



# Omics and AI in breast cancer

Omics driven radiotherapy approaches

**Elisabet Rodríguez Tomàs**

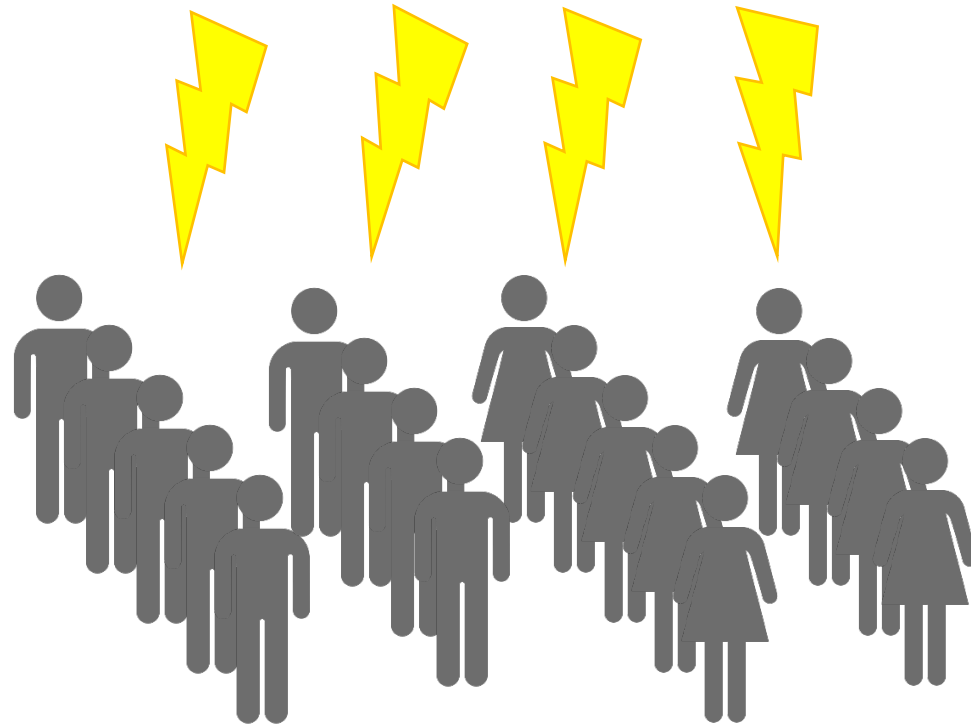
Radiation Oncology Department and Biomedical Research Unit

Universitary Hospital Sant Joan in Reus, Tarragona (Spain)



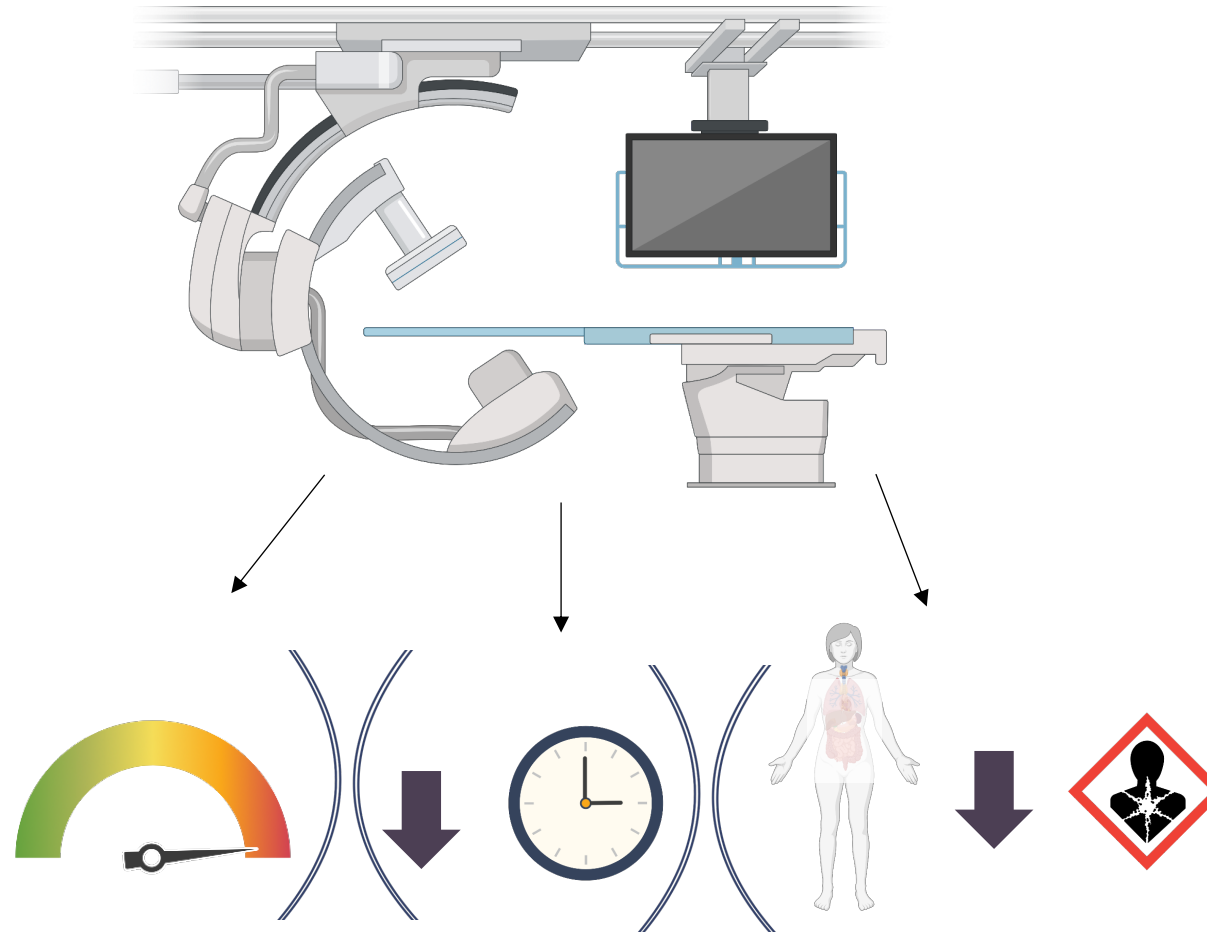
# Clinical considerations

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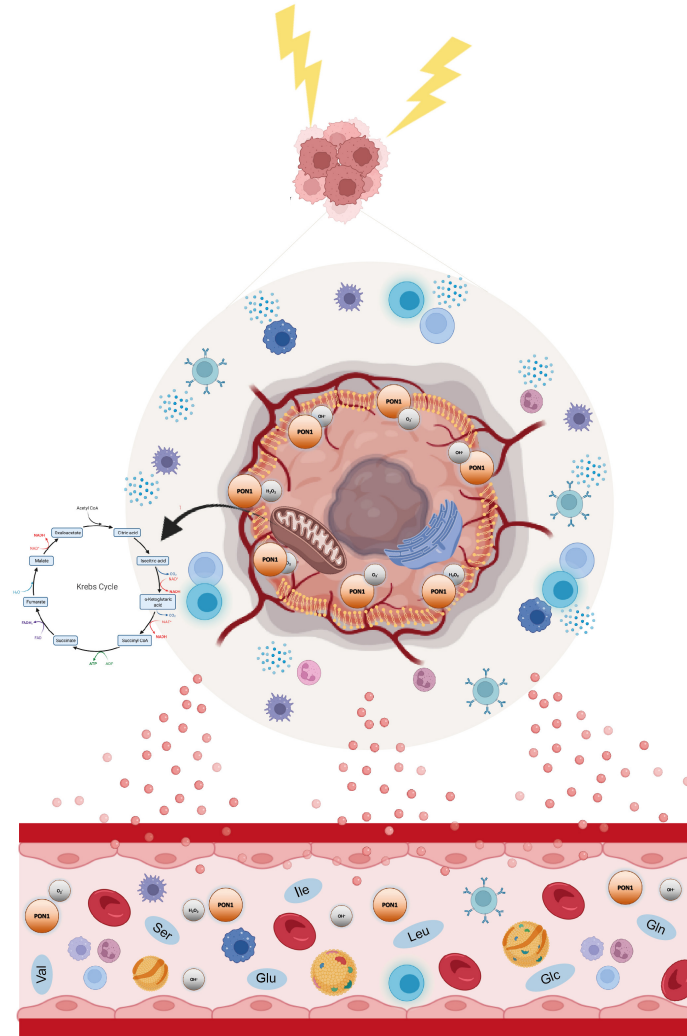


**STANDARD TREATMENT**

# Advances in radiotherapy

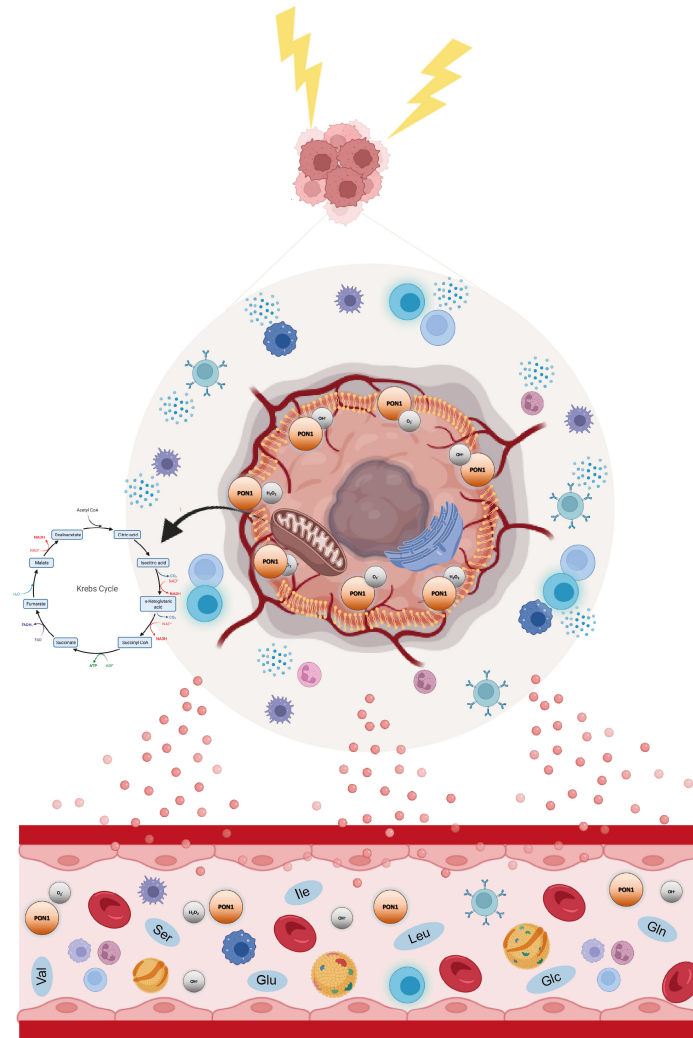


# Radiotherapy gaps



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# Radiotherapy gaps



➔ **BIOMARKER** ➔ **PERSONALIZED THERAPY**

# State of art: The future of omics and big data

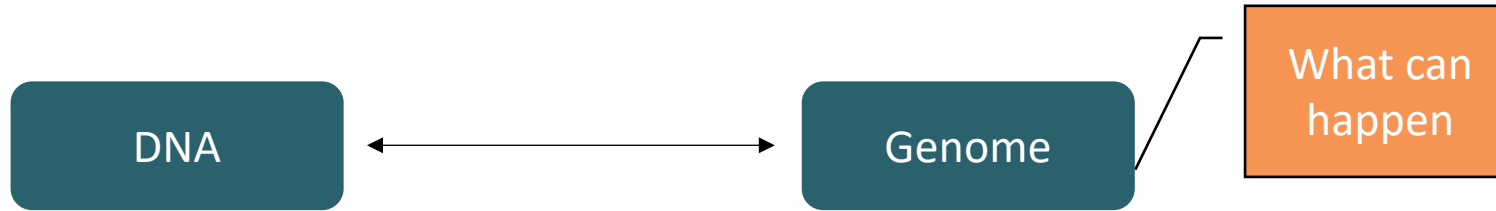
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Molecular techniques help understand how biological entities work

# State of art: The future of omics and big data

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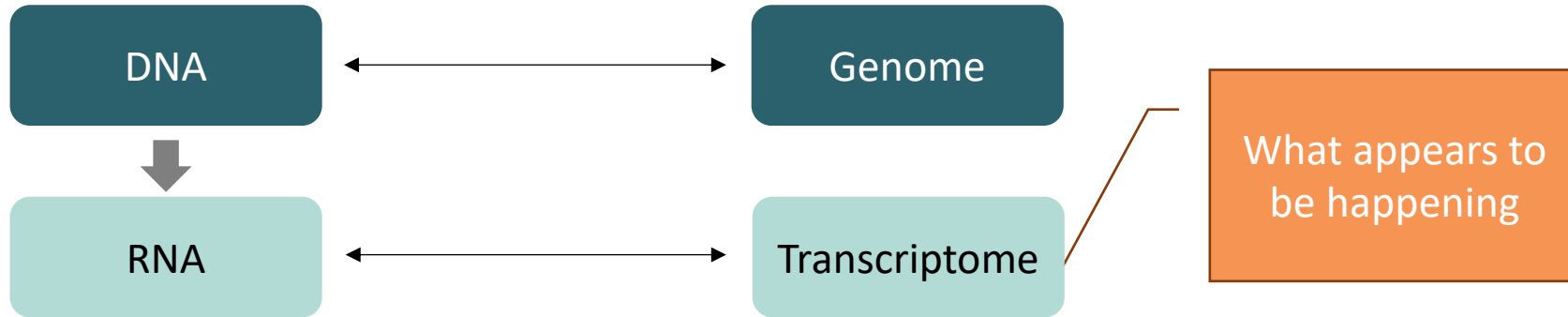
Molecular techniques help understand how biological entities work





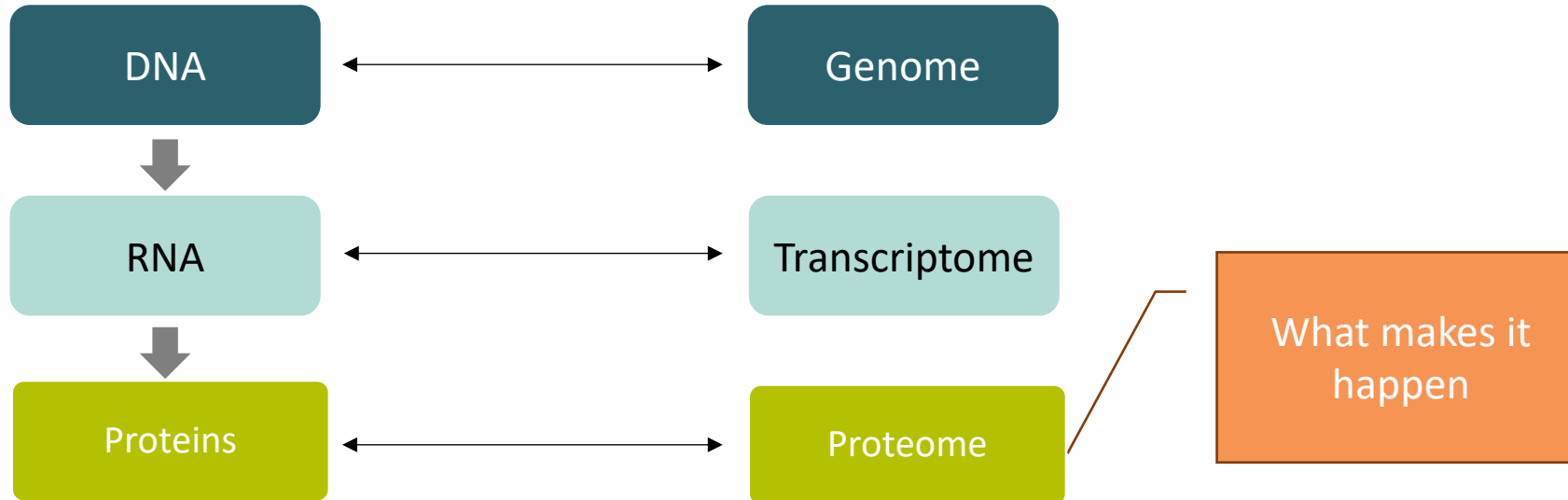
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Molecular techniques help understand how biological entities work



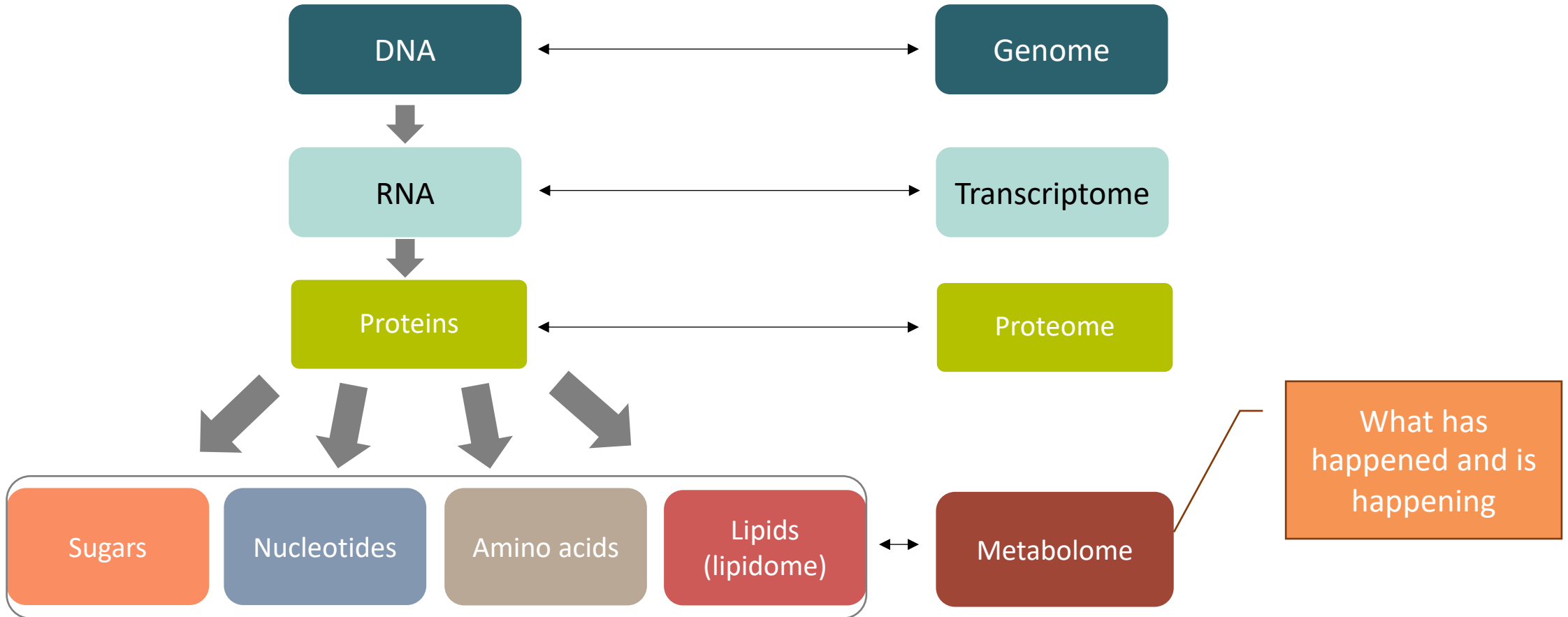
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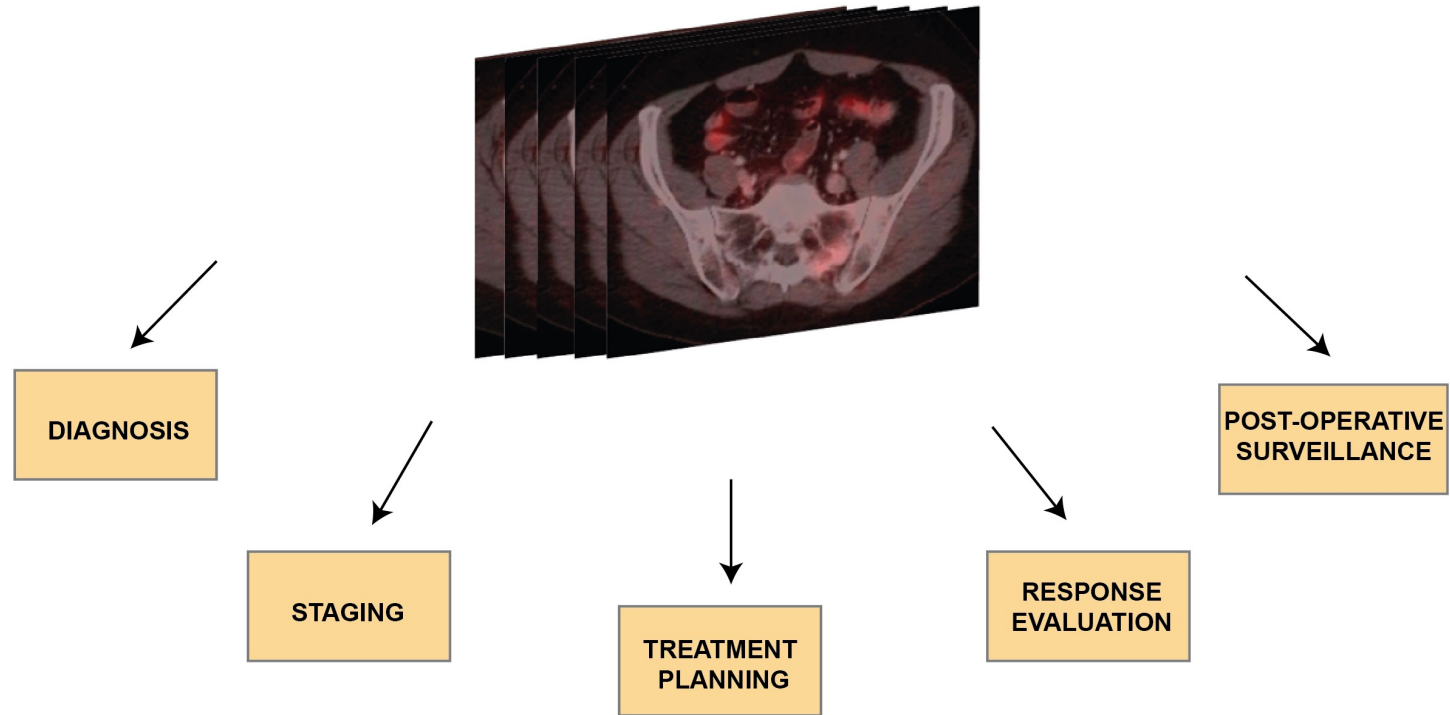
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Molecular techniques help understand how biological entities work



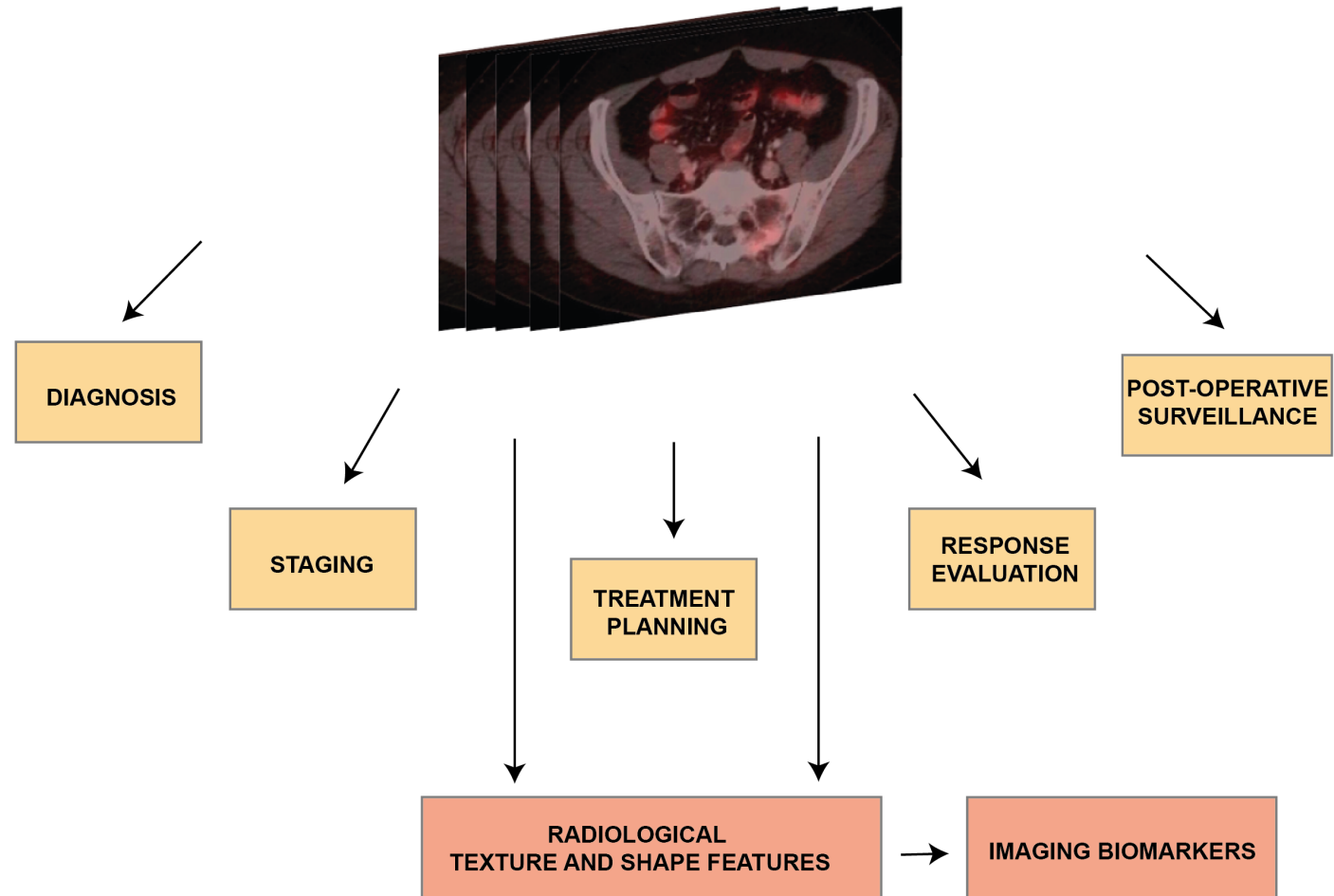
# State of art: The future of omics and big data

## Imaging techniques



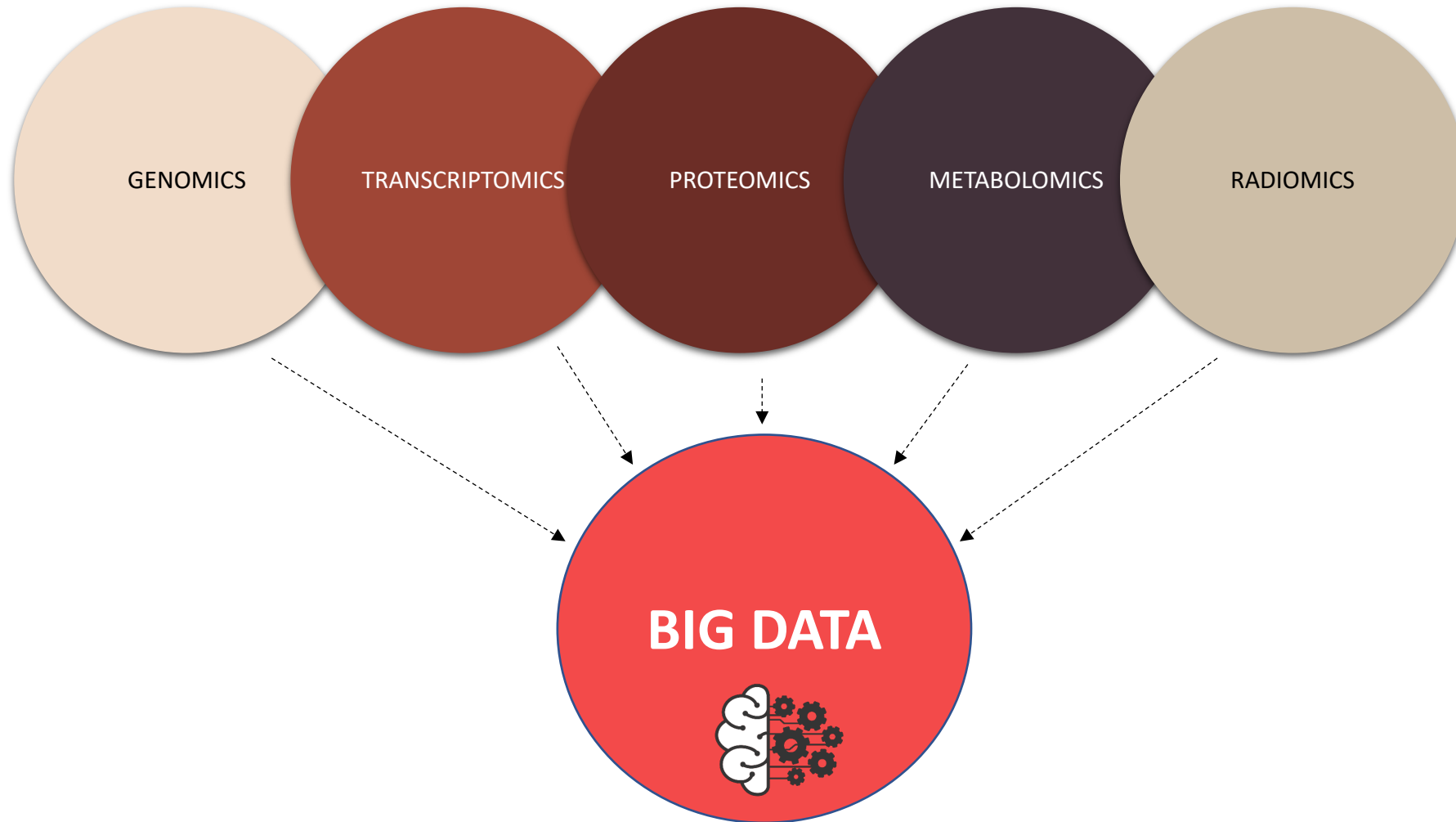
# State of art: The future of omics and big data

## Imaging techniques: RADIOMICS



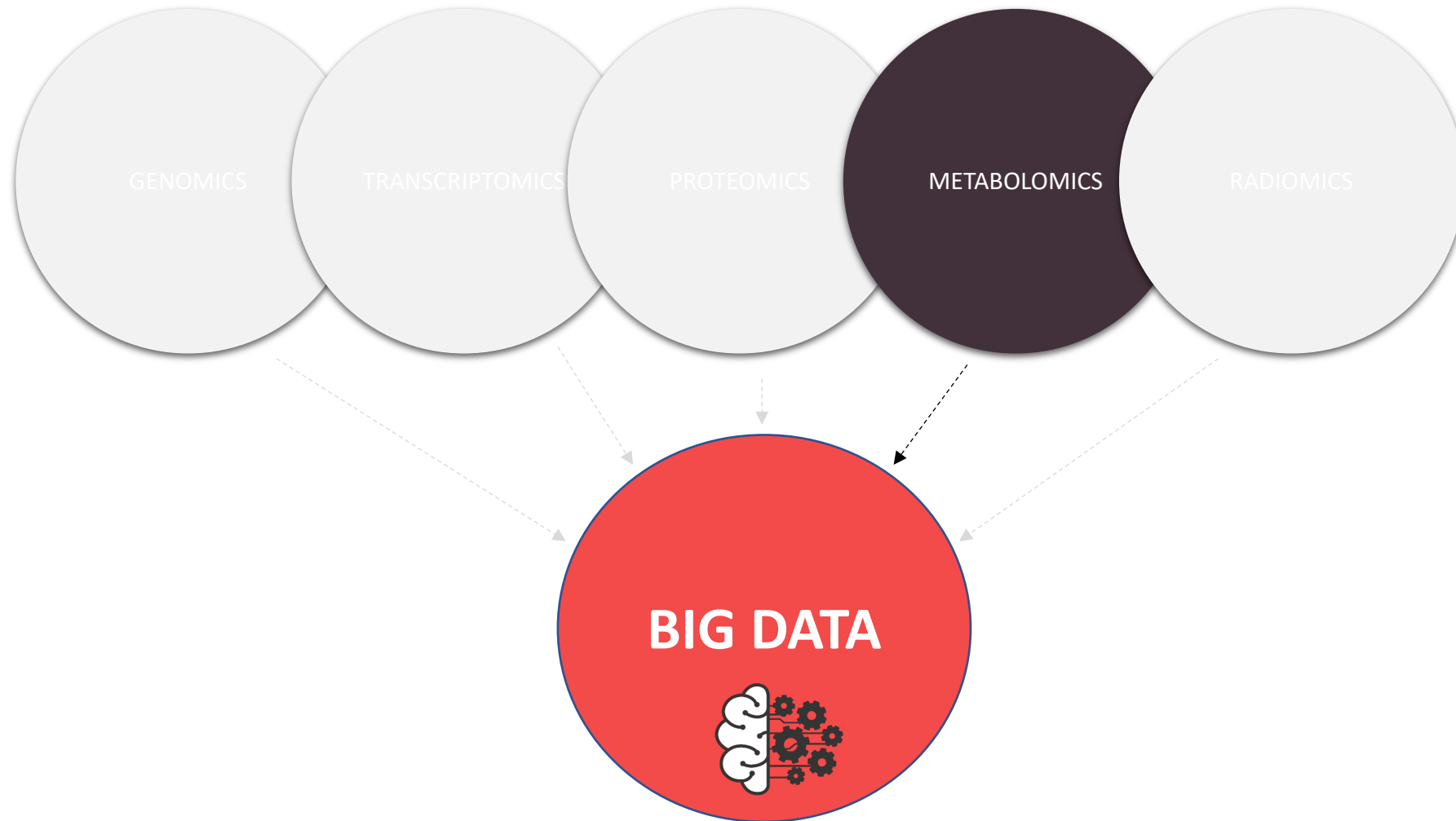
# State of art: The future of omics and big data

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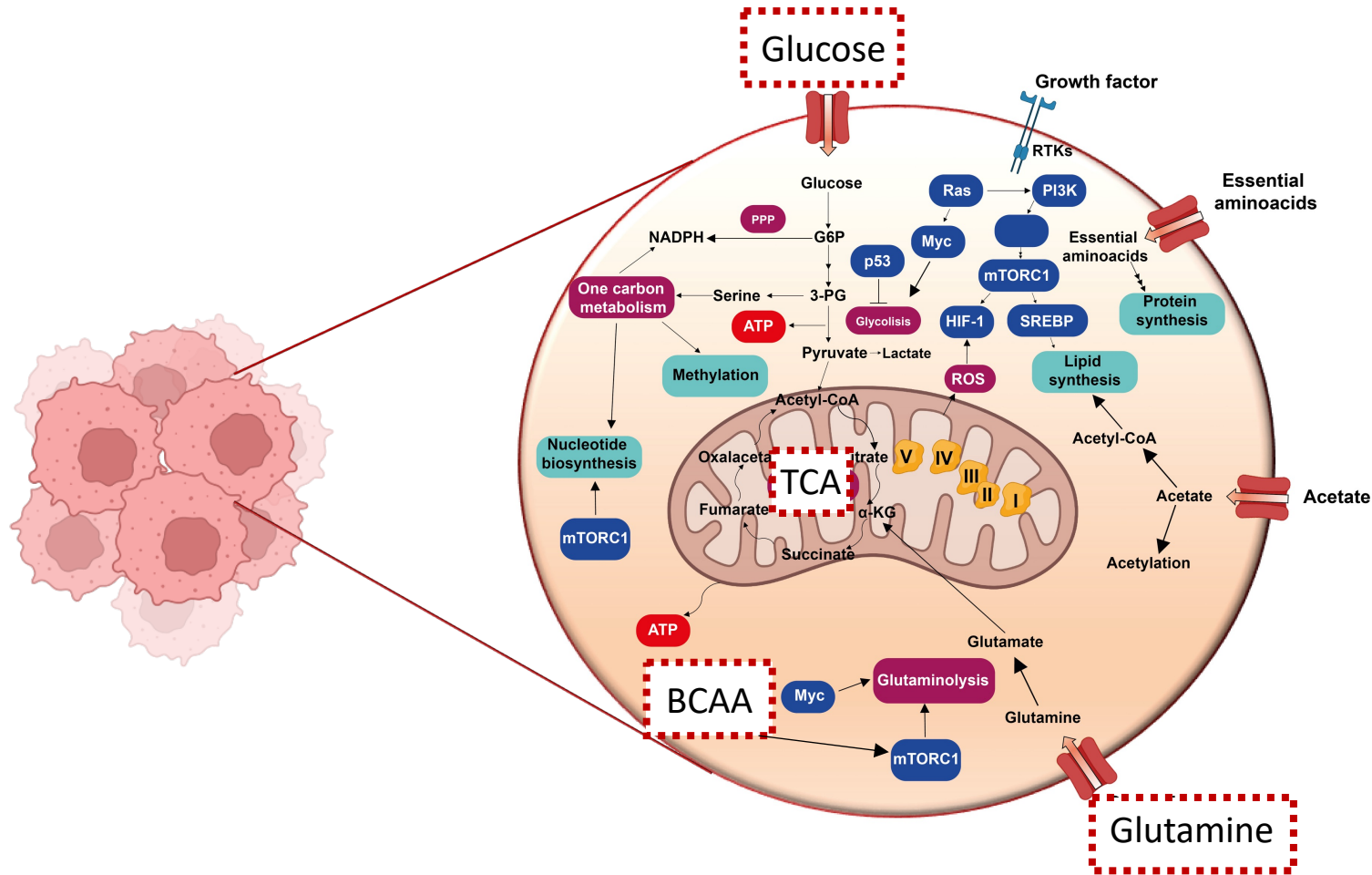


# State of art: The future of omics and big data

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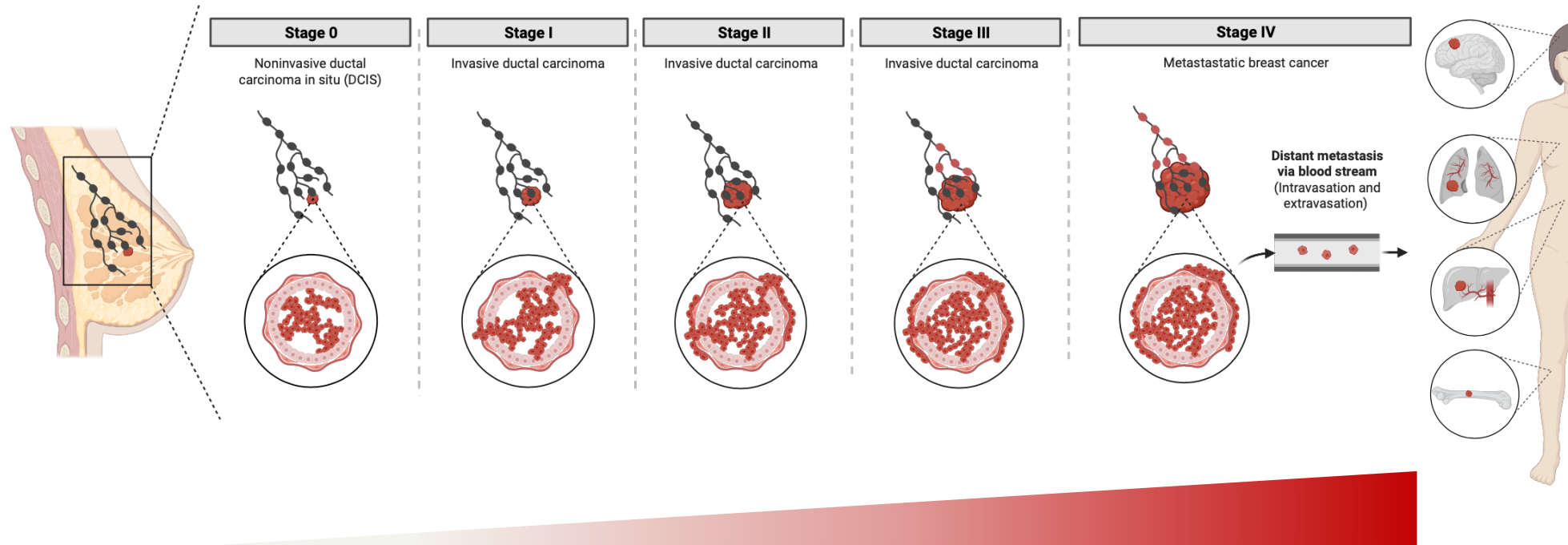


# Cancer metabolism under the spotlight



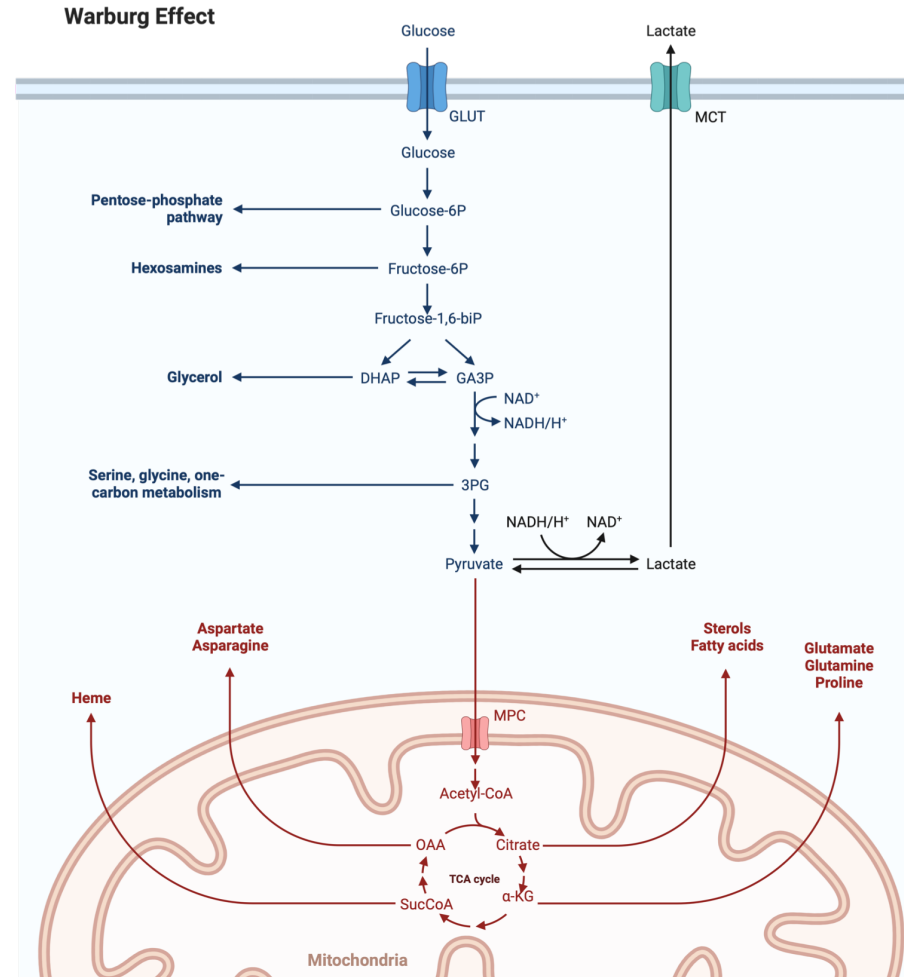
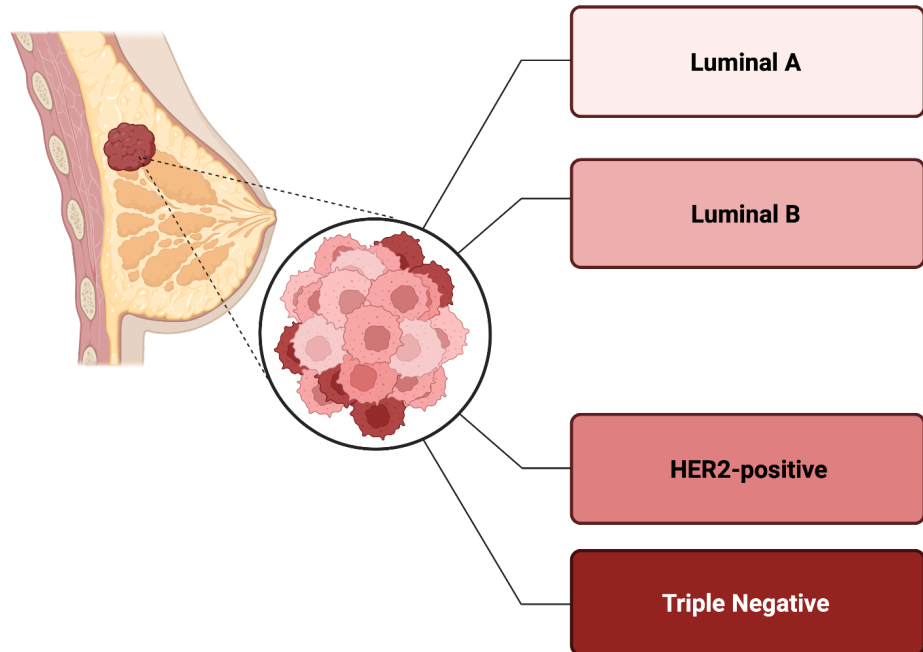


# Breast cancer metabolism under the spotlight



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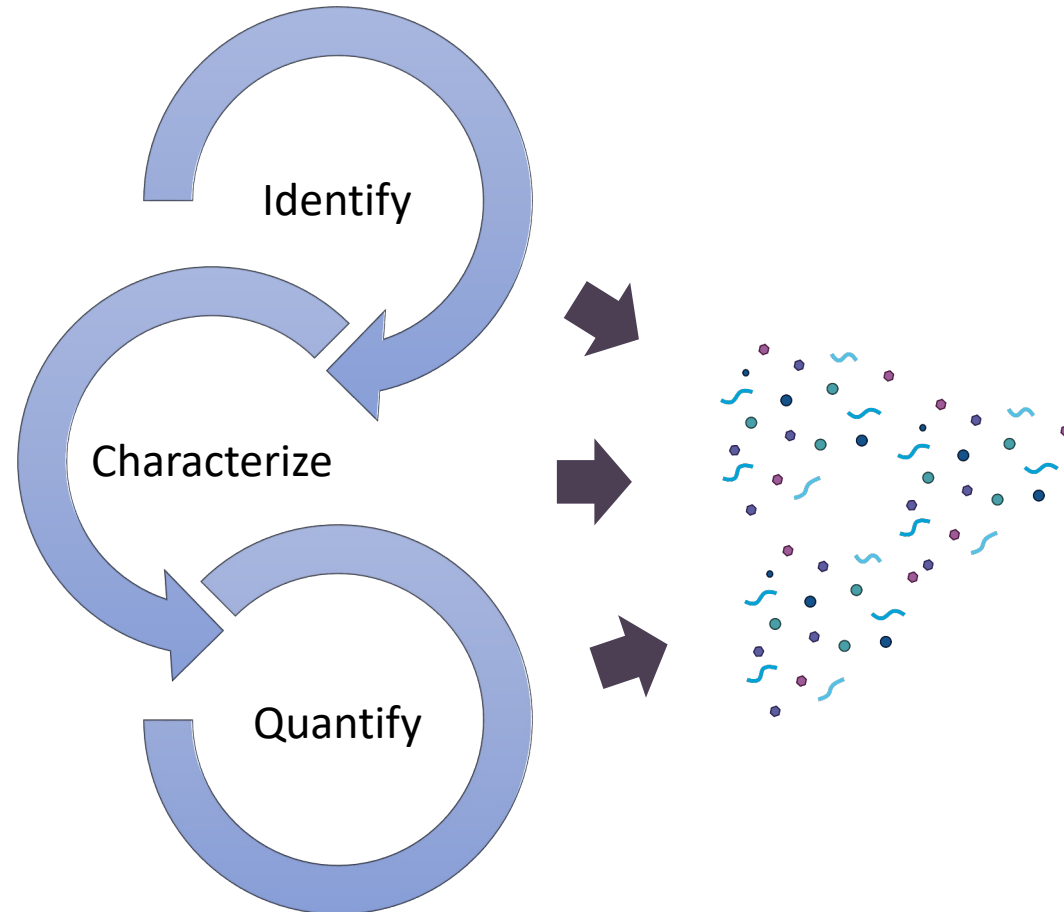
# Breast cancer metabolism under the spotlight



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# Cancer research focus: Targeted metabolomics

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# Research goals

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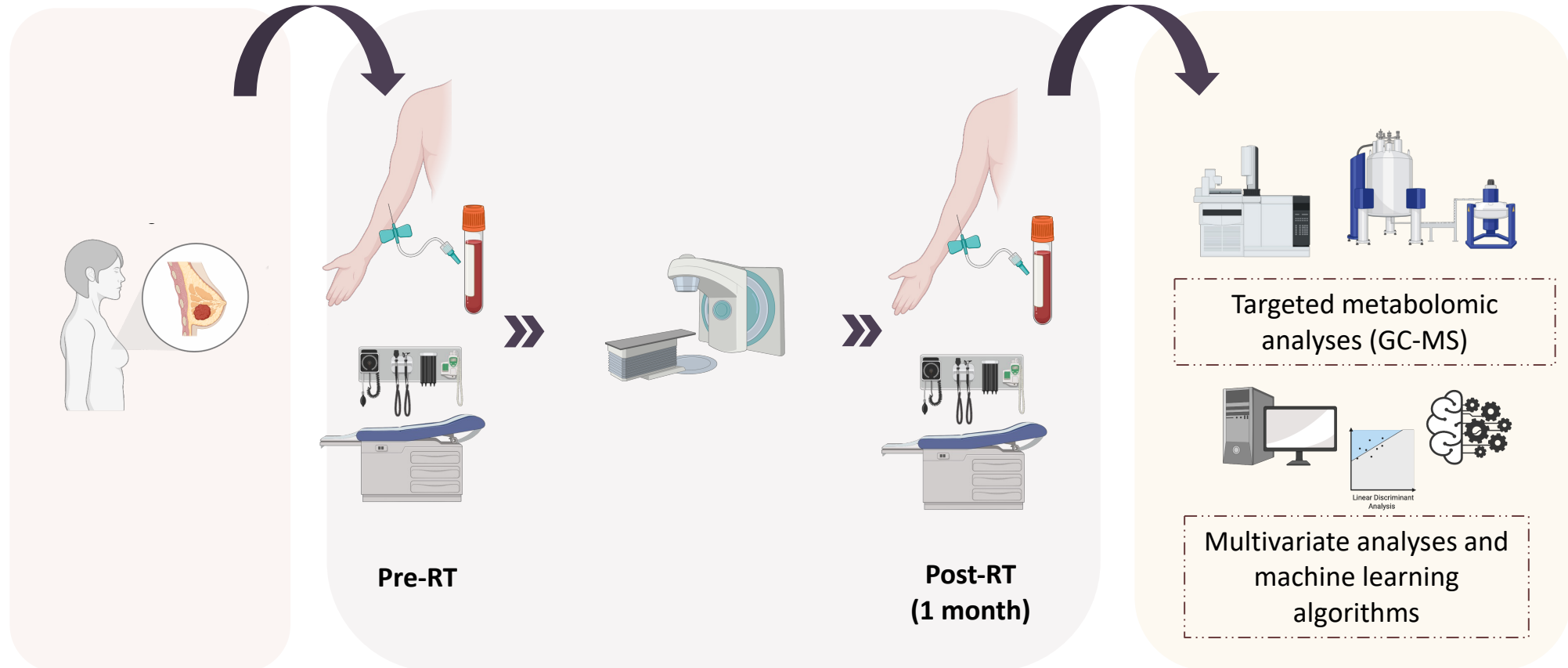
- Investigate circulating alterations in energy-balance-related metabolites.
- Correlate these changes with:
  - Clinicopathological characteristics
  - Response to radiation treatment and toxicity
- Identify potential biomarkers that could be implemented for diagnosis, prognosis and response towards treatment.

# Methodology

PATIENTS  
RECRUITMENT

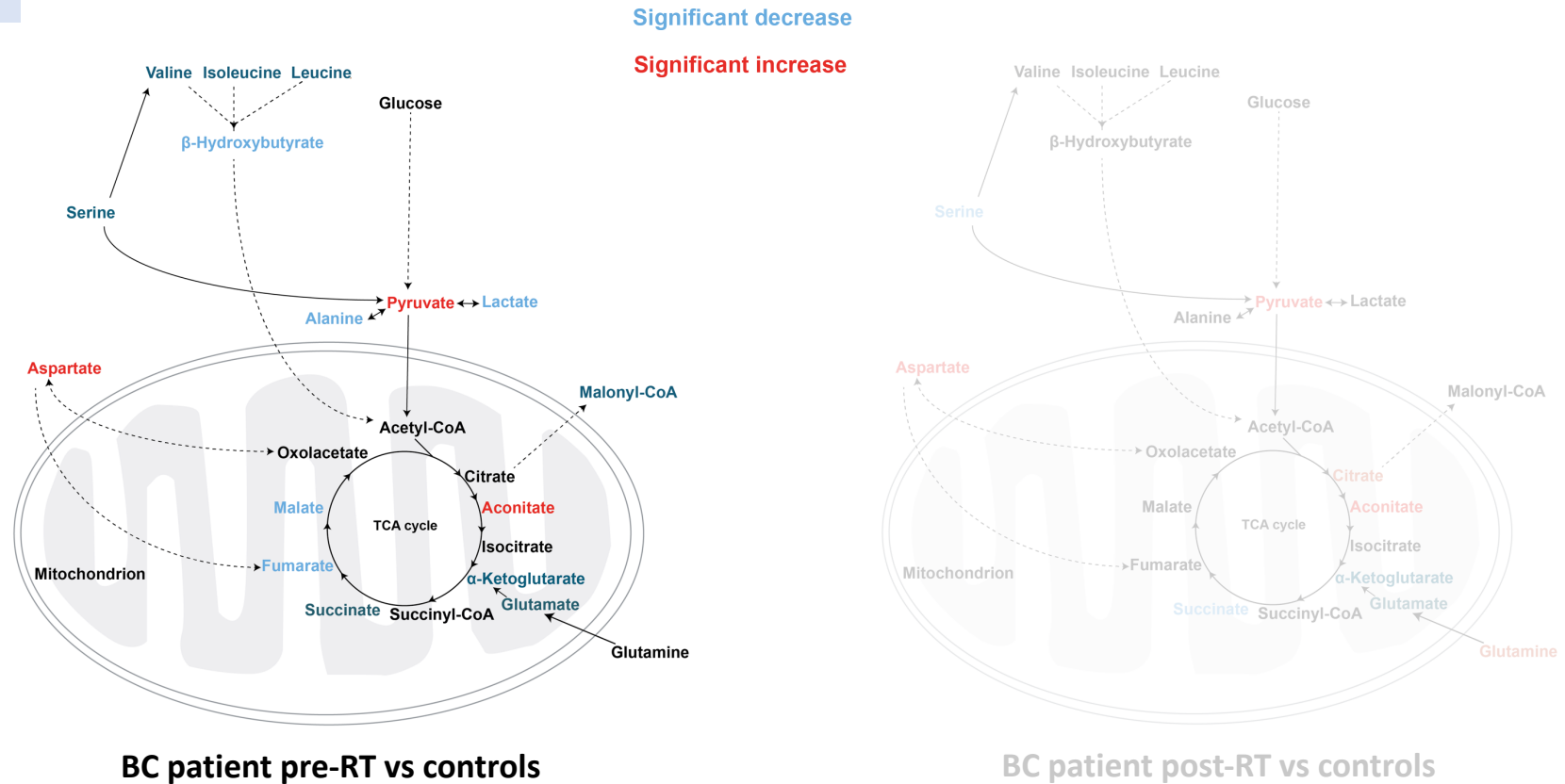
SAMPLE AND CLINICAL DATA COLLECTION

SAMPLE AND  
COMPUTATIONAL ANALYSIS



# Metabolite normalization with local radiotherapy following breast tumor resection

Breast cancer (n= 151)  
Controls (n= 44)



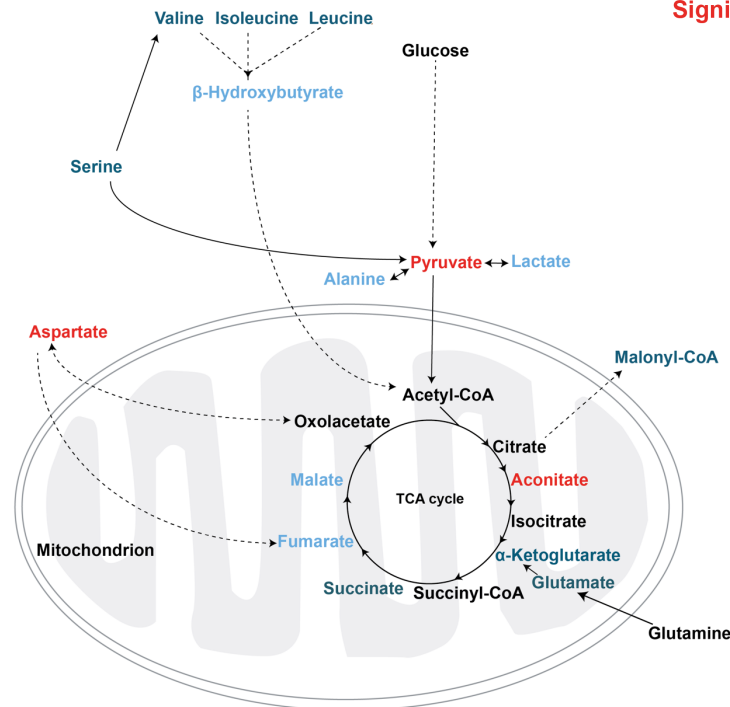
Arenas M, Rodríguez E, *et al.* PLoS One. (2018)

# Metabolite normalization with local radiotherapy following breast tumor resection

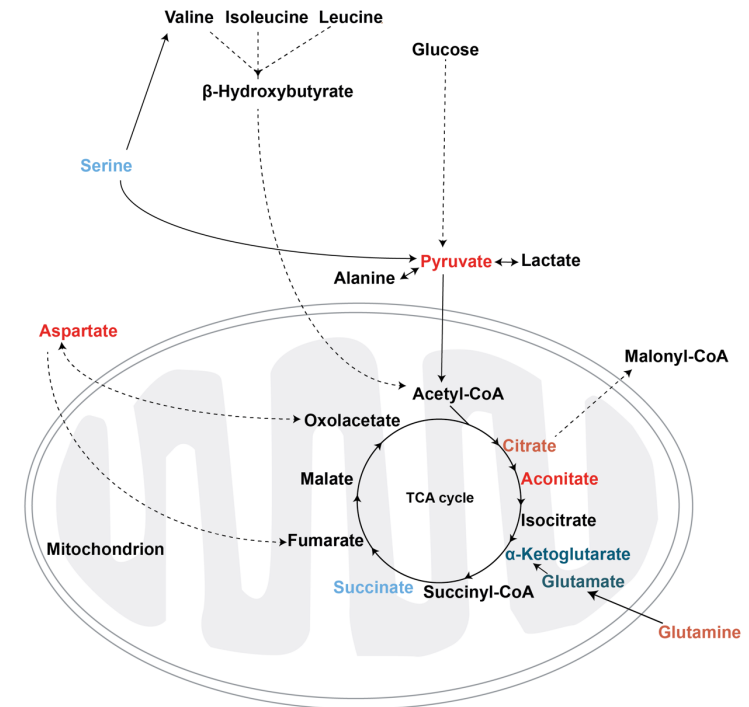
Breast cancer (n= 151)  
Controls (n= 44)

Significant decrease

Significant increase



BC patient pre-RT vs controls

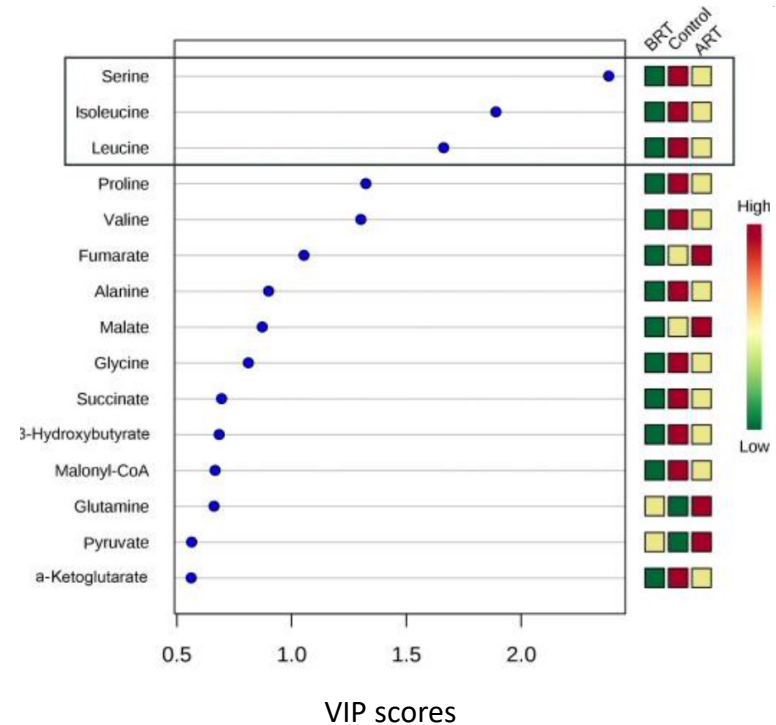
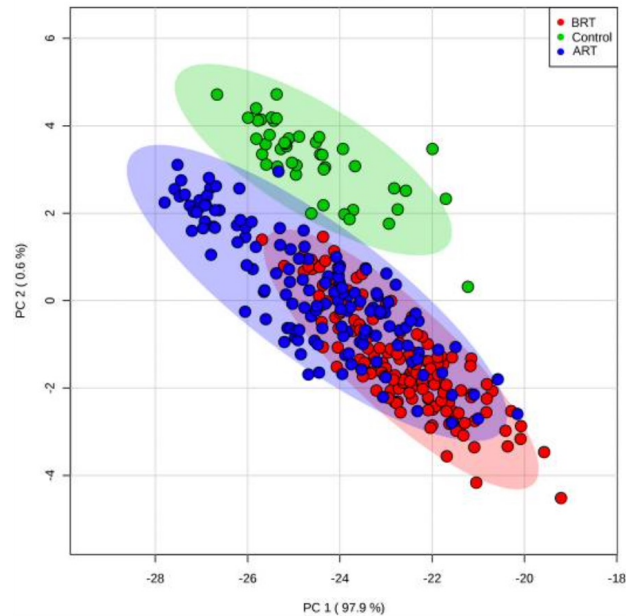


BC patient post-RT vs controls

Arenas M, Rodríguez E, et al. PLoS One. (2018)

# Metabolite normalization with local radiotherapy following breast tumor resection

Breast cancer (n= 151)  
Controls (n= 44)



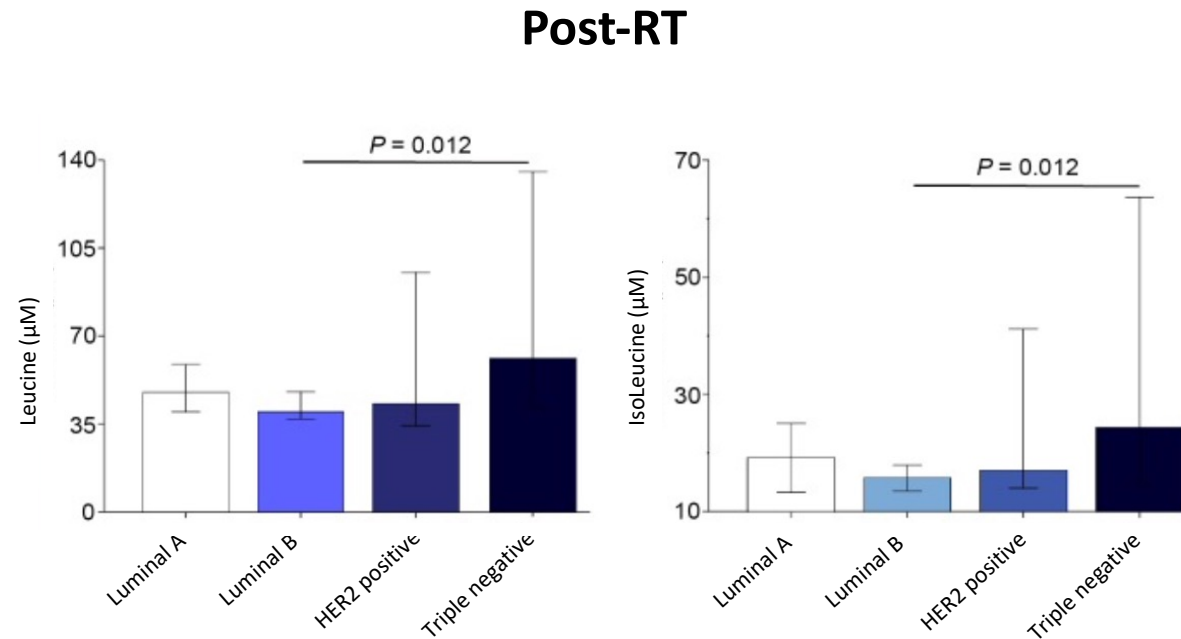
**BRT:** before radiotherapy  
**ART:** after radiotherapy

Arenas M, Rodríguez E, *et al.* PLoS One. (2018)



# Metabolite normalization with local radiotherapy following breast tumor resection

Breast cancer (n= 151)  
Controls (n= 44)



Arenas M, Rodríguez E, et al. PLoS One. (2018)

# Effects of radiotherapy on plasma energy metabolites in patients with breast cancer who received neoadjuvant chemotherapy

Breast cancer (n= 37)  
Controls (n= 44)

Variable	Control group (n=44)	BC patients pre-RT (n=37)		BC patients post-RT (n=37)	
		Partial response (n=24)	Complete response (n=13)	Partial response (n=24)	Complete response (n=13)
Pyruvate (µm)	23.81 (12.07)	61.22 (23.24) <sup>c</sup>	60.86 (19.69) <sup>c</sup>	61.50 (31.98) <sup>c</sup>	86.69 (26.63) <sup>c,d</sup>
Lactate (µm)	559.46 (62.55)	449.74 (60.20) <sup>c</sup>	445.42 (29.62) <sup>c</sup>	508.55 (130.10) <sup>b</sup>	594.59 (89.67) <sup>d</sup>
Alanine (µm)	199.99 (66.81)	142.99 (54.71) <sup>c</sup>	139.31 (52.65) <sup>c</sup>	193.32 (86.79)	271.42 (97.18) <sup>a,d</sup>
Hydroxybutyrate (µm)	24.18 (18.07)	22.35 (20.47)	18.68 (12.88)	22.36 (21.30)	24.88 (16.91)
Valine (µm)	121.68 (45.37)	68.99 (23.64) <sup>c</sup>	61.60 (23.06) <sup>c</sup>	111.70 (78.37)	174.15 (93.85) <sup>b,d</sup>
Leucine (µm)	68.23 (26.03)	30.21 (11.42) <sup>c</sup>	24.17 (11.22) <sup>c</sup>	52.08 (34.43) <sup>a</sup>	88.51 (56.67) <sup>a,d</sup>
Isoleucine (µm)	32.77 (15.91)	10.69 (4.87) <sup>c</sup>	<u>8.60 (5.57)<sup>c</sup></u>	21.38 (16.74) <sup>a</sup>	<u>36.17 (27.77)</u>
Proline (µm)	93.37 (29.73)	43.62 (20.35) <sup>c</sup>	39.03 (30.20) <sup>c</sup>	68.46 (48.44) <sup>c</sup>	119.30 (73.61) <sup>a,d</sup>
Malonyl coenzyme A (µm)	1.60 (0.31)	1.01 (0.34) <sup>c</sup>	<u>0.98 (0.33)<sup>c</sup></u>	1.22 (0.35) <sup>c</sup>	<u>1.59 (0.28)<sup>c</sup></u>
Glycine (µm)	135.40 (43.91)	73.68 (30.52) <sup>c</sup>	<u>70.52 (27.49)<sup>c</sup></u>	97.63 (34.99) <sup>c</sup>	<u>130.63 (26.69)<sup>d</sup></u>
Succinate (µm)	13.79 (4.20)	8.27 (2.35) <sup>c</sup>	<u>8.41 (2.79)<sup>c</sup></u>	10.44 (3.18) <sup>c</sup>	<u>12.67 (2.70)<sup>d</sup></u>
Fumarate (µm)	0.37 (0.17)	0.38 (0.23)	0.37 (0.16)	0.49 (0.31)	0.64 (0.31) <sup>a</sup>
Serine (µm)	56.51 (20.88)	15.75 (14.06) <sup>c</sup>	<u>9.18 (9.38)<sup>c</sup></u>	28.91 (23.95) <sup>c</sup>	<u>55.35 (36.10)<sup>d</sup></u>
Oxaloacetate (µm)	26.56 (7.32)	30.88 (11.70)	30.81 (17.55)	34.22 (12.65) <sup>a</sup>	43.31 (14.59) <sup>c</sup>
Malate (µm)	1.58 (0.76)	1.58 (1.21)	1.63 (0.68)	1.89 (1.03)	2.58 (1.18)
Aspartate (µm)	2.07 (0.91)	13.57 (4.07) <sup>c</sup>	15.51 (5.59)	13.36 (5.76)	17.17 (5.35) <sup>c</sup>
Ketoglutarate (µm)	7.48 (9.45)	3.52 (1.67) <sup>a</sup>	3.74 (1.64)	3.90 (2.35) <sup>a</sup>	4.61 (1.63)
Glutamate (µm)	135.47 (45.77)	46.06 (28.00) <sup>c</sup>	49.94 (23.39) <sup>c</sup>	49.18 (28.39) <sup>c</sup>	46.06 (20.71) <sup>c</sup>
Aconitate (µm)	0.12 (0.05)	0.85 (0.71) <sup>c</sup>	0.57 (0.53) <sup>c</sup>	0.61 (0.31) <sup>c</sup>	0.50 (0.31) <sup>c</sup>
Citrate (µm)	33.13 (5.86)	45.53 (6.27) <sup>a</sup>	43.48 (7.97)	49.57 (16.31) <sup>b</sup>	60.32 (11.58) <sup>c,d</sup>
Glutamine (µm)	36.73 (8.32)	49.15 (8.51)	52.22 (12.25)	54.60 (21.96) <sup>a</sup>	115.10 (21.53) <sup>c,e</sup>

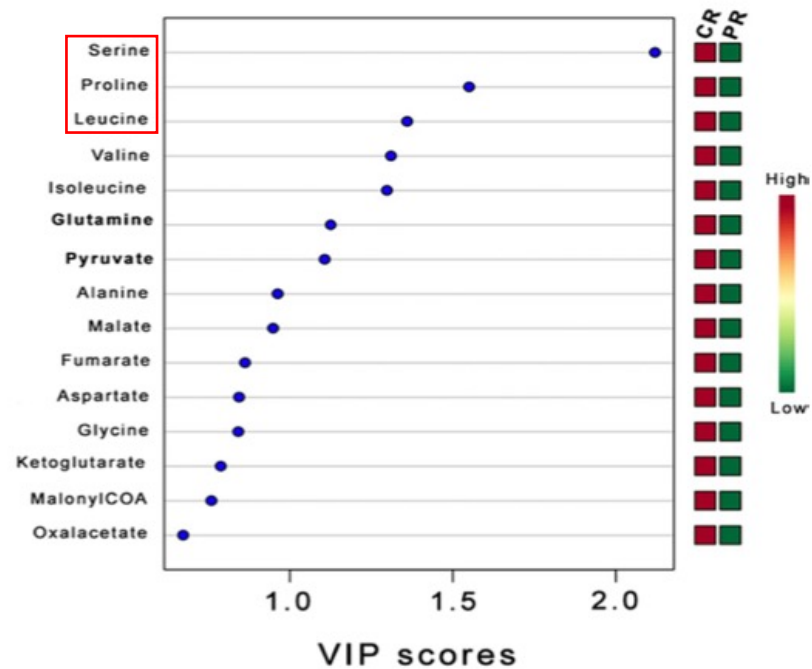
<sup>a</sup> p < 0.05; <sup>b</sup> p < 0.01; <sup>c</sup> p < 0.001, with respect to control.

<sup>d</sup> p < 0.05; <sup>e</sup> p < 0.01, with respect to partial response.

Arenas M, Fernández-Arroyo S, Rodríguez E, et al. Clin Trans Oncol. (2020)

# Effects of radiotherapy on plasma energy metabolites in patients with breast cancer who received neoadjuvant chemotherapy

Breast cancer (n= 37)  
Controls (n= 44)



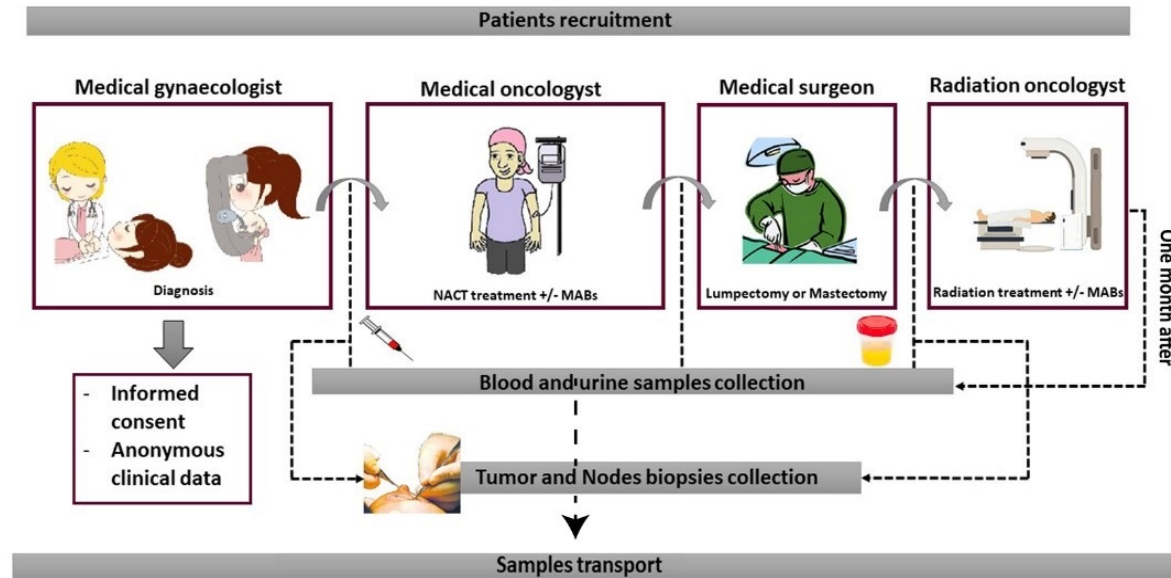
Complete response (CR) vs partial response (PR)

Arenas M, Fernández-Arroyo S, Rodríguez E, *et al.* Clin Trans Oncol. (2020)

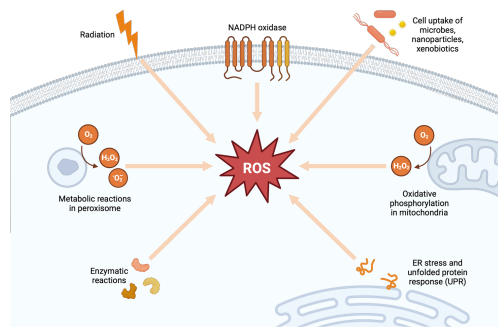
# MAMACOR study

**WORK IN PROGRESS**

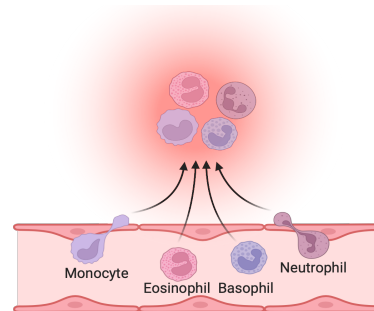
Breast cancer (n= 87)  
Controls (n= 50)



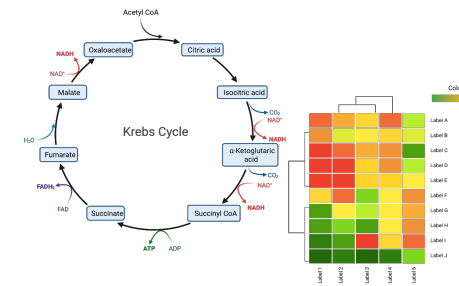
## Oxidative stress markers analysis



## Inflammatory markers analysis



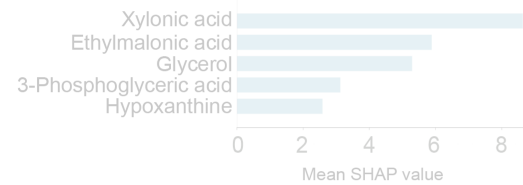
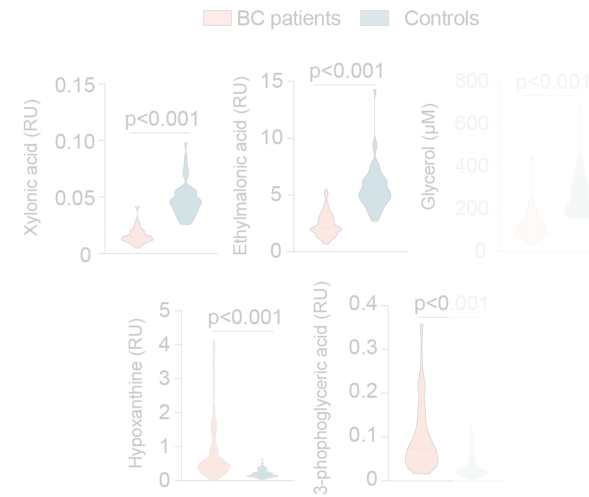
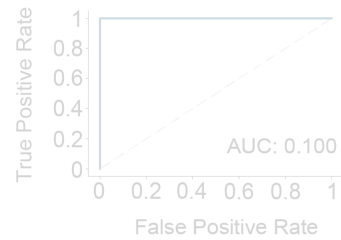
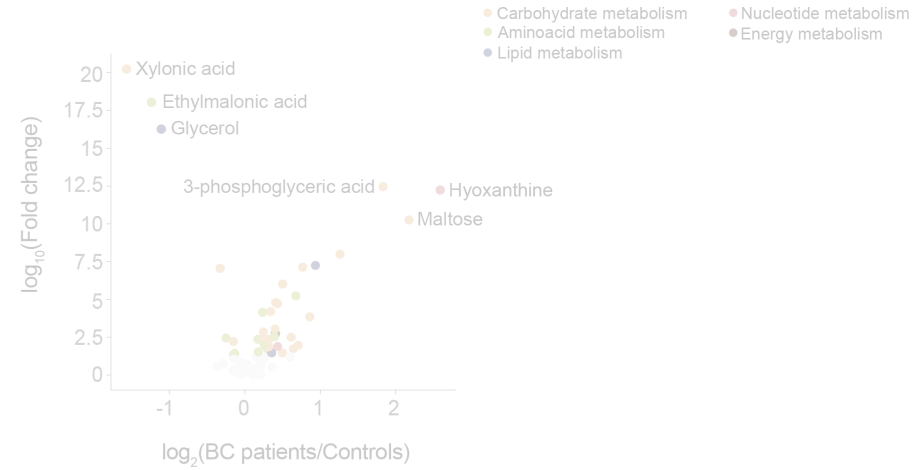
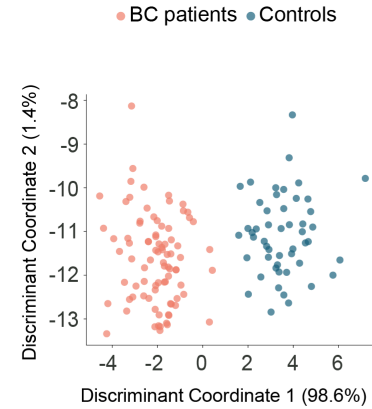
## Metabolomics analysis



# MAMACOR study

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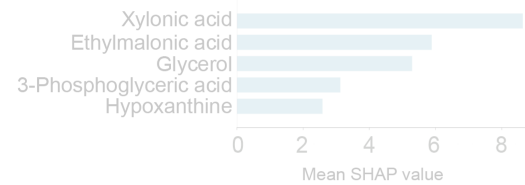
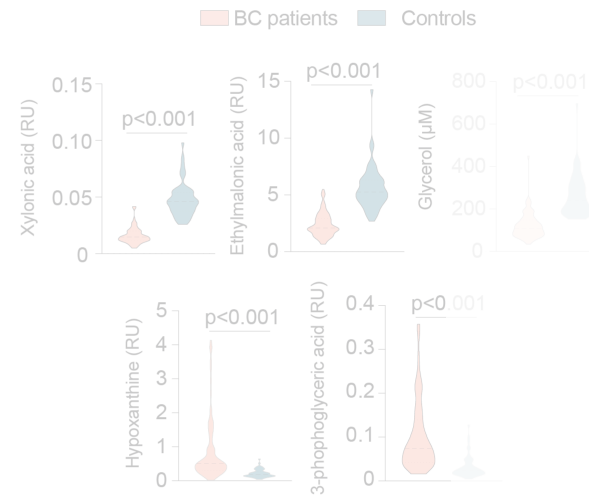
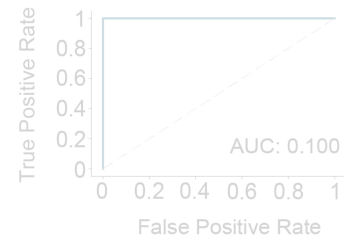
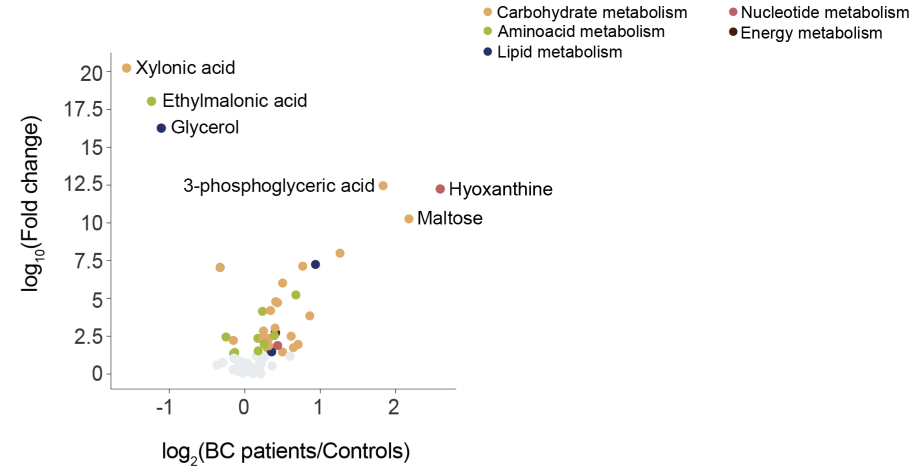
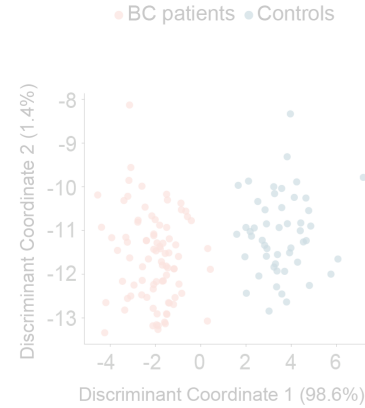
Breast cancer (n= 87)  
Controls (n= 50)



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WORK IN  
PROGRESS

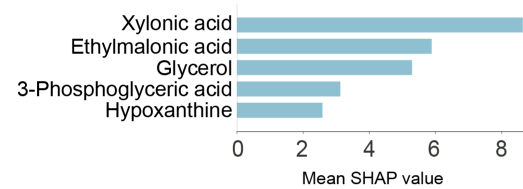
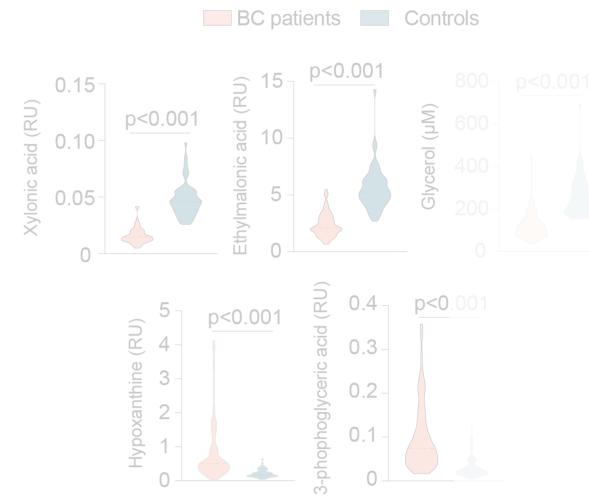
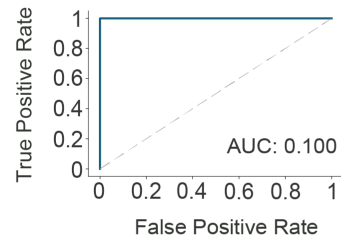
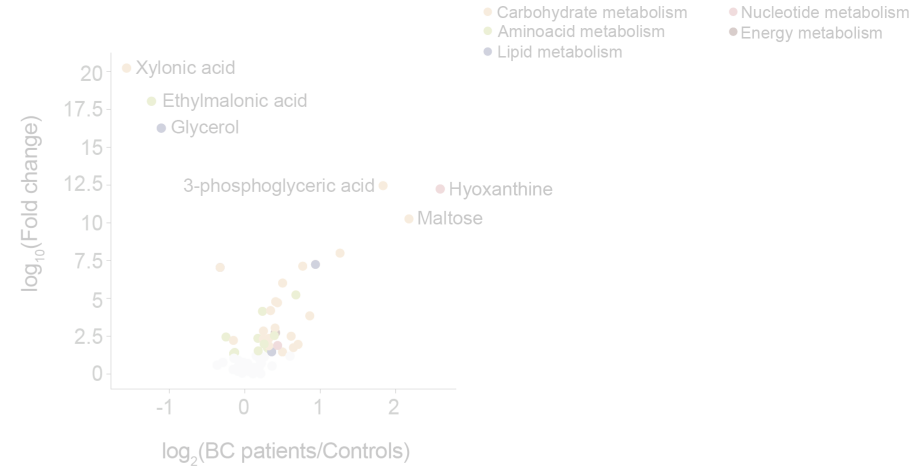
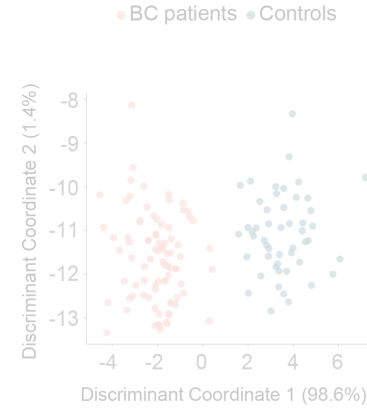
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# MAMACOR study

WORK IN  
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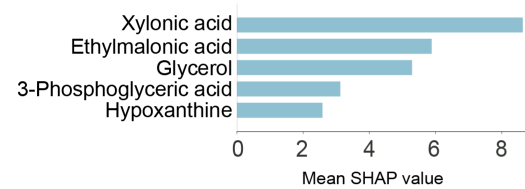
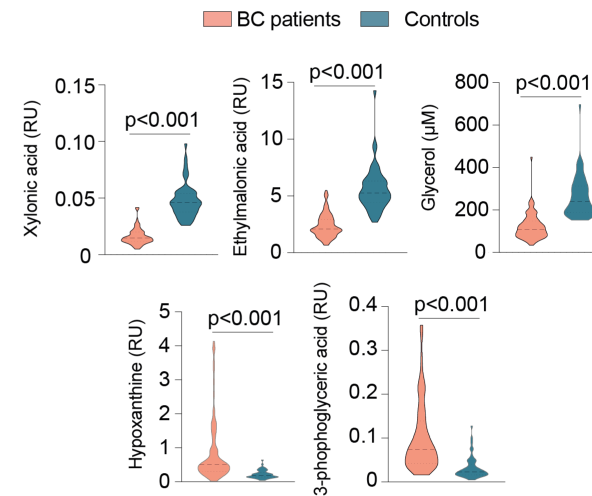
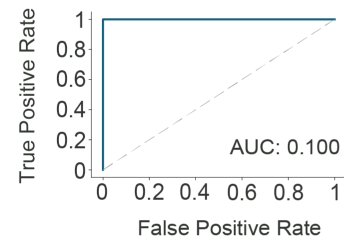
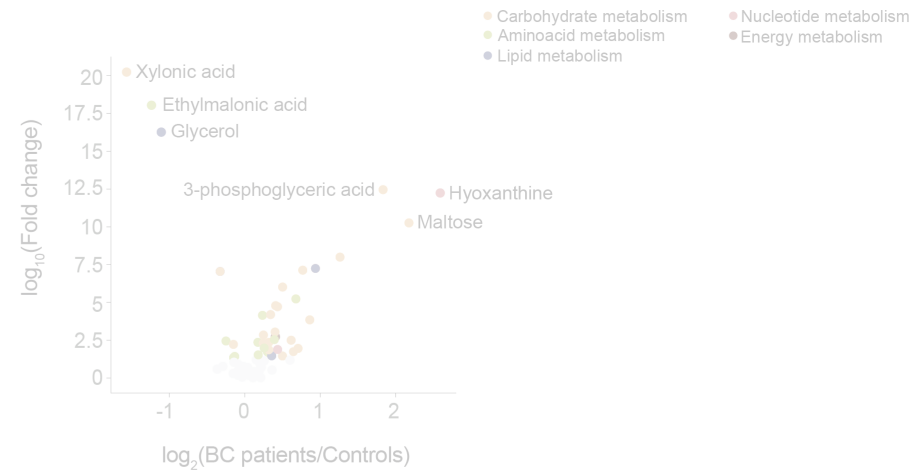
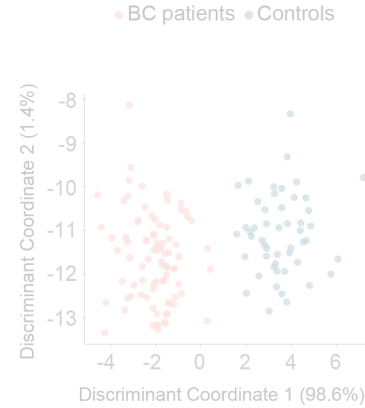
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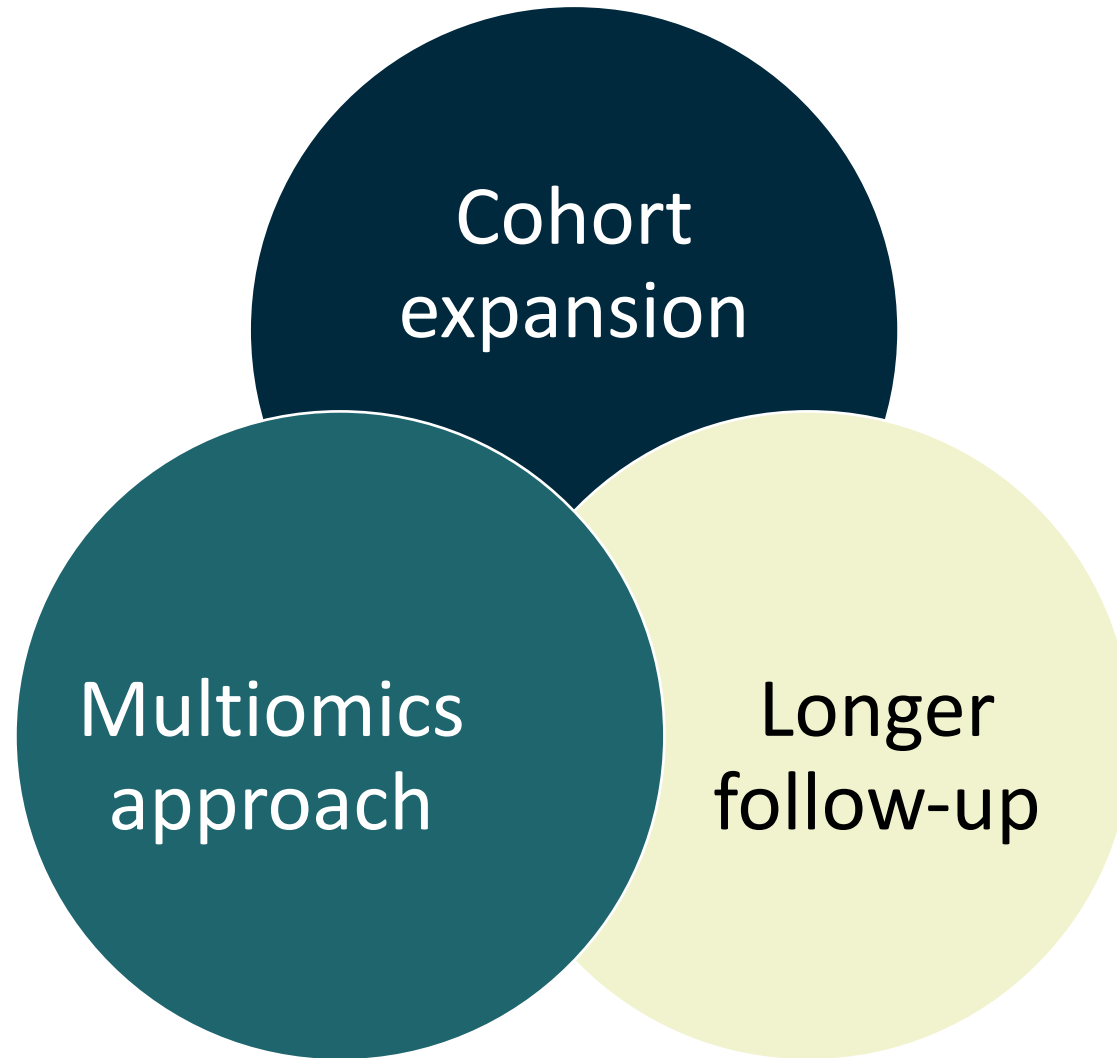
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Controls (n= 50)





# Next steps in our research group

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# Aknowledgements

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## **MRO organizers**

**Dr. Meritxell Arenas** from the Radiation Oncology Department in the  
Universitary Hospital Sant Joan in Reus

**Dr. Jorge Joven and Dr. Jordi Camps** from the Biomedical Research Unit  
in the Universitary Hospital Sant Joan in Reus



# Omics and AI in breast cancer

Omics driven radiotherapy approaches

**Elisabet Rodríguez Tomàs**

Radiation Oncology Department and Biomedical Research Unit

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