Supplementary Information for

## Cascaded filter deterministic lateral displacement microchips for isolation and molecular analysis of circulating tumor cells and fusion cells

Zongbin Liu,\*<sup>a</sup> Yuqing Huang,<sup>b</sup> Wenli Liang,<sup>c</sup> Jing Bai,<sup>a</sup> Hongtao Feng,<sup>b</sup> Zhihao Fang,<sup>b</sup> Geng Tian,<sup>c</sup>

Yanjuan Zhu,<sup>d</sup> Haibo Zhang,<sup>d</sup> Yuanxiang Wang,<sup>e</sup> Aixue Liu,\*<sup>c</sup> and Yan Chen\*<sup>b</sup>

<sup>a</sup>Shenzhen Zigzag Biotechnology Co., Ltd., Shenzhen, 518107, China

<sup>b</sup>CAS Key Laboratory of Health Informatics, Shenzhen Institutes of Advanced Technology, Chinese Academy of Sciences, Shenzhen, 518055, China

<sup>c</sup>Tumor Department, Shenzhen Second People's Hospital, The First Affiliated Hospital of Shenzhen University, Shenzhen, 518035, China

<sup>d</sup>Department of Oncology, Guangdong Provincial Hospital of Traditional Chinese Medicine, The Second Clinical Medical College of Guangzhou University of Chinese Medicine, Guangzhou, 510120, China

<sup>e</sup>Department of Cardiothoracic Surgery, Shenzhen Children's Hospital, Shenzhen, 518038, China

Corresponding author: yan.chen@siat.ac.cn

zongbin.liu@zigbio.com

liuaixue\_76@sina.com

# S1. Measurement of critical separation size in deterministic lateral displacement (DLD) structures.



**Fig. S1.** The performance of microbeads separation in rectangular micro-post chip and filter-DLD chip. (A) Schematic chip design with two inlets and two outlets. (B) Zigzag mode movement of 12  $\mu$ m fluorescent microbeads in the rectangular micro-post chip. Scale bar, 150  $\mu$ m. (C-D) Images of collected small beads (C) and large beads (D) from the two outlets of the rectangular micro-post chip. The diameter range of input beads is from 9  $\mu$ m to 18  $\mu$ m. Scale bar, 50  $\mu$ m. (E) Bumping mode movement of 12  $\mu$ m fluorescent microbeads in the filter-DLD chip. Scale bar, 150  $\mu$ m. (F-G) Images of collected small beads (F) and large beads (G) from the two outlets of the filter-DLD chip. The diameter range of input beads is from 9  $\mu$ m to 18  $\mu$ m. Scale bar, 50  $\mu$ m. From the collected microbeads, the critical sizes (D<sub>c</sub>) of filter-DLD and rectangular micro-post DLD were estimated to be around 10  $\mu$ m and 15  $\mu$ m, respectively. S2. Schematic chip designs of deterministic lateral displacement (DLD) structures.



**Fig. S2.** Schematic chip designs of deterministic lateral displacement (DLD) structures. (A) Rectangular micro-post DLD design, (B) Filter-DLD design. The two structures have the same micro-post side length, gap width, and critical separation size ( $D_c$ ). Red dotted lines indicate inclination angles of micro-posts.

#### **S3.** Evaluation of white blood cells removal rate.

Healthy blood	Count of retained WBCs from CTCs outlet			
samples	(WBCs number in 1 mL blood)			
#1	47 WBCs/mL			
#2	79 WBCs/mL			
#3	134 WBCs/mL			
#4	213 WBCs/mL			
#5	232 WBCs/mL			

**Table S1.** Count of recovered white blood cells (WBCs) in healthy blood samples.

Characteristic	Evaluable	CTC prevalence in 91 blood samples			
	patients, n=35	CTCs=0, n(%)	1≦CTCs<5, n(%)	CTCs≧5, n(%)	
Age, years					
Average	61	N/A	N/A	N/A	
Range	27-89	N/A	N/A	N/A	
Gender					
Male	15	N/A	N/A	N/A	
Female	20	N/A	N/A	N/A	
Tumor histology					
AC	23	11(18%)	15(24%)	36(58%)	
SCC	12	1(3%)	13(45%)	15(52%)	
Disease status					
SD	N/A	12(23%)	24(45%)	17(32%)	
PD	N/A	1(3%)	3(8%)	34(89%)	

#### S4. Evaluation of CTCs from NSCLC patient samples.

Abbreviations: NSCLC, non-small-cell lung cancer; AC, adenocarcinoma; SCC, squamous cell carcinoma; SD, stable disease; PD, progressive disease; N/A, not applicable.

**Table S2.** Demographics of 35 stage IV NSCLC patients and CTCs prevalence in 91 blood samples.

### **Details of Supplementary Videos**

Movie S1. Passage of blood cells through filter deterministic lateral displacement structures.

Movie S2. Passage of a 12  $\mu m$  microbead in filter-DLD and rectangular micro-post DLD structures.

**Movie S3.** Passage of cells with different sizes in filter-DLD and rectangular micro-post DLD structures.