

SUPPLEMENTARY MATERIALS

Clonal relationship and directionality of progression of synchronous endometrial and ovarian carcinomas in patients with DNA mismatch repair-deficiency associated syndromes

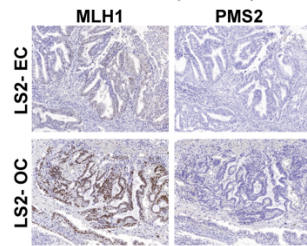
Moukarzel et al

Supplementary Figures S1-S3

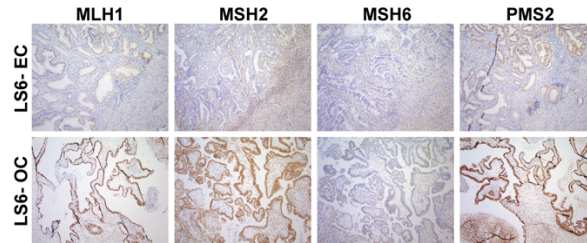
Supplementary Table S1

Supplementary Figure S1

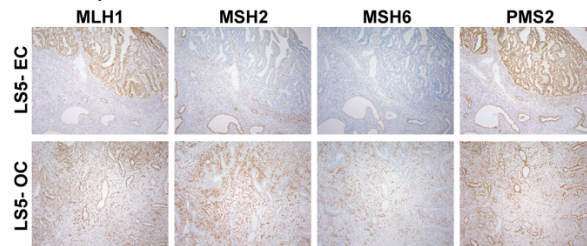
LS2-CMMRD: *PMS2* p.S8fs* (homozygous)



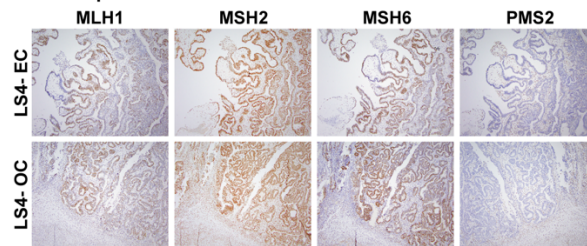
LS6: *MSH6* p.R1334P



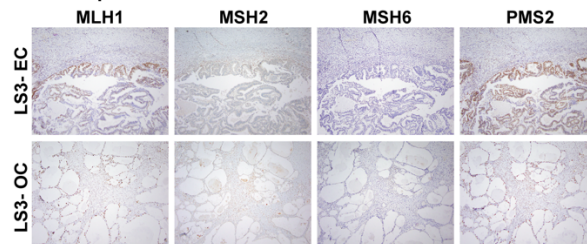
LS5: *MSH2* p.Y521*



LS4: *PMS2* p.Q30*



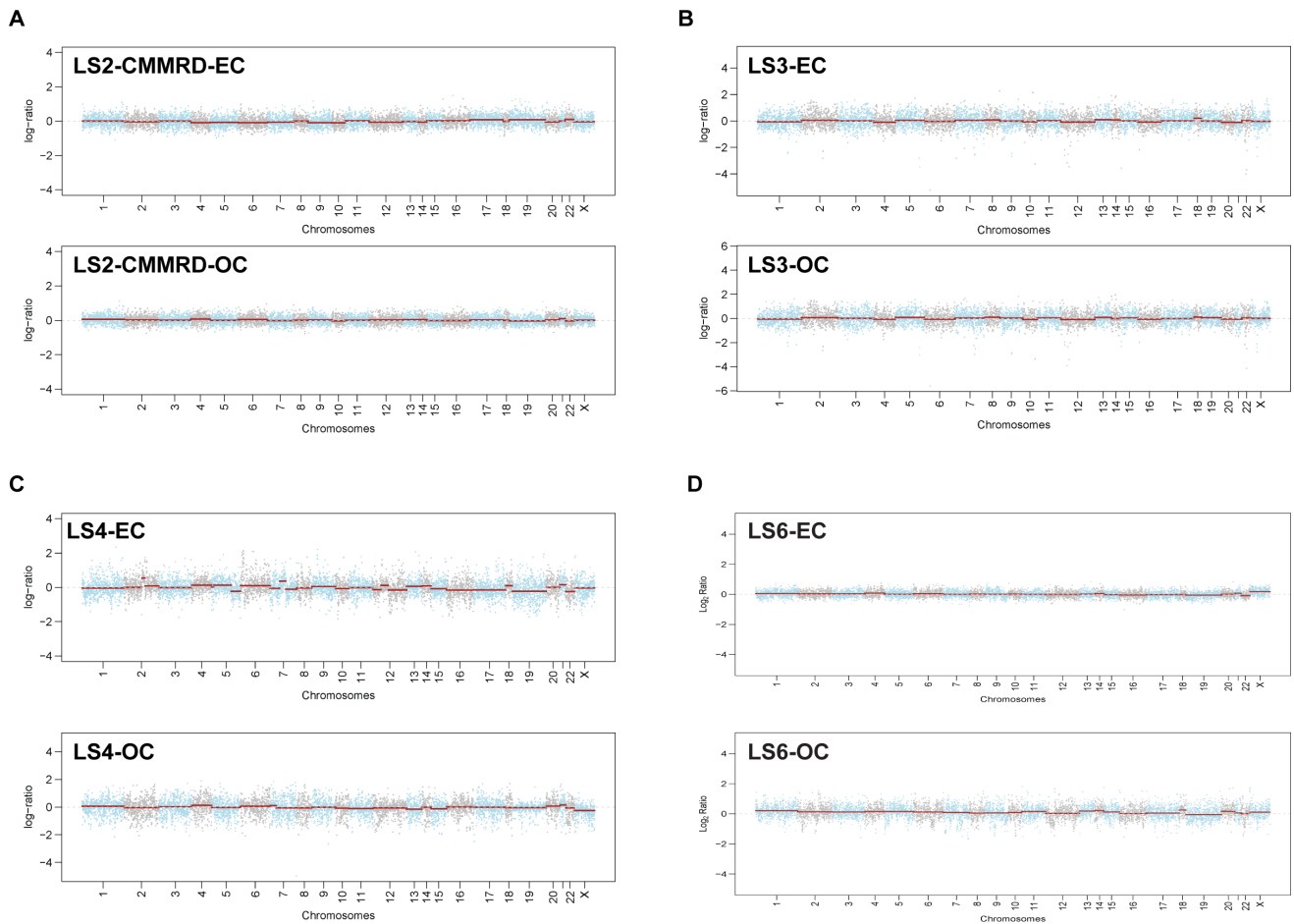
LS3: *MSH2* p.Q324FS*



Supplementary Figure S1: Immunohistochemical analysis of DNA mismatch repair protein expression in synchronous endometrial and ovarian cancers from patients with germline DNA mismatch repair gene mutations.

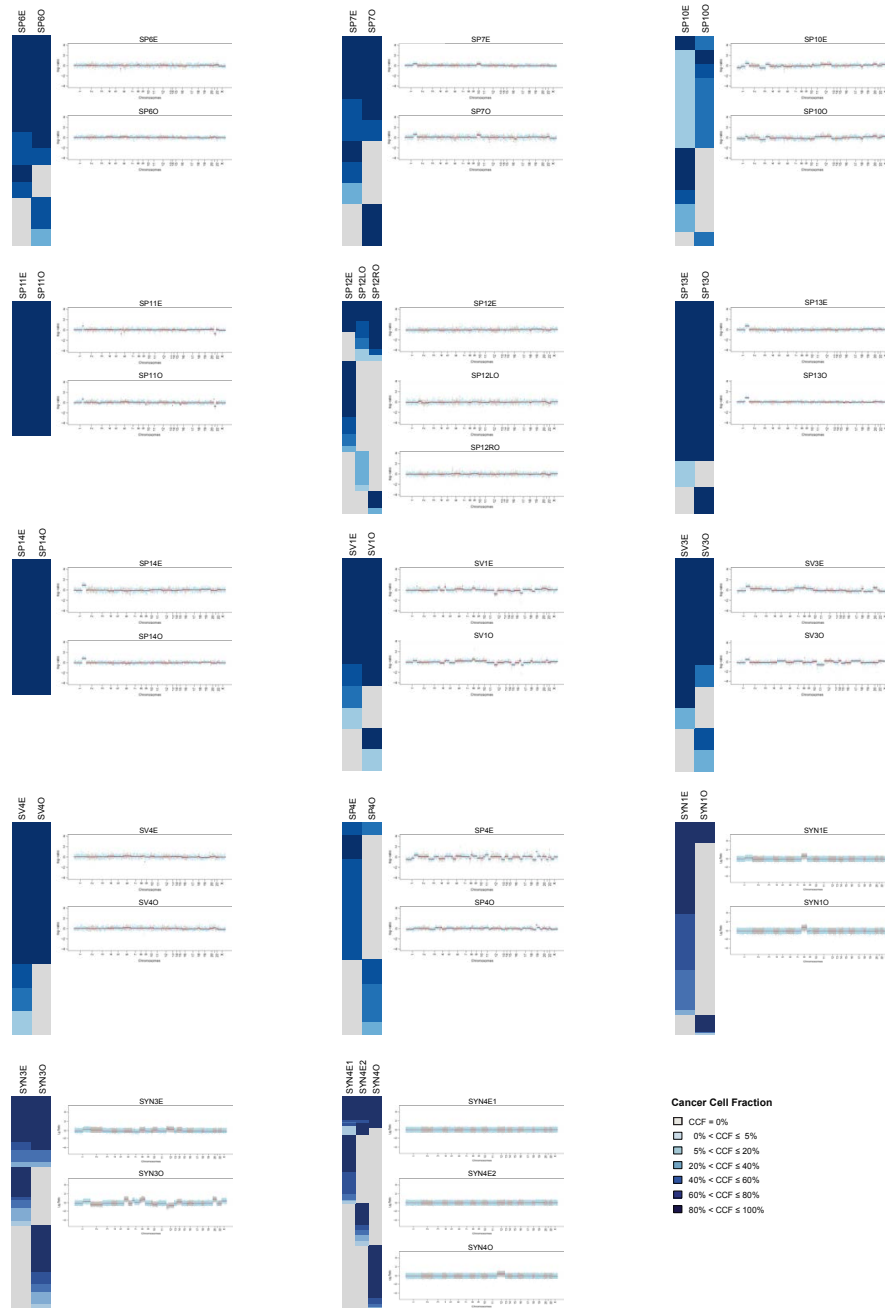
MLH1, MSH2, MSH6 and PMS2 protein expression in the synchronous endometrial (ECs) and ovarian cancers (OCs) from patients LS2-CMMRD, LS6, LS5, LS4 and LS3 harboring germline DNA mismatch repair mutations as indicated.

Supplementary Figure S2



Supplementary Figure 2: Chromosome plots of synchronous endometrial and ovarian cancers from patients with germline DNA mismatch repair mutations. Log₂-ratios are plotted on the y-axis and the genomic positions are plotted on the x-axis for the synchronous endometrial cancer (EC) and ovarian cancer (OC) cases (A) LS2-CMMRD, (B) LS3, (C) LS4 and (D) LS6.

Supplementary Figure S3



Supplementary Figure S3: Clonal composition analysis of sporadic synchronous endometrioid endometrial and endometrioid ovarian cancers.

Re-analysis of sporadic synchronous endometrial (ECs) and ovarian cancers (OCs) (from Schultheis et al, JNCI 2016) subjected to whole-exome sequencing (SYN1, SYN3, SYN4) or targeted MSK-IMPACT sequencing (SP6, SP7, SP10, SP11, SV12, SP13, SP14, SV1, SV3, SV4, SP4). ECs and OCs of a given cases were clonally related, however for these cases, chronology of development could not be inferred based on clonal composition analysis. For each case, cancer cell fractions of somatic mutations identified in the EC and synchronous OC (left), color coded according to the legend, and copy number plots with the Log_2 -ratios on the y-axis and the genomic positions on the x-axis are depicted.

Gene	Accession	Length (bp)	GC (%)	GC3 (%)	GC4 (%)	GC5 (%)	GC6 (%)	GC7 (%)	GC8 (%)	GC9 (%)	GC10 (%)	GC11 (%)	GC12 (%)	GC13 (%)	GC14 (%)	GC15 (%)	GC16 (%)	GC17 (%)	GC18 (%)	GC19 (%)	GC20 (%)	GC21 (%)	GC22 (%)	GC23 (%)	GC24 (%)	GC25 (%)	GC26 (%)	GC27 (%)	GC28 (%)	GC29 (%)	GC30 (%)	GC31 (%)	GC32 (%)	GC33 (%)	GC34 (%)	GC35 (%)	GC36 (%)	GC37 (%)	GC38 (%)	GC39 (%)	GC40 (%)	GC41 (%)	GC42 (%)	GC43 (%)	GC44 (%)	GC45 (%)	GC46 (%)	GC47 (%)	GC48 (%)	GC49 (%)	GC50 (%)	GC51 (%)	GC52 (%)	GC53 (%)	GC54 (%)	GC55 (%)	GC56 (%)	GC57 (%)	GC58 (%)	GC59 (%)	GC60 (%)	GC61 (%)	GC62 (%)	GC63 (%)	GC64 (%)	GC65 (%)	GC66 (%)	GC67 (%)	GC68 (%)	GC69 (%)	GC70 (%)	GC71 (%)	GC72 (%)	GC73 (%)	GC74 (%)	GC75 (%)	GC76 (%)	GC77 (%)	GC78 (%)	GC79 (%)	GC80 (%)	GC81 (%)	GC82 (%)	GC83 (%)	GC84 (%)	GC85 (%)	GC86 (%)	GC87 (%)	GC88 (%)	GC89 (%)	GC90 (%)	GC91 (%)	GC92 (%)	GC93 (%)	GC94 (%)	GC95 (%)	GC96 (%)	GC97 (%)	GC98 (%)	GC99 (%)	GC100 (%)
Gene 1	Accession 1	Length 1	GC 1	GC3 1	GC4 1	GC5 1	GC6 1	GC7 1	GC8 1	GC9 1	GC10 1	GC11 1	GC12 1	GC13 1	GC14 1	GC15 1	GC16 1	GC17 1	GC18 1	GC19 1	GC20 1	GC21 1	GC22 1	GC23 1	GC24 1	GC25 1	GC26 1	GC27 1	GC28 1	GC29 1	GC30 1	GC31 1	GC32 1	GC33 1	GC34 1	GC35 1	GC36 1	GC37 1	GC38 1	GC39 1	GC40 1	GC41 1	GC42 1	GC43 1	GC44 1	GC45 1	GC46 1	GC47 1	GC48 1	GC49 1	GC50 1	GC51 1	GC52 1	GC53 1	GC54 1	GC55 1	GC56 1	GC57 1	GC58 1	GC59 1	GC60 1	GC61 1	GC62 1	GC63 1	GC64 1	GC65 1	GC66 1	GC67 1	GC68 1	GC69 1	GC70 1	GC71 1	GC72 1	GC73 1	GC74 1	GC75 1	GC76 1	GC77 1	GC78 1	GC79 1	GC80 1	GC81 1	GC82 1	GC83 1	GC84 1	GC85 1	GC86 1	GC87 1	GC88 1	GC89 1	GC90 1	GC91 1	GC92 1	GC93 1	GC94 1	GC95 1	GC96 1	GC97 1	GC98 1	GC99 1	GC100 1
Gene 2	Accession 2	Length 2	GC 2	GC3 2	GC4 2	GC5 2	GC6 2	GC7 2	GC8 2	GC9 2	GC10 2	GC11 2	GC12 2	GC13 2	GC14 2	GC15 2	GC16 2	GC17 2	GC18 2	GC19 2	GC20 2	GC21 2	GC22 2	GC23 2	GC24 2	GC25 2	GC26 2	GC27 2	GC28 2	GC29 2	GC30 2	GC31 2	GC32 2	GC33 2	GC34 2	GC35 2	GC36 2	GC37 2	GC38 2	GC39 2	GC40 2	GC41 2	GC42 2	GC43 2	GC44 2	GC45 2	GC46 2	GC47 2	GC48 2	GC49 2	GC50 2	GC51 2	GC52 2	GC53 2	GC54 2	GC55 2	GC56 2	GC57 2	GC58 2	GC59 2	GC60 2	GC61 2	GC62 2	GC63 2	GC64 2	GC65 2	GC66 2	GC67 2	GC68 2	GC69 2	GC70 2	GC71 2	GC72 2	GC73 2	GC74 2	GC75 2	GC76 2	GC77 2	GC78 2	GC79 2	GC80 2	GC81 2	GC82 2	GC83 2	GC84 2	GC85 2	GC86 2	GC87 2	GC88 2	GC89 2	GC90 2	GC91 2	GC92 2	GC93 2	GC94 2	GC95 2	GC96 2	GC97 2	GC98 2	GC99 2	GC100 2
Gene 3	Accession 3	Length 3	GC 3	GC3 3	GC4 3	GC5 3	GC6 3	GC7 3	GC8 3	GC9 3	GC10 3	GC11 3	GC12 3	GC13 3	GC14 3	GC15 3	GC16 3	GC17 3	GC18 3	GC19 3	GC20 3	GC21 3	GC22 3	GC23 3	GC24 3	GC25 3	GC26 3	GC27 3	GC28 3	GC29 3	GC30 3	GC31 3	GC32 3	GC33 3	GC34 3	GC35 3	GC36 3	GC37 3	GC38 3	GC39 3	GC40 3	GC41 3	GC42 3	GC43 3	GC44 3	GC45 3	GC46 3	GC47 3	GC48 3	GC49 3	GC50 3	GC51 3	GC52 3	GC53 3	GC54 3	GC55 3	GC56 3	GC57 3	GC58 3	GC59 3	GC60 3	GC61 3	GC62 3	GC63 3	GC64 3	GC65 3	GC66 3	GC67 3	GC68 3	GC69 3	GC70 3	GC71 3	GC72 3	GC73 3	GC74 3	GC75 3	GC76 3	GC77 3	GC78 3	GC79 3	GC80 3	GC81 3	GC82 3	GC83 3	GC84 3	GC85 3	GC86 3	GC87 3	GC88 3	GC89 3	GC90 3	GC91 3	GC92 3	GC93 3	GC94 3	GC95 3	GC96 3	GC97 3	GC98 3	GC99 3	GC100 3
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Gene	Accession	Length (bp)	GC (%)	GC3 (%)	GC4 (%)	GC5 (%)	GC6 (%)	GC7 (%)	GC8 (%)	GC9 (%)	GC10 (%)	GC11 (%)	GC12 (%)	GC13 (%)	GC14 (%)	GC15 (%)	GC16 (%)	GC17 (%)	GC18 (%)	GC19 (%)	GC20 (%)	GC21 (%)	GC22 (%)	GC23 (%)	GC24 (%)	GC25 (%)	GC26 (%)	GC27 (%)	GC28 (%)	GC29 (%)	GC30 (%)	GC31 (%)	GC32 (%)	GC33 (%)	GC34 (%)	GC35 (%)	GC36 (%)	GC37 (%)	GC38 (%)	GC39 (%)	GC40 (%)	GC41 (%)	GC42 (%)	GC43 (%)	GC44 (%)	GC45 (%)	GC46 (%)	GC47 (%)	GC48 (%)	GC49 (%)	GC50 (%)	GC51 (%)	GC52 (%)	GC53 (%)	GC54 (%)	GC55 (%)	GC56 (%)	GC57 (%)	GC58 (%)	GC59 (%)	GC60 (%)	GC61 (%)	GC62 (%)	GC63 (%)	GC64 (%)	GC65 (%)	GC66 (%)	GC67 (%)	GC68 (%)	GC69 (%)	GC70 (%)	GC71 (%)	GC72 (%)	GC73 (%)	GC74 (%)	GC75 (%)	GC76 (%)	GC77 (%)	GC78 (%)	GC79 (%)	GC80 (%)	GC81 (%)	GC82 (%)	GC83 (%)	GC84 (%)	GC85 (%)	GC86 (%)	GC87 (%)	GC88 (%)	GC89 (%)	GC90 (%)	GC91 (%)	GC92 (%)	GC93 (%)	GC94 (%)	GC95 (%)	GC96 (%)	GC97 (%)	GC98 (%)	GC99 (%)	GC100 (%)
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Gene	Accession	Length (bp)	GC (%)	GC3 (%)	GC4 (%)	GC5 (%)	GC6 (%)	GC7 (%)	GC8 (%)	GC9 (%)	GC10 (%)	GC11 (%)	GC12 (%)	GC13 (%)	GC14 (%)	GC15 (%)	GC16 (%)	GC17 (%)	GC18 (%)	GC19 (%)	GC20 (%)	GC21 (%)	GC22 (%)	GC23 (%)	GC24 (%)	GC25 (%)	GC26 (%)	GC27 (%)	GC28 (%)	GC29 (%)	GC30 (%)	GC31 (%)	GC32 (%)	GC33 (%)	GC34 (%)	GC35 (%)	GC36 (%)	GC37 (%)	GC38 (%)	GC39 (%)	GC40 (%)	GC41 (%)	GC42 (%)	GC43 (%)	GC44 (%)	GC45 (%)	GC46 (%)	GC47 (%)	GC48 (%)	GC49 (%)	GC50 (%)	GC51 (%)	GC52 (%)	GC53 (%)	GC54 (%)	GC55 (%)	GC56 (%)	GC57 (%)	GC58 (%)	GC59 (%)	GC60 (%)	GC61 (%)	GC62 (%)	GC63 (%)	GC64 (%)	GC65 (%)	GC66 (%)	GC67 (%)	GC68 (%)	GC69 (%)	GC70 (%)	GC71 (%)	GC72 (%)	GC73 (%)	GC74 (%)	GC75 (%)	GC76 (%)	GC77 (%)	GC78 (%)	GC79 (%)	GC80 (%)	GC81 (%)	GC82 (%)	GC83 (%)	GC84 (%)	GC85 (%)	GC86 (%)	GC87 (%)	GC88 (%)	GC89 (%)	GC90 (%)	GC91 (%)	GC92 (%)	GC93 (%)	GC94 (%)	GC95 (%)	GC96 (%)	GC97 (%)	GC98 (%)	GC99 (%)	GC100 (%)
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Gene 2	Accession 2	Length 2	GC 2	GC3 2	GC4 2	GC5 2	GC6 2	GC7 2	GC8 2	GC9 2	GC10 2	GC11 2	GC12 2	GC13 2	GC14 2	GC15 2	GC16 2	GC17 2	GC18 2	GC19 2	GC20 2	GC21 2	GC22 2	GC23 2	GC24 2	GC25 2	GC26 2	GC27 2	GC28 2	GC29 2	GC30 2	GC31 2	GC32 2	GC33 2	GC34 2	GC35 2	GC36 2	GC37 2	GC38 2	GC39 2	GC40 2	GC41 2	GC42 2	GC43 2	GC44 2	GC45 2	GC46 2	GC47 2	GC48 2	GC49 2	GC50 2	GC51 2	GC52 2	GC53 2	GC54 2	GC55 2	GC56 2	GC57 2	GC58 2	GC59 2	GC60 2	GC61 2	GC62 2	GC63 2	GC64 2	GC65 2	GC66 2	GC67 2	GC68 2	GC69 2	GC70 2	GC71 2	GC72 2	GC73 2	GC74 2	GC75 2	GC76 2	GC77 2	GC78 2	GC79 2	GC80 2	GC81 2	GC82 2	GC83 2	GC84 2	GC85 2	GC86 2	GC87 2	GC88 2	GC89 2	GC90 2	GC91 2	GC92 2	GC93 2	GC94 2	GC95 2	GC96 2	GC97 2	GC98 2	GC99 2	GC100 2
Gene 3	Accession 3	Length 3	GC 3	GC3 3	GC4 3	GC5 3	GC6 3	GC7 3	GC8 3	GC9 3	GC10 3	GC11 3	GC12 3	GC13 3	GC14 3	GC15 3	GC16 3	GC17 3	GC18 3	GC19 3	GC20 3	GC21 3	GC22 3	GC23 3	GC24 3	GC25 3	GC26 3	GC27 3	GC28 3	GC29 3	GC30 3	GC31 3	GC32 3	GC33 3	GC34 3	GC35 3	GC36 3	GC37 3	GC38 3	GC39 3	GC40 3	GC41 3	GC42 3	GC43 3	GC44 3	GC45 3	GC46 3	GC47 3	GC48 3	GC49 3	GC50 3	GC51 3	GC52 3	GC53 3	GC54 3	GC55 3	GC56 3	GC57 3	GC58 3	GC59 3	GC60 3	GC61 3	GC62 3	GC63 3	GC64 3	GC65 3	GC66 3	GC67 3	GC68 3	GC69 3	GC70 3	GC71 3	GC72 3	GC73 3	GC74 3	GC75 3	GC76 3	GC77 3	GC78 3	GC79 3	GC80 3	GC81 3	GC82 3	GC83 3	GC84 3	GC85 3	GC86 3	GC87 3	GC88 3	GC89 3	GC90 3	GC91 3	GC92 3	GC93 3	GC94 3	GC95 3	GC96 3	GC97 3	GC98 3	GC99 3	GC100 3
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