

## The building blocks of organoid research

Organoid models include three-dimensional (3D) cell culture systems that closely resemble *in vivo* organs or tissues. These 3D systems reproduce the complex spatial morphology of a differentiated tissue to allow biologically relevant cell–cell and cell–matrix interactions, ideally sharing similar physiological responses with *in vivo* differentiated tissues.

The major focus of future organoid studies, beside the study of developmental processes, will most likely be in drug testing and modeling of diseases such as developmental disorders, genetic conditions, cancer, and degenerative disorders.

The hope is that organoids are another step in the long way towards *in vitro* construction of tissues and organs for transplantation.

Organoids	Proteins and growth factors	References
Brain organoids	BDNF	Lancaster MA and Knoblich JA (2014) Nat Protoc 9:2329.
	DKK-1	Li Y et al. (2017) Cell Stem Cell 20:1.
	EGF	Mariani J et al. (2015) Cell 162:375.
	FGF-basic	
	GDNF	
	Noggin	
Germinal center-like organoids	IL-4	Purwada A and Singh A (2017) Nat Protoc 12:168.
Inner ear organoids	BMP-4	Koehler KR and Hashino E (2014) Nat Protoc 9:1229.
	FGF-basic	
Kidney organoids	Activin A	Takasato M et al. (2014) Nat Cell Biol 16:118.
	BMP-2	Xia Y et al. (2013) Nat Cell Biol 15:1507.
	BMP-4	Xia Y et al. (2014) Nat Protoc 9:2693.
	BMP-7	
	FGF-9	
	FGF-basic	
Liver organoids	BMP-4	Broutier L et al. (2016) Nat Protoc 11:1724.
	EGF	Sato T and Clevers H (2015) Cell 161:1700.
	FGF-10	Takebe T et al. (2013) Nature 499:481.
	FGF-basic	
	HGF	
	Noggin	
	R-spondin-1	
	Wnt-3a	

Organoids	Proteins and growth factors	References
Lung organoids	Activin A	Dye BR et al. (2015) <i>eLife</i> 4:e05098.
	FGF-basic	
	FGF-4	
	Noggin	
Mammary organoids	EGF	Jamieson PR et al. (2017) Development 144:1065.
	FGF-10	Jardé T et al. (2016) Nat Commun 7:13207.
	FGF-basic	
	Heregulin β-1	
	Noggin	
	Prolactin	
	R-spondin-1	
	R-spondin-2	
	Wnt-3a	
	EGF	Boj SF et al. (2015) Cell 160:324.
	FGF-10	Broutier L et al. (2016) Nat Protoc 11:1724.
Pancreatic organoids	Noggin	Sato T and Clevers H (2015) Cell 161:1700.
	R-spondin-1	
	Wnt-3a	
Prostate organoids	Activin A	Drost J et al. (2016) Nat Protoc 11:347.
	EGF	Karthaus WR et al. (2014) Cell 159:163.
	FGF-10	
	FGF-basic	
	Noggin	
	R-spondin-1	
Retina organoids	Sonic hedgehog (Shh)	Nakano T et al. (2012) Cell Stem Cell 10:771.
	Wnt-3a	
Small intestinal and colonic organoids	EGF	Jung P et al. (2011) Nat Med 17:1225.
	Noggin	Mahe M et al. (2013) Curr Protoc Mouse Biol 3:217.
	R-spondin-1	Sato T et al. (2011) Gastroenterology 141:1762.
	Wnt-3a	Sato T and Clevers H (2009) Nature 459:262.
		Sato T and Clevers H (2015) Cell 161:1700.
Stomach organoids	EGF	Bartfeld S et al. (2015) Gastroenterology 148:126.
	FGF-10	Mahe M et al. (2013) Curr Protoc Mouse Biol 3:217.
	Noggin	Sato T and Clevers H (2015) Cell 161:1700.
	R-spondin-1	
	Wnt-3a	

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