

Patient-ID Age Gender Site of origin Grade Description seen with low g background of	AA1716 59 Male Peritoneal metastasis Low-grade A strip of neoplastic mucinous epithelium is grade cytologic and architectural features in a abundant extracellular mucin.
background of	abundant extracellular mucin.
seen with low g background of	rade cytologic and architectural features in a abundant extracellular mucin.

Patient-ID Age Gender Site of origin Grade Description	App2312 64 Female Peritoneal metastasis Low-grade Papillary and glandular formations of peoplastic
Description	Papillary and glandular formations of neoplastic
architectural fe	nelium are seen with low grade cytologic and eatures in a background of extracellular mucin
and blood.	

Patient-ID	AA1816
Age	42
Gender	Male
Site of origin	Peritoneal metastasis
Grade	Low-grade
Description	A single glandular structure composed of
neoplastic mu	cinous epithelium with low grade cytologic and
architectureal	features is seen embedded in fibrous tissue.

Patient-ID	AA1830
Age	72
Gender	Female
Site of origin	Peritoneal metastasis
Grade	Low-grade
Description	Strips of neoplastic mucinous epithelium are
seen with low	grade cytologic and architectural features,
associated with	h abundant extracellular mucin.

Patient-ID	AA1924					
Age	62					
Gender	Male					
Site of origin	Peritoneal metastasis					
Grade	Low-grade					
Description	Strips and papillary formations of neoplastic					
mucinous epi	thelium are seen with low grade cytologic and					
architectural features, associated with abundant extracellular						
mucin.						

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Patient-ID	AA1934
Age	50
Gender	Female
Site of origin	Peritoneal metastasis
Grade	High-grade
Description	Crowded aggregates of floating peoplastic
Description	Crowded aggregates of floating neoplastic
mucinous epith	helium are seen with complex architecture and
high grade cyt	ologic features.

Patient-ID	AA2021
Age	45
Gender	Female
Site of origin	Peritoneal metastasis
Grade	Low-grade
Description	Small strip of neoplastic mucinous epithelium
with low grade	e cytologic and architectural features is seen
associated wi	th abundant extracellular mucin.

Patient-ID	AA2051
Age	51
Gender	Female
Site of origin	Peritoneal metastasis
Grade	Low-grade
Description	A strip of neoplastic mucinous epithelium is seen
with low grade	e cytologic and architectural features, associated
with abundan	t extracellular mucin.

Patient-ID	AA2055
Age	54
Gender	Male
Site of origin	Peritoneal metastasis
Grade	Low-grade
Description	A strip of neoplastic mucinous epithelium is seen
with low grade	e cytologic and architectural features, associated
with abundan	t extracellular mucin.

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Figure S1: H&E stained histology sections from the discovery group. The sections presented were selected as representing the diagnosis of the patient, including histological grade, and containing a high fraction of tumor cells for visualization purpose. For each section, the location, grade and description of the area displayed are indicated.



Figure S2: Distribution of the Mutant Allelic Fraction at all positions determined to be somatically mutated by whole exome sequencing of 10 MCPs.

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							Blood		Tumor				
Patient ID Specime	Spacimon	n Gene	Mutation	Chr	Coord	Ref	A 1+	Ref	Alt	Allelic	Ref	Alt	Allelic
	Specimen						~"	coverage	Coverage	Fraction	Coverage	Coverage	Fraction
AA1811	MCP	KRAS	G12V	chr12	25398284	С	Α	198236	153	0.08%	95681	1227	1.27%
AA1811	LMNA	KRAS	G12V	chr12	25398284	С	Α	198239	153	0.08%	180783	4488	2.42%
AA1811	MCP	GNAS	R201C	chr20	57484420	С	Т	177522	218	0.12%	117171	4074	3.36%
AA1811	LMNA	GNAS	R201C	chr20	57484420	С	Т	177524	218	0.12%	249506	8961	3.47%



substrates

Figure S3: Analysis of matched low-grade MCP and primary LAMN from patient AA1811. (a) Summary table presenting the coverage and mutant allelic fraction observed at KRAS G12V and GNAS R201C mutations in both specimen. (b-g) Immunohistochemistry. The low grade MCP (bd) and matched primary LAMN (e-f) sections were stained using Anti-pErk staining (b,e), Anti-pAkt staining (c,f) and Anti-phospho-PKA substrates (d,g).



Figure S4: Evidence of chromosome 18 Loss of Heterozygosity. The minor allele frequency observed in the tumor at chromosome 18 heterozygous SNPs. We selected SNPs with Normal Allele frequency between 0.3 and 0.7 and cumulative coverage in normal and tumor greater than 100 fold.



Figure S5: Confirmation by Sanger sequencing of the coding mutations identified in the discovery group in the genes TP53, FAT3/4, SMAD2/3/4 and TFBR1/2. Above each panel the samples, the coding mutations as well as nucleic acid substitution are indicated.



Figure S6: Digital Droplet PCR. Representative 2D histograms displaying the distribution of droplets (colored dots) as a function of the intensity in WT probe fluorescence (y axis) versus mutant probe fluorescence (x axis). Clusters of droplets are framed in white (WT) or blue (mutant) (A-C) KRAS-G12V/A/D assay, representing a positive control containing three mutations (A) along a G12V positive (B) and a G12D positive sample (C). **(D-F)** GNAS-R201C assay representing a positive control (D: AA2051 after LMD) along with positive (E) and negative (F) samples. **(G-I)** GNAS-R201H assay representing a positive control (G: AA2004 after LMD) along with negative (H) and positive (I) samples.