



# Clinically relevant multidrug transporters are regulated by different microRNAs along the human intestine

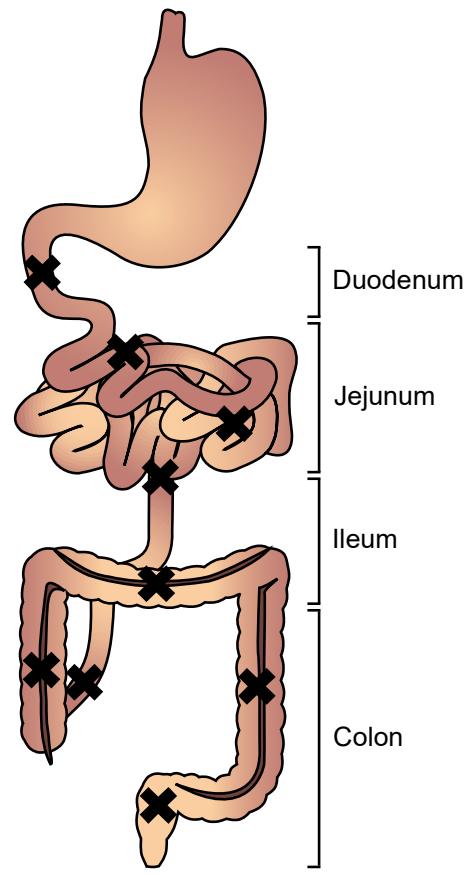
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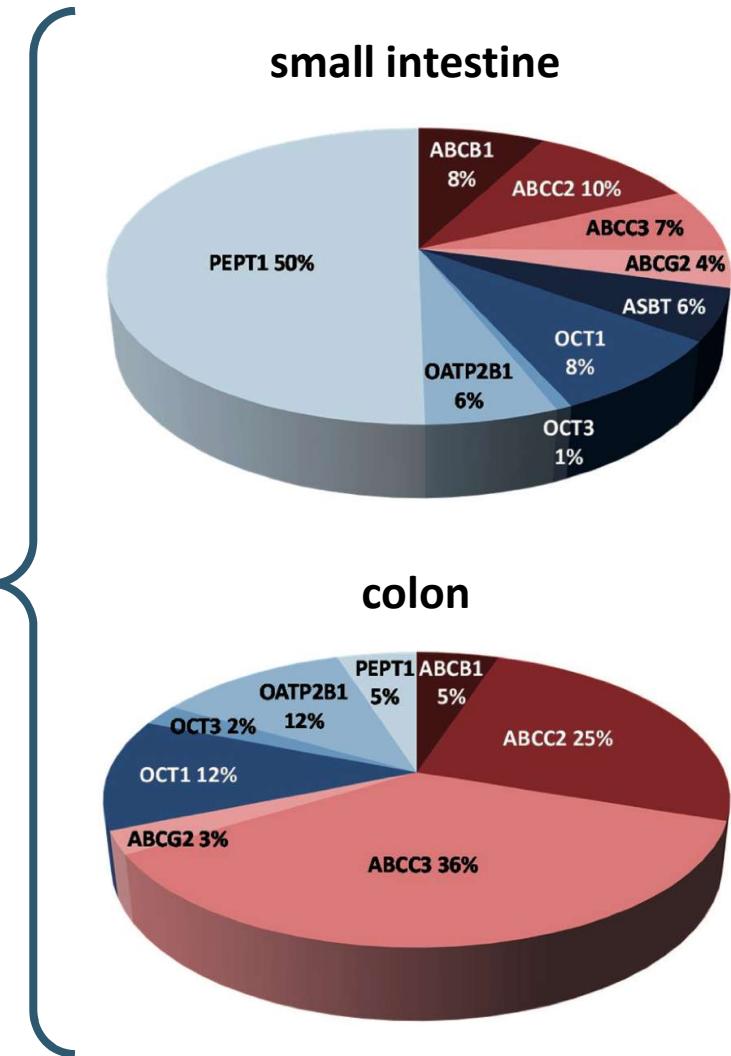


# Site-specific mRNA and protein levels of transporters along the human intestine

## Localisation of biopsies from organ donors



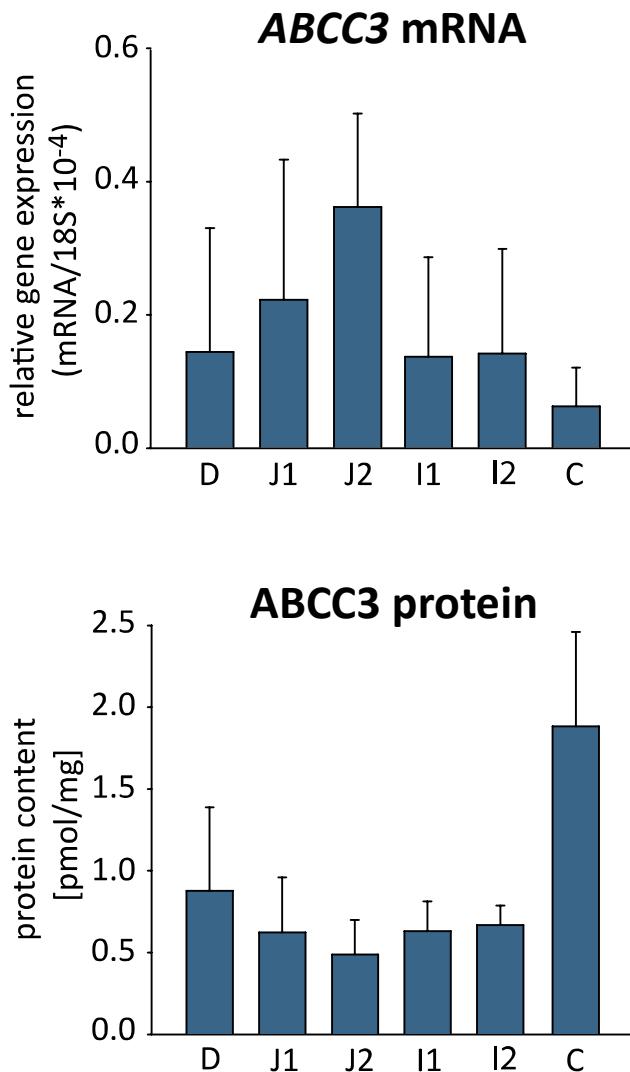
targeted proteomics



✖ positions of tissue biopsies

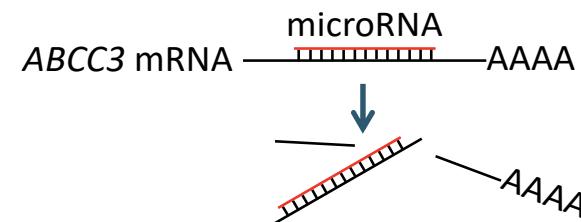
Drozdzik, M. et al. Protein Abundance of Clinically Relevant Multidrug Transporters Along the Entire Length of the Human Intestine. *Mol. Pharmaceutics* 2014, 11 (10), 3547–3555.

# Discrepancy between mRNA and protein levels

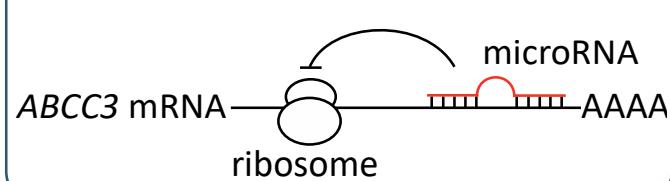


## Role of microRNAs?

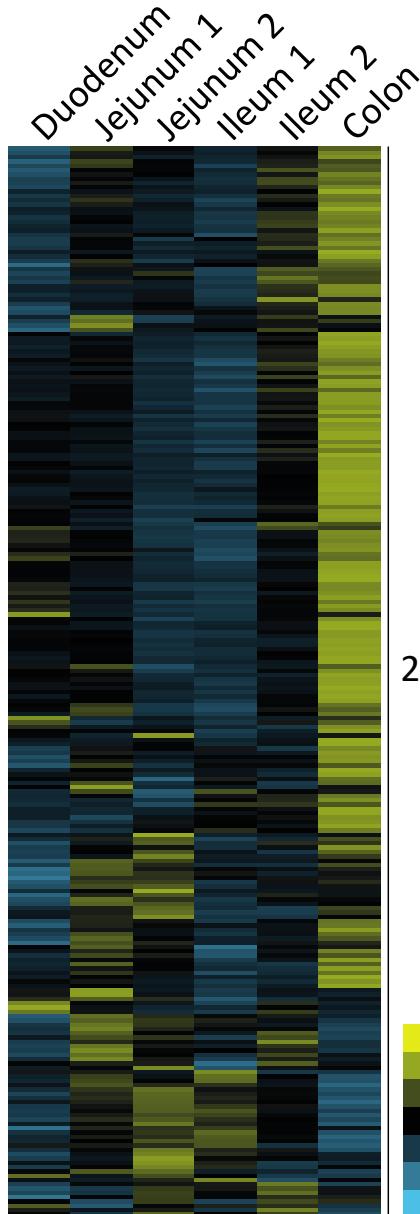
### mRNA degradation



### transcriptional repression



# Site-specific microRNA expression along the human intestine



## Correlation microRNA/protein + *in silico*-target prediction

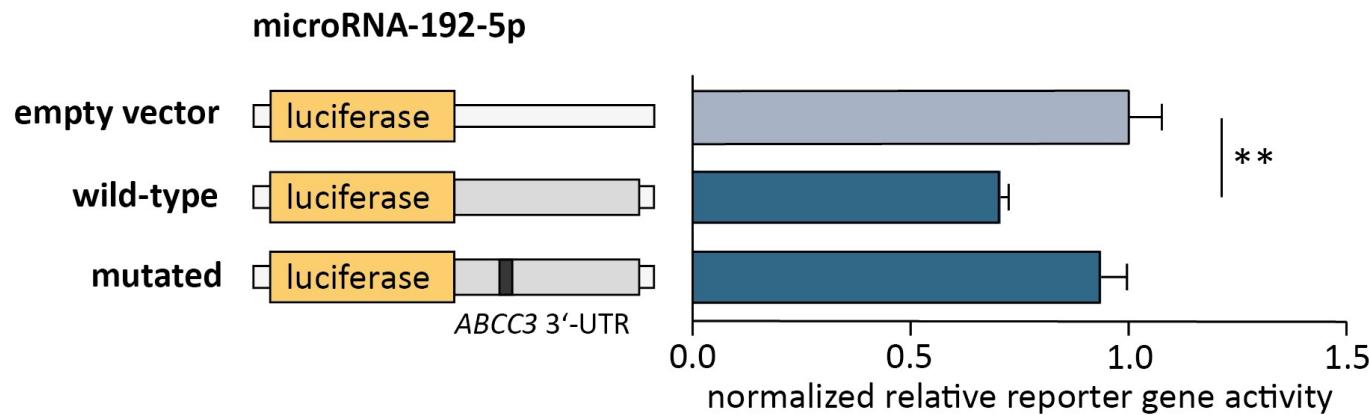
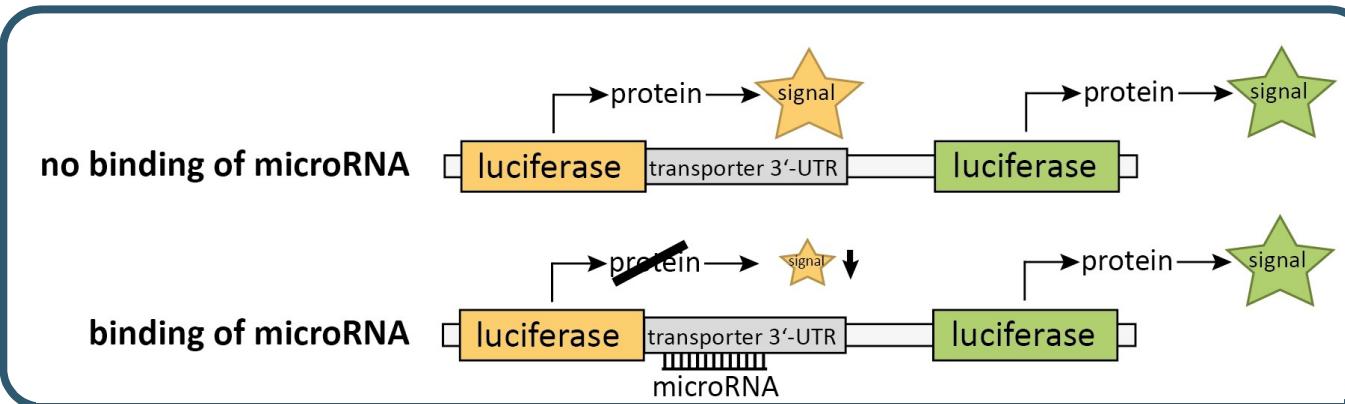
### correlation coefficients

	ABCB1	ABCC2	ABCC3	ABCG2	PEPT1
<b>mir-27a-3p</b>	-0.21	n.s.	n.s.	-0.37*	-0.50**
<b>miR-100-5p</b>	n.s.	n.s.	n.s.	-0.44**	-0.48**
<b>miR-192-5p</b>	n.s.	n.s.	-0.52**	n.s.	-0.48**
<b>miR-193a-3p</b>	n.s.	n.s.	n.s.	-0.37*	-0.49**
<b>miR-193b-3p</b>	n.s.	n.s.	n.s.	-0.35*	-0.53**
<b>miR-215-3p</b>	n.s.	n.s.	-0.63**	n.s.	-0.55**
<b>miR-365a-3p</b>	n.s.	n.s.	n.s.	-0.40*	-0.48**
<b>miR-409-3p</b>	-0.41*	n.s.	n.s.	-0.48**	-0.36**
<b>miR-766-3p</b>	-0.34*	n.s.	n.s.	-0.47**	-0.48**

Pearson correlation: p-value \* < 0.05, \*\* < 0.01; n.s. = not significant

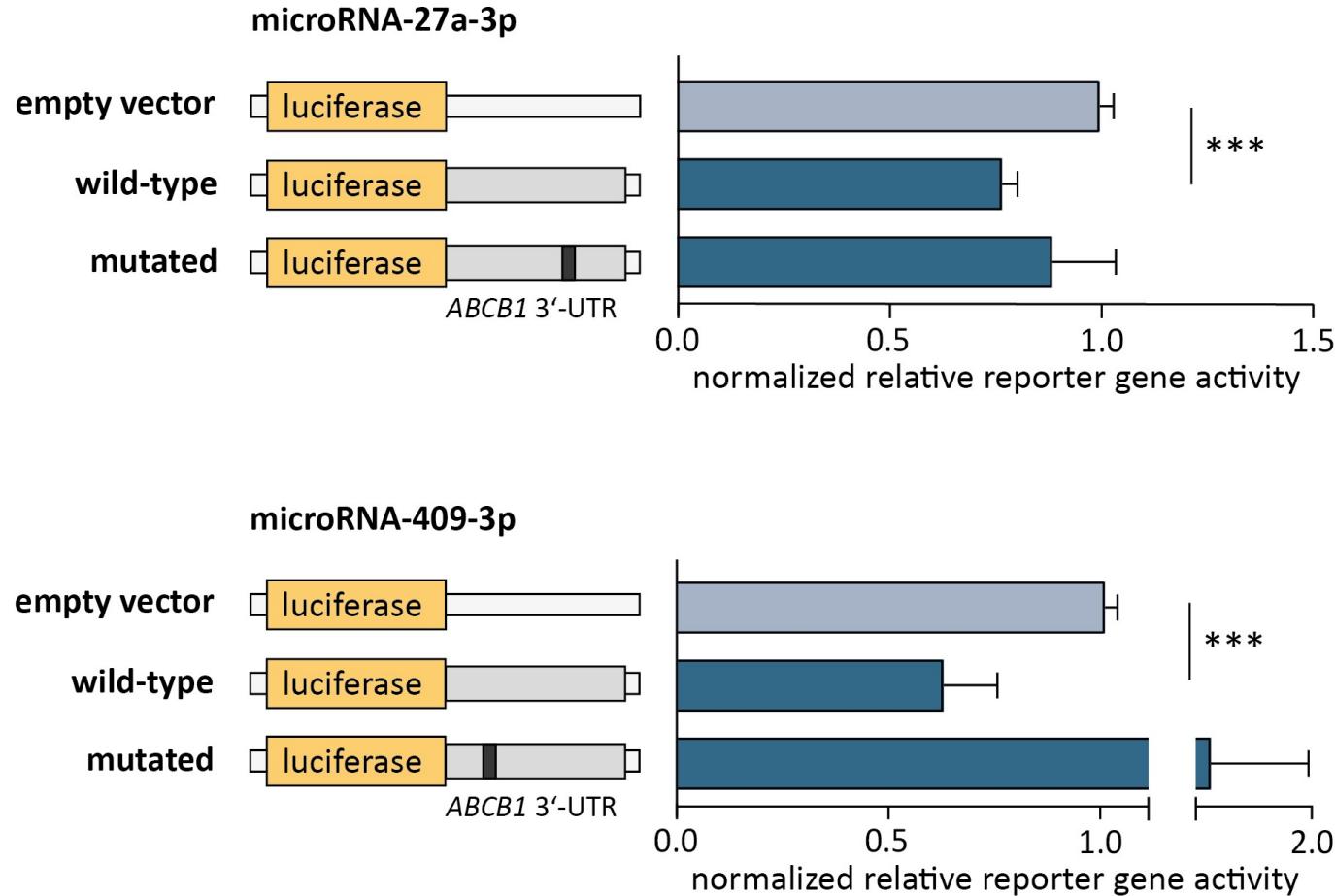
# Verification of miR-192-5p / ABCC3 3'-UTR interaction

## Principle of reporter gene assay



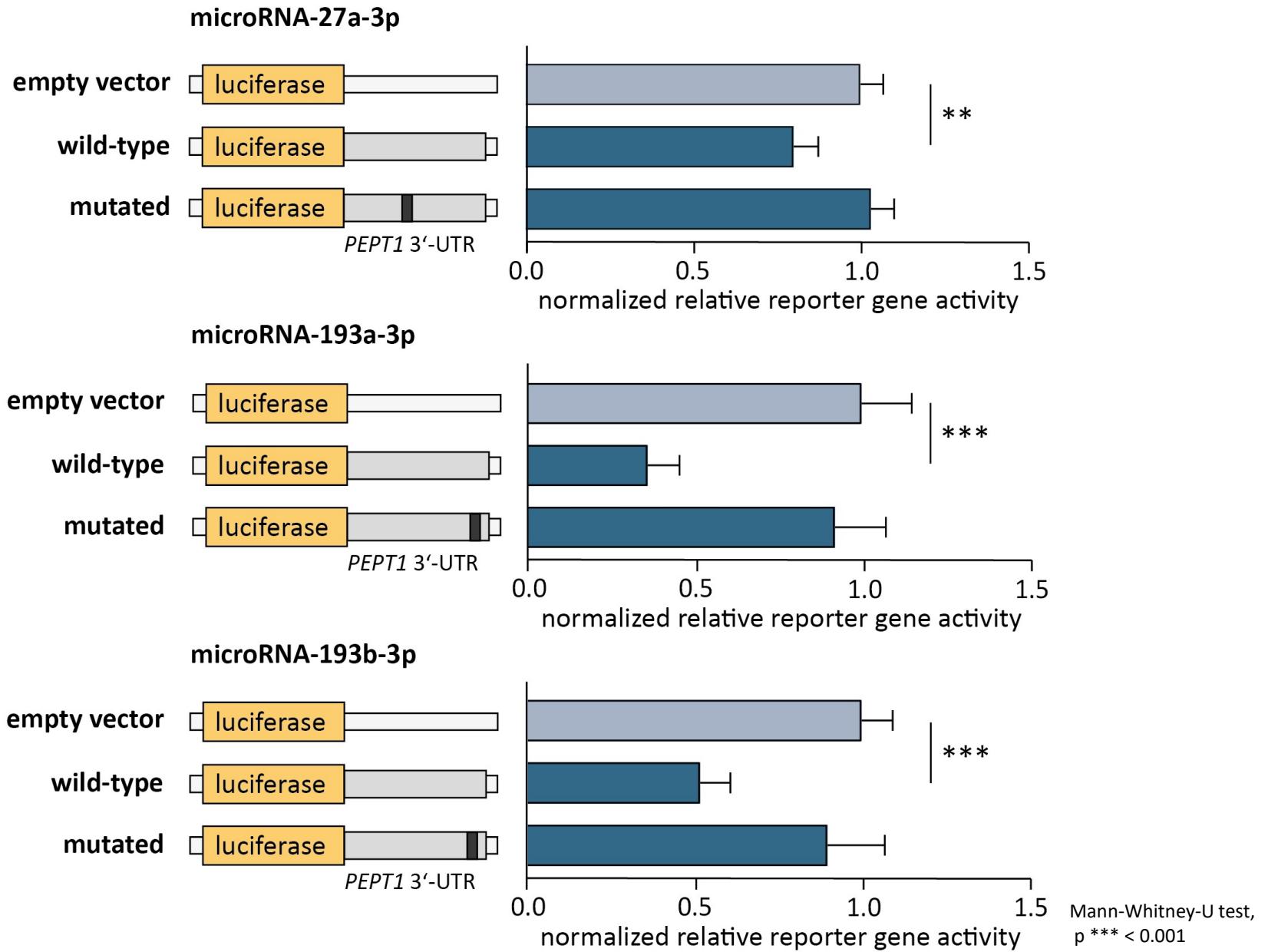
Mann-Whitney-U test, p \*\* < 0.01

# Verification of miR-27a-3p / miR-409-3p / *ABCB1* 3'-UTR interactions

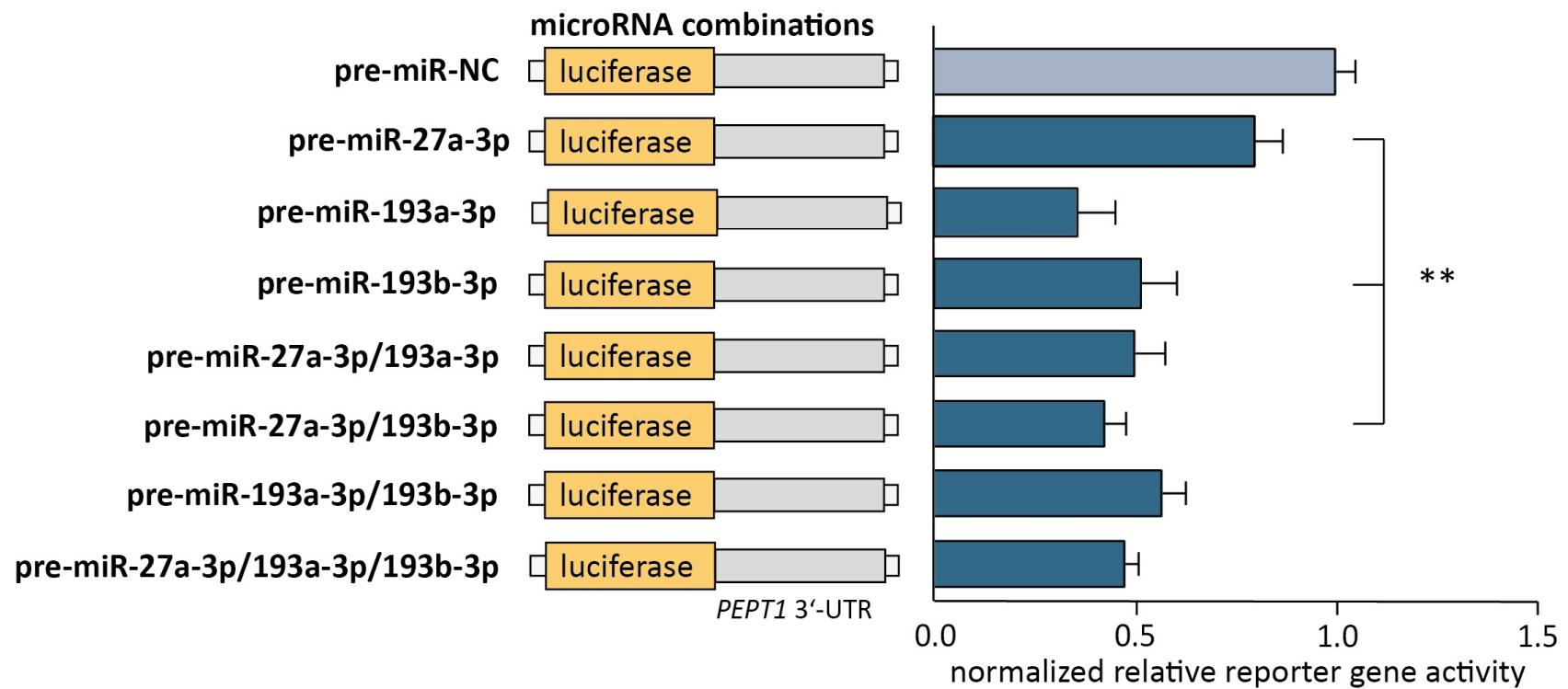


Mann-Whitney-U test, p \*\*\* < 0.001

# Verification of miR-27a-3p / miR-193a-3p / miR-193b-3p / *PEPT1* (*SLC15A1*) 3'-UTR interactions

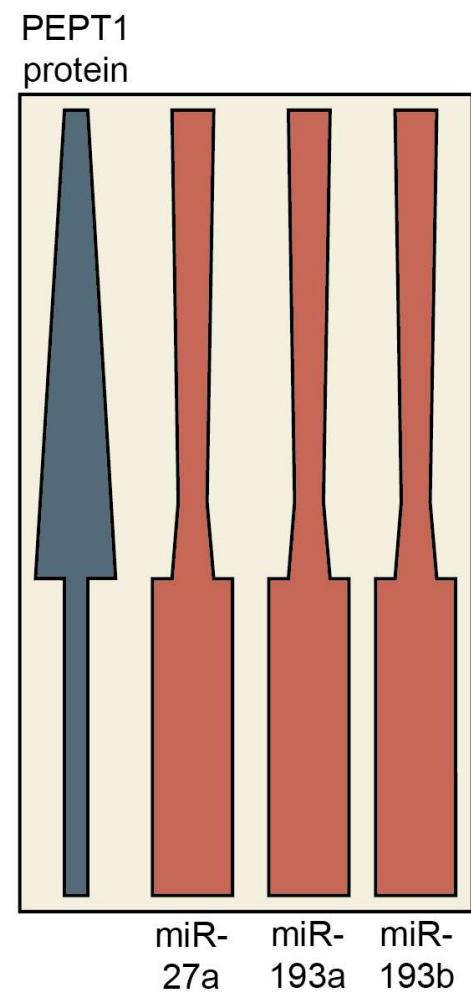
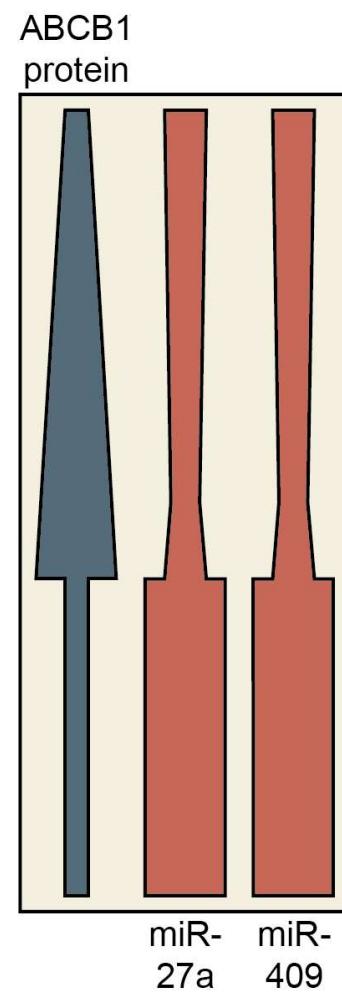
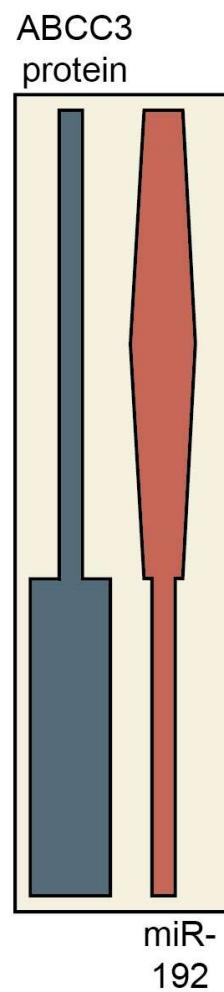
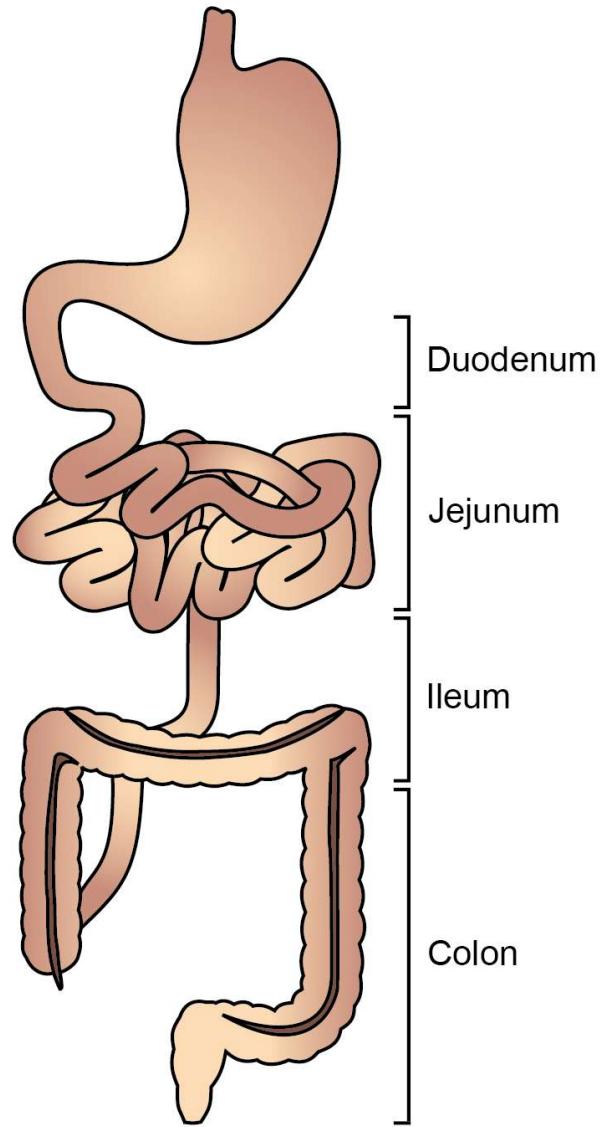


# Combined effects of multiple microRNAs on *PEPT1* (*SLC15A1*) 3'-UTR



Mann-Whitney-U test, p \*\*< 0.01

# Summary



# Acknowledgement

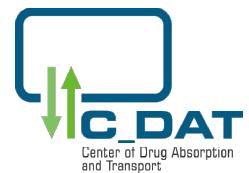
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