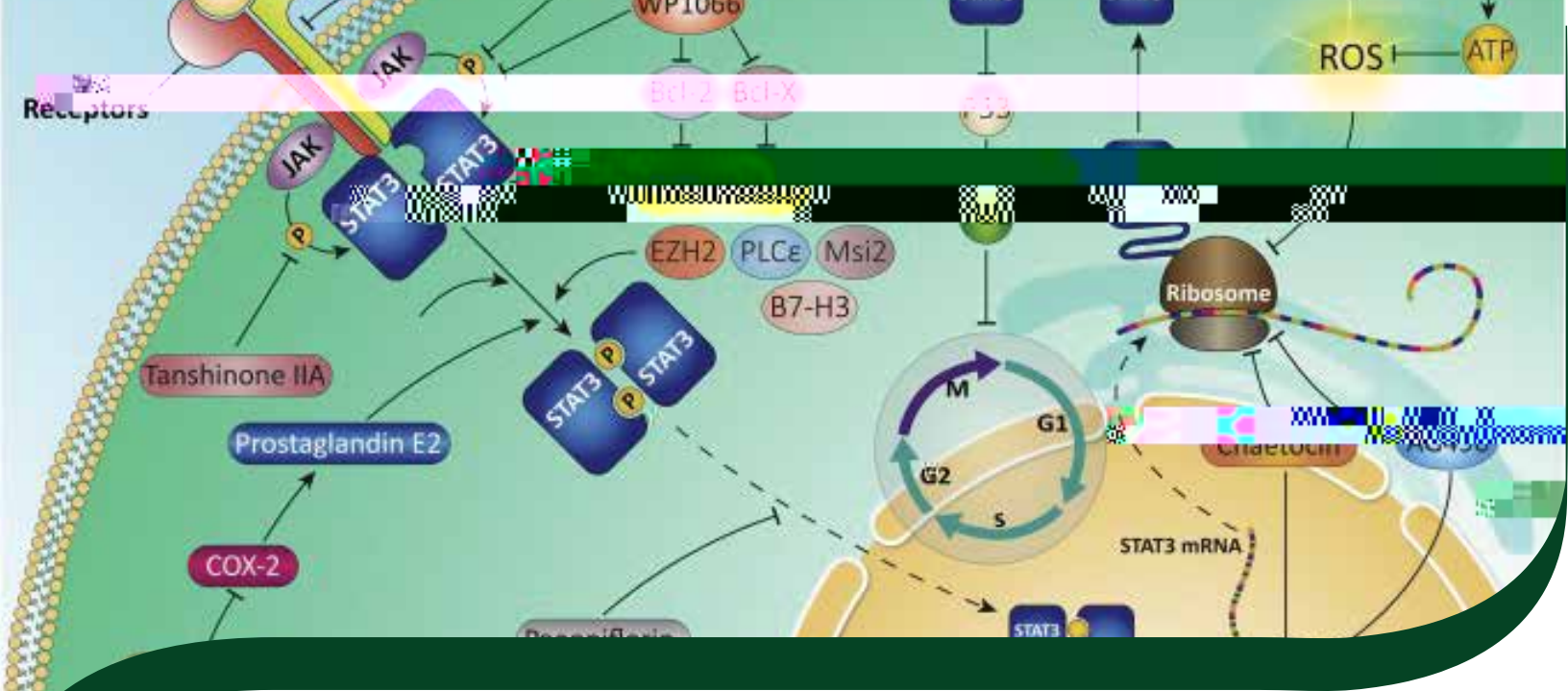


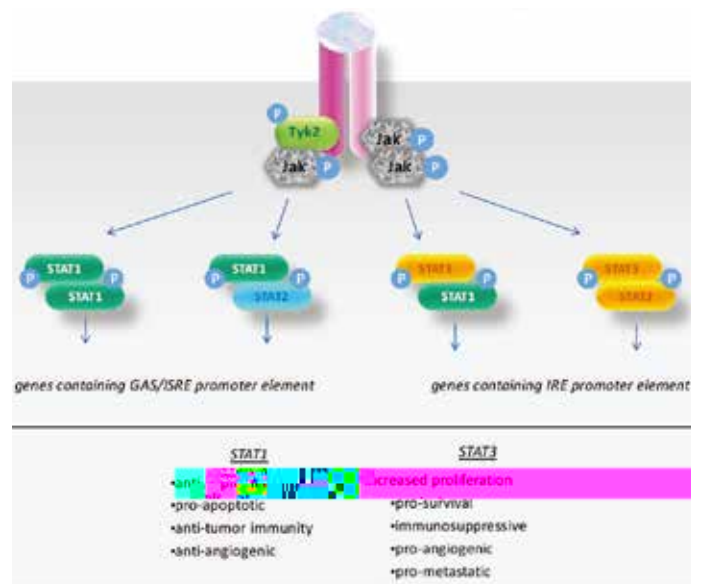


STAT3

STAT3

STAT3

STAT3

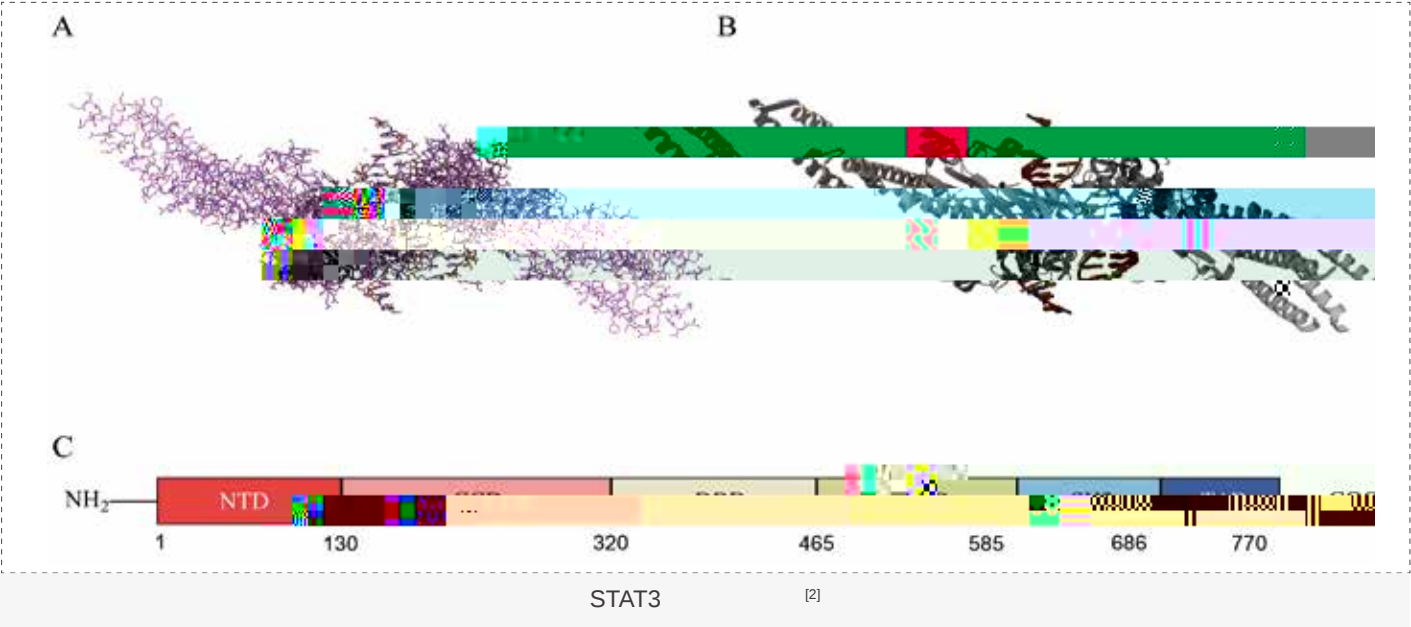


STAT3

STAT		STAT1	STAT2	STAT3	STAT4	STAT5A	STAT5B	STAT6	STAT3	
	STAT3α	STAT3β	STAT3α		STAT3β	STAT3α	C	55		STAT3β
STAT3α		DNA		STAT3				STAT3		17
		770								



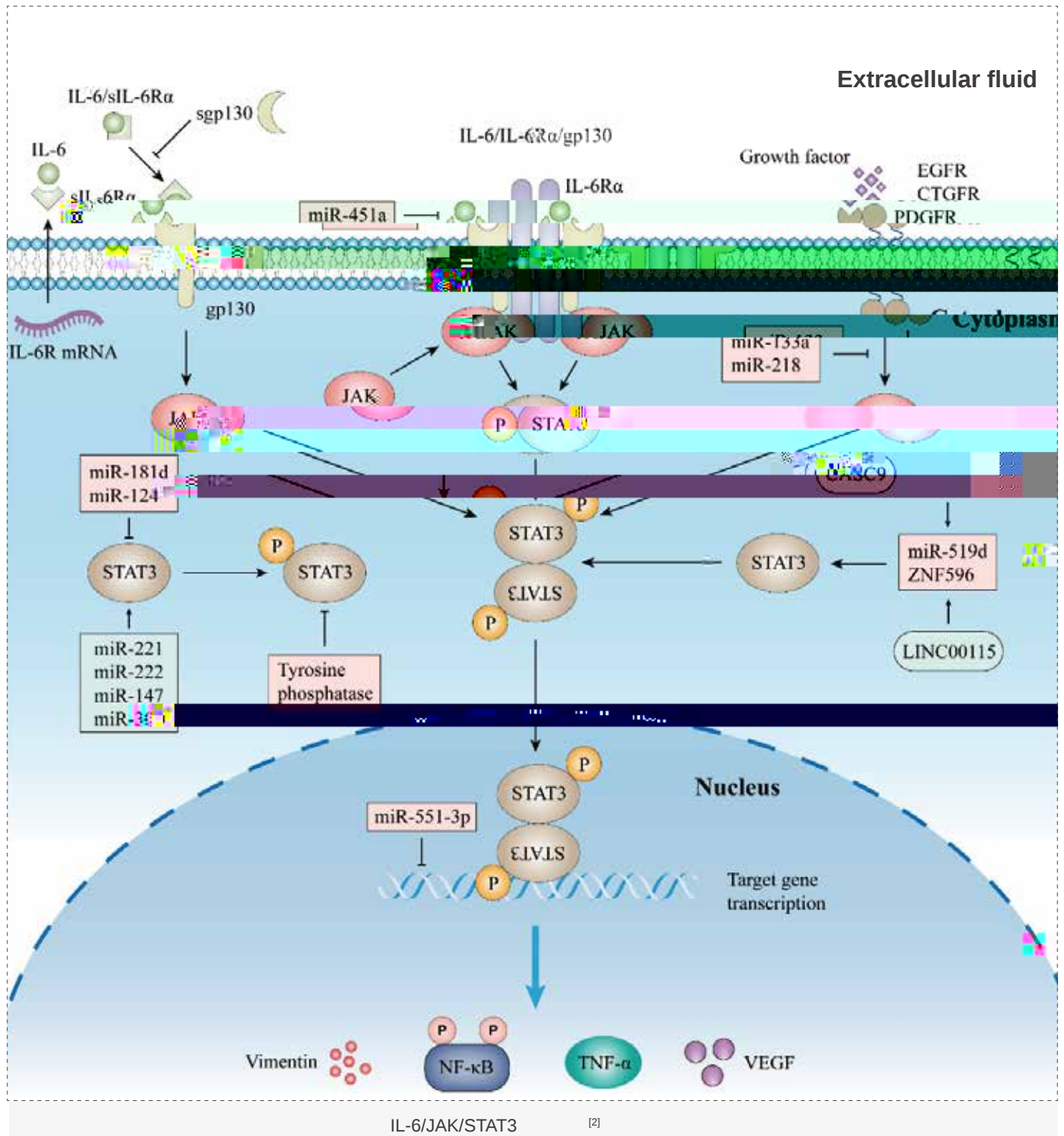
●	N	(NTD)	STAT							
●		(CCD)		STAT3						
●	DNA	(DBD)			DNA					
●		(LD)	DBD	SH2	DNA					
●	SRC	2	(SH2)		STAT		STAT3			
			STAT3							
●		(TAD)								



STAT3

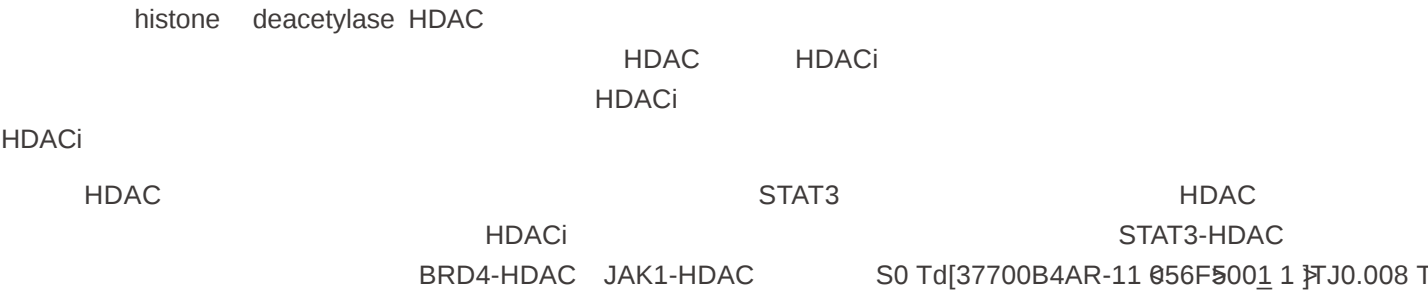
JAK/STAT										
(IL)		IL-6		TME	IL-6					
		IL-6			TME					
IL-6				IL-6	JAK/STAT3					JAK
IL-6/JAK/STAT3										JAK
JAK1	JAK2	JAK3	TYK2	JAK3						

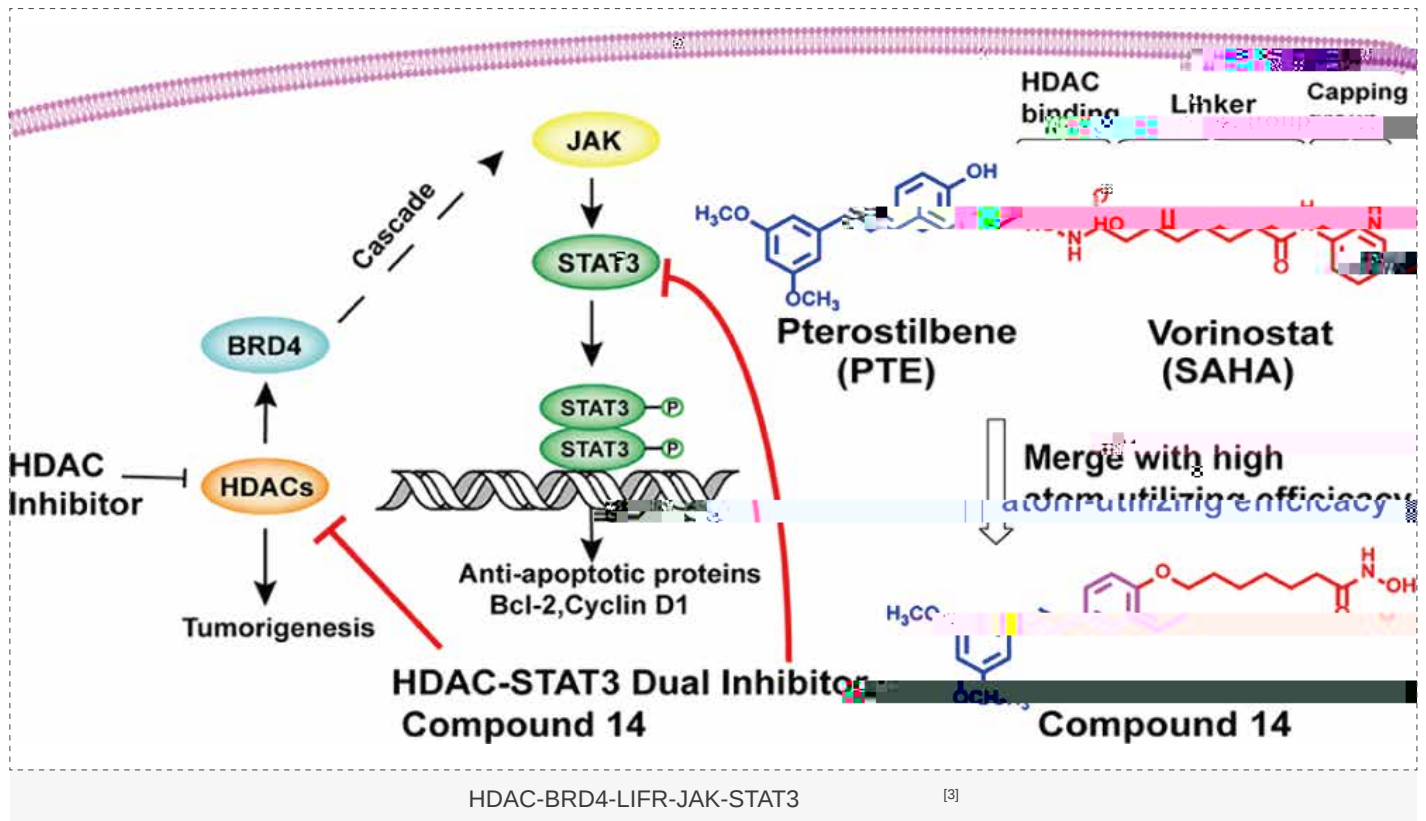
(JH1-7)		JH1	JH2	SH2	FERM		
IL-6	IL-6	IL-6	α (IL-6R α)	gp130		IL-6/IL-6R/gp130	STAT3
JAK	JAK		gp130				STAT3
	gp130	JAK	STAT3	705			STAT3



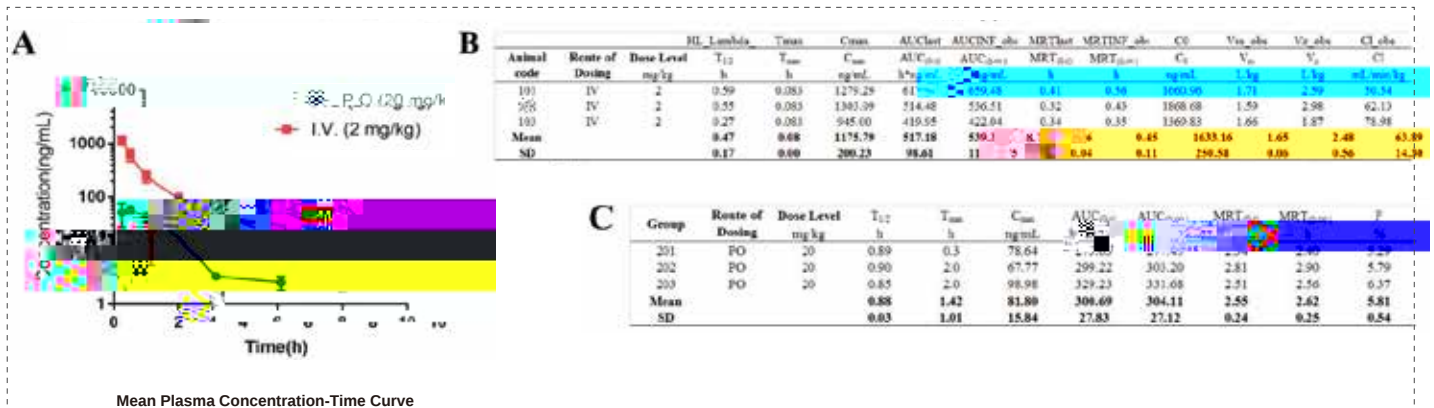
STAT3

🏆 STAT3-HDAC





Compound 14	HDAC	IC ₅₀	23.15 nM	Compound 14	STAT3	K _D =33 nM	
HCT116	IC ₅₀	1.07 μM	Compound 14	MDA-MB-231	IC ₅₀	0.78 μM	
Compound 14			Compound 14				
53%	30 mg/kg		64%	Compound 14	15 mg/kg		54%
Compound 14			62%	Compound 14			
Compound 14							
Compound 14	PK		Compound 14	SD			(PK)
(CI)	63.89 mL/min/kg		Compound 14	2 mg/kg	V _{ss}		1.65 L/kg
5.81%	T _{1/2}	0.88h	T _{1/2}	0.47h	Compound 14	20 mg/kg	F



Pharmacokinetics of Compound 14 in SD rats^[3]

YY201

2023 7

STAT3

—YY201

NMPA

AI

STAT3

“YY201”

First-in-class

STAT3

STAT3

Tyr705

Ser727

YY201 STAT3

STAT3

1-10

1-10 nM

YY201

STAT3

STAT3

STAT3

STAT3

PROTAC

STAT3-PROTAC

STAT3

STAT3

STAT3

[1]Courtney Nicholas and Gregory B. Lesinski. The Jak-STAT Signal Transduction Pathway in Melanoma. Breakthroughs in Melanoma Research.

[2]Xin Li, et al. STAT3 Inhibitors: A Novel Insight for Anticancer Therapy of Pancreatic Cancer. Biomolecules. 2022 Oct 9;12(10):1450. doi: 10.3390/biom12101450.

[3]Yuhao Ren, et al. Discovery of STAT3 and Histone Deacetylase (HDAC) Dual-Pathway Inhibitors for the Treatment of Solid Cancer. J Med Chem. 2021 Jun 10;64(11):7468-7482. doi: 10.1021/acs.jmedchem.1c00136.

[4]Sailan Zou, et al. Targeting STAT3 in Cancer Immunotherapy. Mol Cancer. 2020 Sep 24;19(1):145.DOI: 10.1186/s12943-020-01258-7.