Supplementary

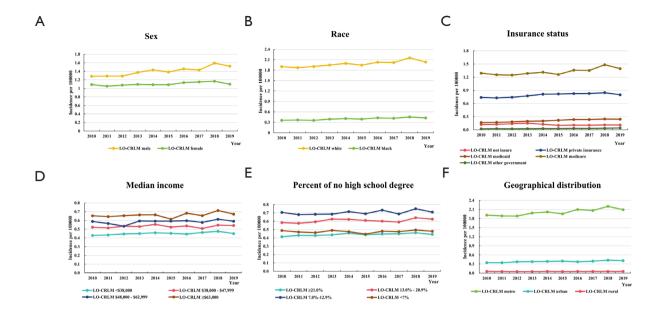


Figure S1 Annual incidence of LO-CRLM among sub-population stratified by demographics from 2010 to 2019. (A) Sex. (B) Race. (C) Insurance. (D) Incoming. (E) Percent of no high school degree. (F) Geographical distribution. LO-CRLM, late-onset colorectal liver metastases.

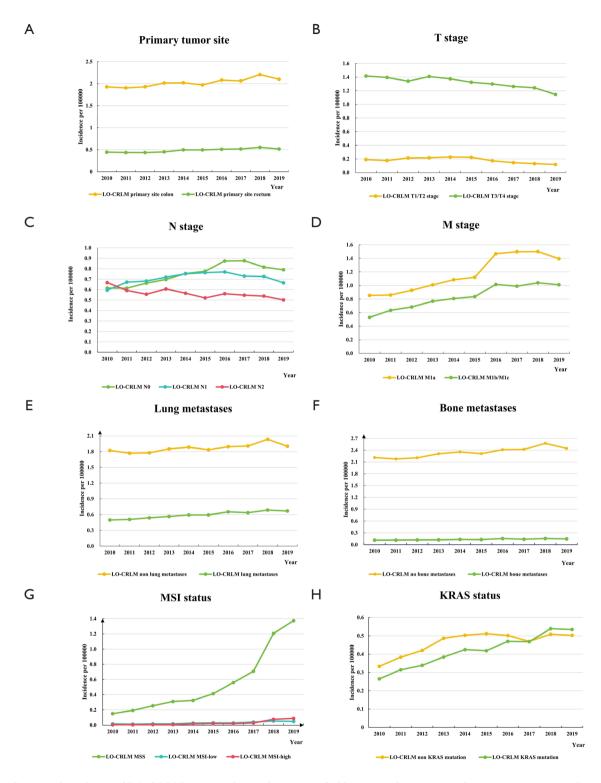


Figure S2 Annual incidence of LO-CRLM among sub-population stratified by tumor characteristics from 2010 to 2019. (A) Primary tumor sites. (B) T stage. (C) N stage. (D) M stage. (E) Lung metastases. (F) Bone metastases. (G) MSI status. (H) KRAS status. LO-CRLM, late-onset colorectal liver metastases; microsatellite instability, MSI; Kirsten rat sarcoma viral oncogene homolog, KRAS.

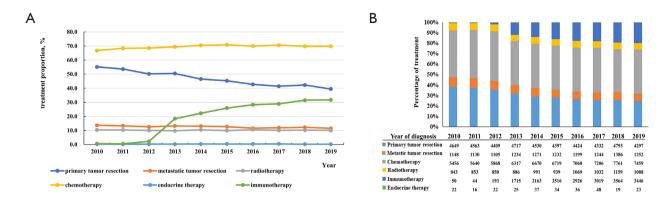


Figure S3 Treatment trends of LO-CRLM patients from 2010 to 2019. (A) Trends in proportion of treatment modalities used by LO-CRLM patients from 2010 to 2019. (B) Temporal trends for treatment from 2010 to 2019. LO-CRLM, late-onset colorectal liver metastases.

demographics, tumor characteristics and treatment	1, EO-CRLM and LO-CRLM in the whole population and sub-population stratified 1-, 3- and 5-year OS rate (%)		
Variables	CRLM	EO-CRLM	LO-CRLM
None stratifying	59.1, 26.3, 15.0	77.5, 39.1, 23.7	55.8, 24.0, 13.4
Charlson-Deyo score	62.2, 28.4, 16.4	78.3, 39.6, 24.1	58.7, 25.9, 14.7
1	54.5, 22.7, 12.3	76.6, 37.2, 21.6	52.4, 21.4, 11.5
≥2	40.4, 14.9, 7.3	57.4, 27.9, 16.8	39.5, 14.2, 6.7
Sex	60 6 07 0 14 9	76 7 20 7 20 0	E7 0 04 0 10 4
Female Male	60.6, 27.0, 14.8 57.2, 25.5, 15.1	76.7, 38.7, 23.0 78.6, 39.5, 24.4	57.8, 24.9, 13.4 53.3, 22.9, 13.4
Race	,,	, ,	2212, 2212, 1211
White	59.1, 26.6, 15.0	78.6, 40.8, 24.6	55.7, 24.2, 13.4
Black	56.9, 22.6, 13.1	71.3, 29.7, 18.6	54.2, 21.3, 12.1
Others Rural-Urban Grouping	66.4, 32.1, 19.7	81.6, 41.8, 25.6	62.3, 29.5, 18.0
Metro areas	59.0, 26.4, 15.1	77.5, 39.2, 23.8	55.7, 24.0, 13.6
Urban areas	57.9, 24.2, 12.9	76.1, 35.8, 20.6	55.1, 22.4, 11.7
Rural areas	55.0, 22.6, 11.2	75.0, 31.1, 17.3	52.4, 21.6, 10.4
Facility type Community Cancer Program	53.5, 21.5, 11.9	74.7, 34.3, 18.7	51.3, 20.2, 11.2
Comprehensive Community Cancer Program	55.4, 22.6, 11.8	74.9, 35.6, 20.1	53.1, 21.1, 10.8
Academic/Research Program	64.3, 31.0, 19.1	79.9, 41.7, 26.9	61.7, 29.2, 17.7
Integrated Network Cancer Program	56.6, 25.3, 14.2	77.0, 39.4, 24.3	54.1, 23.5, 12.9
Percent of no high school degree ≥21%	58.1, 25.5, 15.0	73.4, 35.6, 22.9	55.2, 23.6, 13.5
13.0-20.9%	57.2, 24.4, 13.5	75.5, 35.6, 21	54.2, 22, 12.3
7.0-12.9%	58.7, 25.8, 14.0	78.6, 39.9, 22.9	55.3, 23.3, 12.5
<7.0%	61.0, 28.2, 16.7	81.1, 43.7, 27.2	57.2, 25.3, 14.8
Median income <\$38,000	55.6, 22.7, 12.6	72.9, 33.7, 19.8	52.7, 20.9, 11.4
\$38,000-\$47,999	57.1, 24.1, 13.2	72.9, 33.7, 19.8 75.1, 35.7, 20.9	54.0, 22.1, 11.9
\$48,000-\$62,999	58.7, 26.3, 15.0	77.5, 39.1, 24.3	55.3, 24.0, 13.3
≥\$63,000	62.0, 29.0, 16.9	81.3, 43.8, 26.6	58.4, 26.2, 15.1
nsurance status	FO 6 05 0 44 0	60 6 07 0 14 7	FG 0 044 44 -
Not Insured Private Insurance/Managed Care	59.6, 25.0, 14.6 72.8, 36.1, 21.3	68.6, 27.9, 14.7 81.3, 42.8, 26.4	56.8, 24.1, 14.5 69.6, 33.6, 19.4
Medicaid Medicaid	62.6, 27.7, 16.3	71.6, 33.1, 19.7	59.1, 25.6, 14.9
Medicare	47.1,18.0, 9.5	62.1, 26.2, 14.6	46.8, 17.9, 9.4
Other Government	62.4, 30.1, 15.1	83.0, 42.9, 25.7	58.0, 27.3, 12.8
Diagnosis year 2010-2012	57.0, 23.6, 12.9	75.4, 35.3, 20.7	53.9, 21.6, 11.5
2013-2015	58.7, 25.9, 15.0	77.0, 37.3, 22.8	55.5, 24.0, 13.6
2016-2019	61.2, 29.3, 16.4	79.5, 44.0, 25.0	57.6, 26.4, 14.8
Primary tumor site			
Primary sites in colon	56.7, 24.7, 14.1	76.0, 37.4, 22.9	53.5, 22.6, 12.7
Primary sites in rectum CEA level	68.4, 32.7, 18.1	81.9, 43.8, 25.7	65.1, 30.0, 16.4
<5 ng/mL	70.1, 38.5, 25.1	83.3, 51.9, 36.7	67.4, 35.7, 22.7
≥5 ng/mL	57.3, 22.9, 12.1	75.8, 34.5, 19.4	53.9, 20.8, 10.7
Grade			
Well differentiation Moderate differentiation	68.6, 34.6, 20.7 68.7, 33.0, 19.0	86.1, 50.0, 33.9 84.4, 46.0, 28.4	65.3, 31.7, 18.2 65.5, 30.4, 17.1
Poor differentiation	49.3, 19.4, 11.1	65.1, 27.9, 16.4	46.5, 17.9, 10.2
Undifferentiation	49.0, 18.2, 10.3	66.8, 26.1, 16.5	46.0, 16.9, 9.2
umor size			
<5cm	68.6, 33.4, 19.3	84.4, 47.3, 28.6	65.5, 30.8, 17.6
≥5cm ymph invasion	62.0, 28.5, 17.0	78.2, 40.2, 26.2	59.0, 26.3, 15.3
No	73.3, 40.4, 25.3	87.0, 53.7, 36.2	70.7, 37.8, 23.1
Yes	66.2, 30.3, 16.9	82.5, 43.5, 25.4	63.0, 27.7, 15.3
Perineural invasion			
No Yes	67.9, 34.1, 20.8 67.7, 30.7, 16.5	83.8, 47.8, 31.5 83.2, 43.9, 24.5	65.1, 31.6, 18.9 64.1, 27.7, 14.7
stage	07.7, 30.7, 10.3	00.2, 40.3, 24.0	04.1, 27.1, 14.1
T1	52.5, 22.8, 12.5	73.1, 36.3, 21.2	48.7, 20.3, 10.9
T2	77.8, 46.9, 31.7	90.0, 59.9, 44.9	74.8, 43.7, 28.5
T3	74.6, 39.2, 23.5	87.6, 52.5, 33.3	71.9, 36.5, 21.5
T4 I stage	59.1, 24.1, 13.0	76.1, 35.1, 19.6	56.1, 22.1, 11.8
N0	58.1, 28.0, 16.8	78.9, 41.9, 27.3	55.1, 25.9, 15.2
N1	66.3, 32.1, 18.5	81.8, 44.7, 27.5	63.2, 29.6, 16.7
N2	63.0, 25.8, 14.1	79.2, 39.2, 21.9	59.4, 22.8, 12.3
1 stage M1a	66.8, 34.2, 20.4	84.5, 50.1, 32.1	63.6, 31.2, 18.3
M1b	50.0, 16.5, 7.9	69.0, 25.0, 12.8	46.4, 15.0, 7.0
M1c	49.8, 14.7, not reach	69.3, 25.9, not reach	45.9, 12.6, not reach
Bone metastases			
No Yes	60.6, 27.4, 15.6 36.3, 8.9, 4.1	79.0, 40.5, 24.6 51.8, 12.4, 5.6	57.3, 25.0, 14.0 33.8, 8.4, 3.8
res Brain metastases	oo.o, o.a, 4.1	U1.0, 12.4, U.U	oo.o, o.4, o.d
No	59.7, 26.7, 15.2	78.0, 39.5, 23.9	56.4, 24.3, 13.6
Yes	30.3, 7.8, 4.6	46.8, 14.1, 9.4	27.5, 6.8, 3.8
ung metastases	00.0.00.0	00.0.40.0.==	F0.0.07
No Yes	62.6, 30.0, 17.6 48.8, 14.8, 6.5	80.3, 43.6, 27.3 68.2, 23.1, 10.6	59.3, 27.4, 15.8 45.7, 13.5, 5.8
(RAS mutation	.5.5, 14.5, 5.5	55.2, 25.1, 10.0	.5., 10.0, 0.0
No	71.8, 34.3, 19.1	84.3, 46.6, 28.2	68.7, 31.2, 16.8
Yes	68.3, 25.5, 13.5	77.9, 31.8, 18.3	66.2, 24.1, 12.5
ASI status	70 6 05 0 00 0	00 7 45 0 07 0	60 5 00 0 10 0
MSI-high	72.6, 35.8, 20.6 73.4, 38.0, 23.1	83.7, 45.8, 27.9 86.1, 49.3, 31.9	69.5, 33.0, 18.6 70.6, 35.5, 21.1
MSI-low	64.9, 36.3, 25.9	79.6, 50.8, 40.1	60.9, 32.3, 21.8
rimary tumor site resection			
No	46.2, 14.6, 6.8	66.2, 23.1, 11.6	43.0, 13.3, 6.1
Yes Matastasas resection	72.6, 38.1, 23.0	86.9, 51.9, 33.1	69.7, 35.3, 21.0
No	55.3, 21.8, 11.6	73.9, 32.6, 18.8	52.3, 20.0, 10.4
Yes	82.2, 53.2, 34.9	92.0, 63.8, 42.3	79.5, 50.1, 32.8
Radiation			
No	57.3, 25.0, 14.2	76.6, 37.9, 23.0	54.1, 22.9, 12.7
Yes Chemotherapy	71.2, 35.7, 21.2	82.6, 46.2, 28.6	68.2, 32.9, 19.3

Immunotherapy
No 53.8, 24.0, 13.9 74.3, 37.6, 23.1 50.6, 21.8, 12.4

 ${\sf OS,\ overall\ survival;\ CRLM,\ colorectal\ liver\ metastases;\ EO-CRLM,\ early-onset\ colorectal\ liver\ metastases;\ LO-CRLM,\ late-onset\ colorectal\ liver\ metastases;\ LO-CRLM,\ late-onset\ liver\ metastases;\ late-onset\ liver\ metastases;\ late-onset\ liver\ metastases;\ late-onset\ liver\ metastases;\ late-onset\ liver\ late-onset\ la$

38.9, 18.0, 12.6

81.6, 41.2, 24.7

79.6, 35.3, 18.8 85.8, 42.8, 24.7 77.9, 33.2, 17.2

20.8, 8.4, 5.3

73.1, 32.9, 18.4

No

Yes

Yes

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19.8, 7.8, 4.9

71.1, 30.9, 17.0