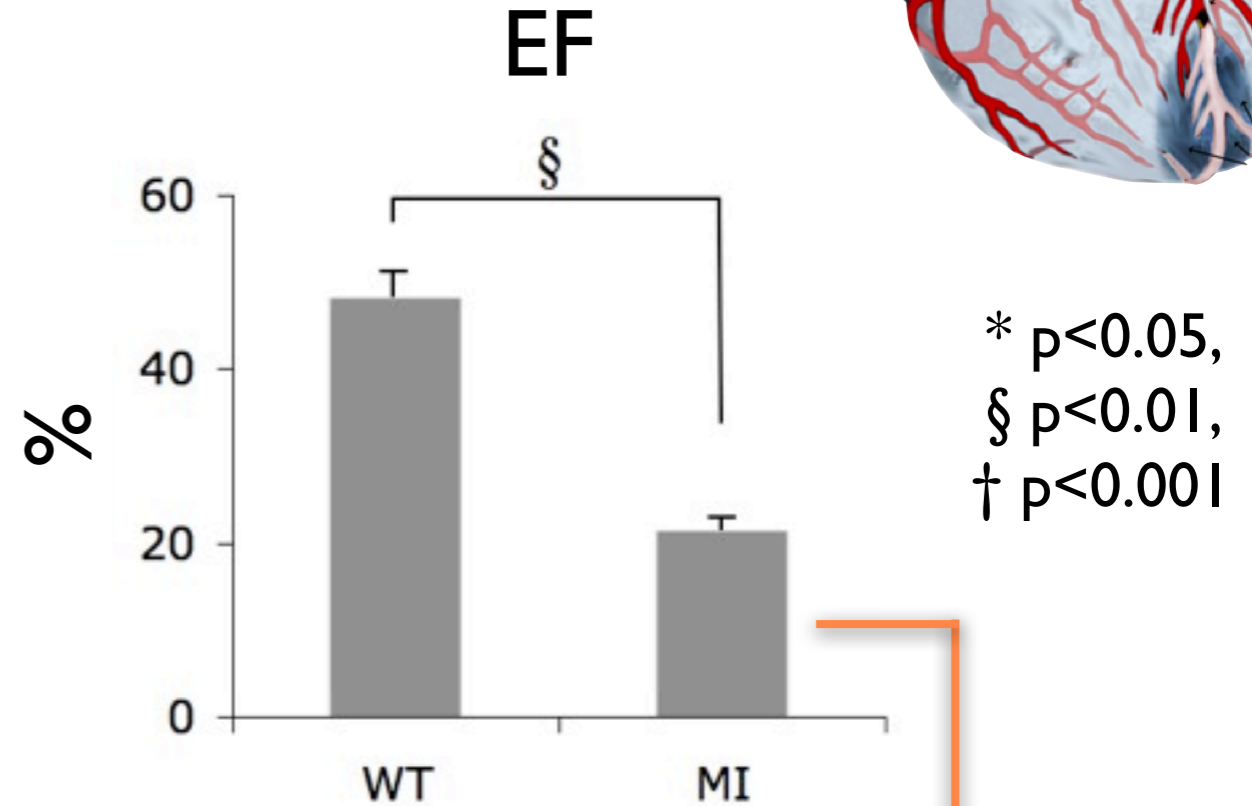
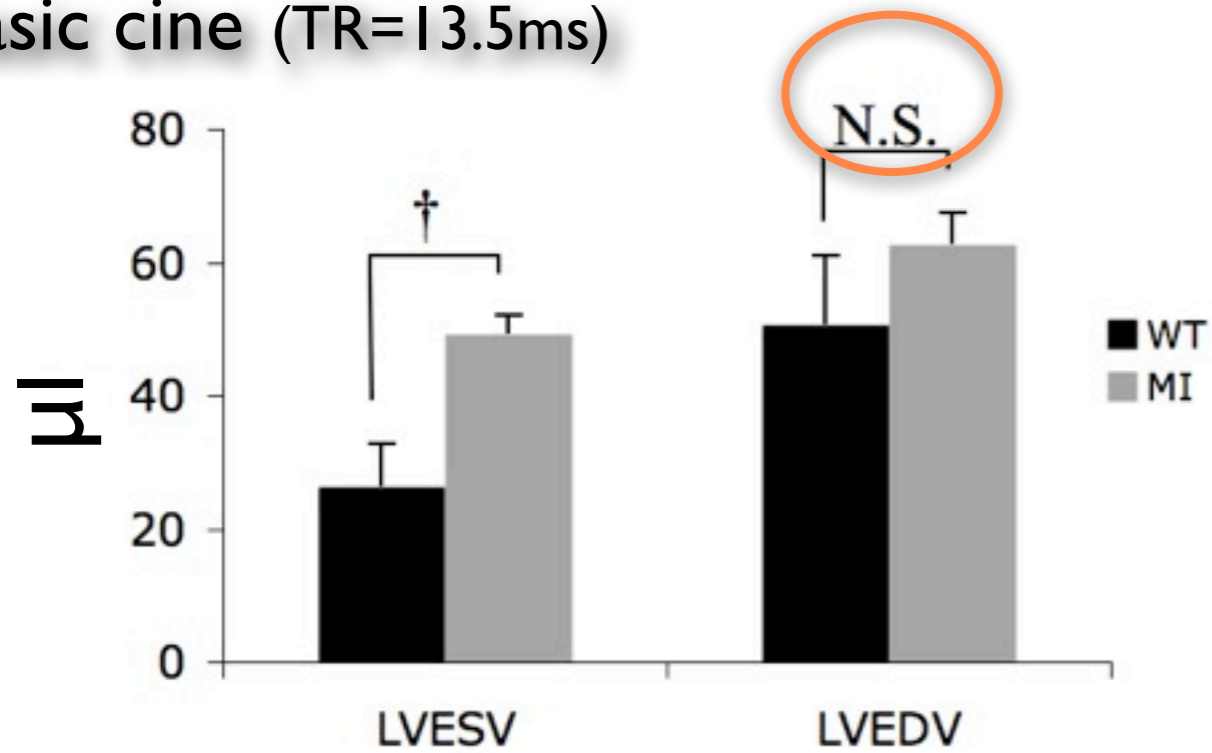
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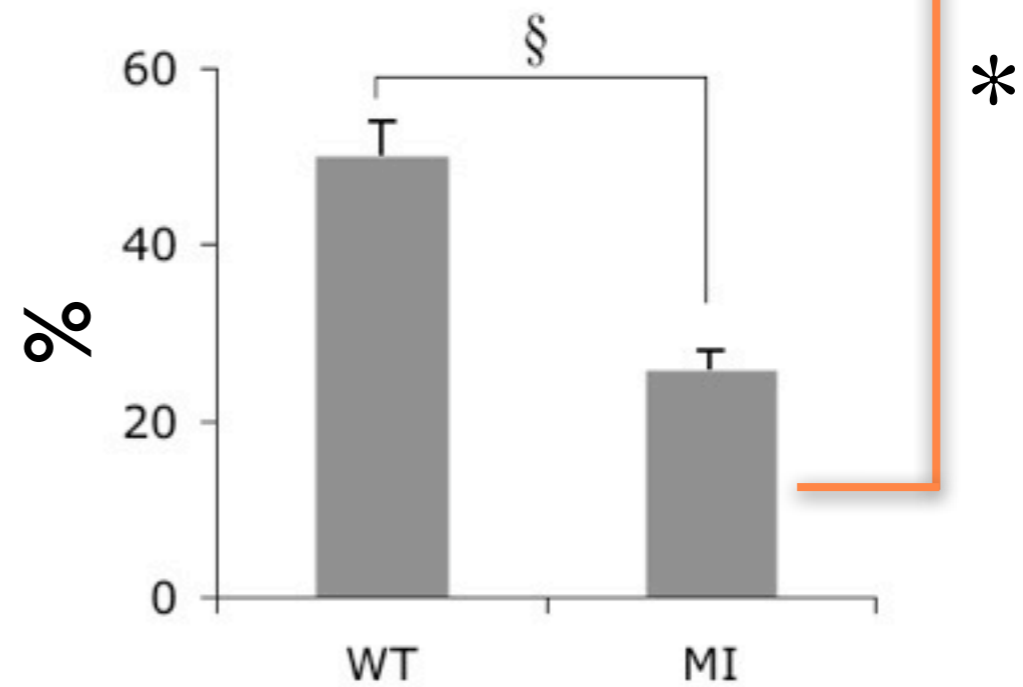
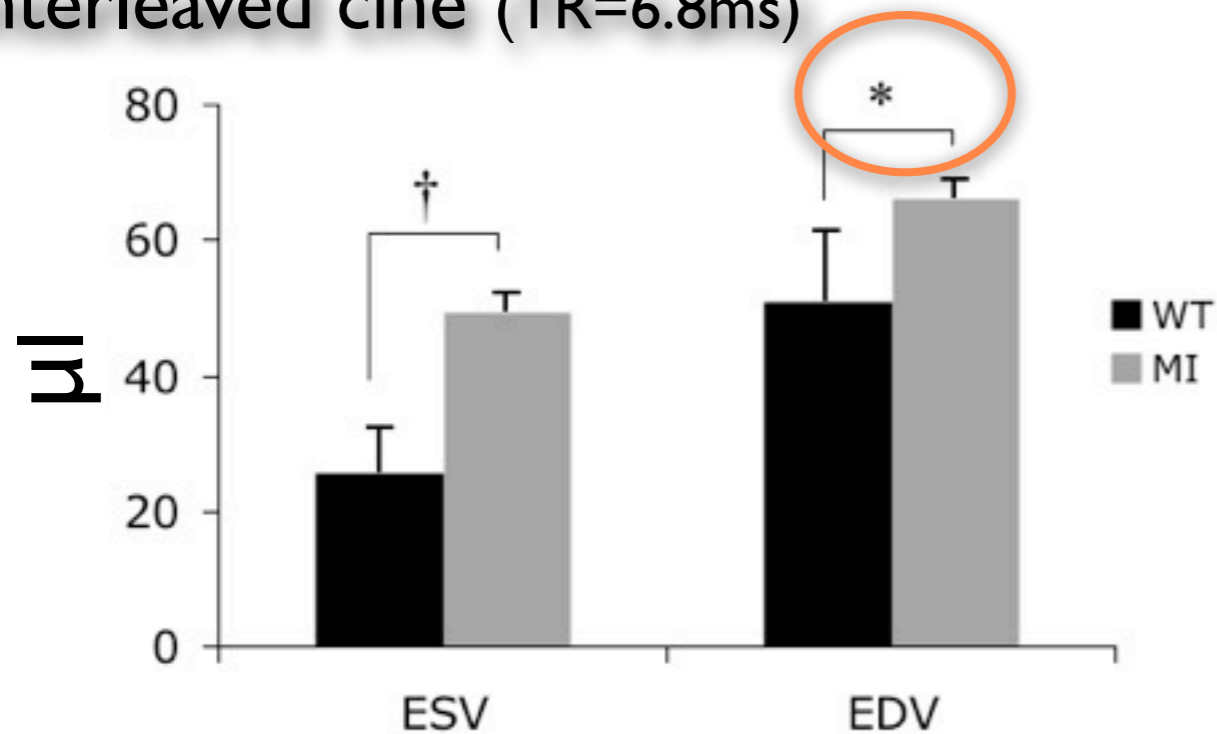
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Interleaved cine (TR=6.8ms)



Conclusions

- Interleaved cine
 - Same total acquisition time as basic cine
 - Data sampling at different time points (no sliding window)
- Denoising algorithm and temporal regularization
 - Fast and non-iterative
 - Works in image domain, still possible to use parallel imaging (SENSE or GRAPPA)
- Sequence available for cardiac assessment in mice
 - Already used in 2 large studies (diabetes, infarction)

Acknowledgments



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Dr. Jean-Noël Hyacinthe



Service of Cardiology

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