

**S3 Table. GG-NER Incision Complex genes.** Burden testing results (based on 1063 MM/MGUS cases and 964 unaffected controls), SGS and prioritized SNV results, and intolerance to missense and loss of function variants (based on ExAC population data).

Gene	Position	Burden (p-value)	SGS	SNV	Intolerance to MS (Z)	Intolerance to LoF (pLI)
<i>USP45</i>	6:99880182-99963565	0.711	significant	Y	-0.74	0.00
<i>CHD1L</i>	1:146714291-146767447	0.006 <sup>‡</sup>			-1.82	0.00
<i>PARP1</i>	1:226548392-226595801	0.036 <sup>‡</sup>			0.57	0.01
<i>RPA2</i>	1:28218049-28241236	0.019 <sup>‡</sup>			0.41	0.89
<i>ERCC3</i>	2:128014866-128051752	0.020 <sup>‡</sup>		Y	1.17	0.00
<i>GTF2H2</i>	5:70330951-70363497	0.005 <sup>‡</sup>			2.27	0.72
<i>GTF2H5</i>	6:158589379-158620376	0.532			-0.70	0.05
<i>GTF2H4</i>	6:30875977-30882814	0.930			0.74	0.26
<i>RPA3</i>	7:7676575-7758238	1.000			-0.73	0.00
<i>XPA</i>	9:100437191-100459691	0.104			0.52	0.00
<i>GTF2H1</i>	11:18343816-18388590	0.001 <sup>‡</sup>			0.59	1.00
<i>DDB2</i>	11:47236493-47260769	0.929			1.25	0.01
<i>DDB1</i>	11:61066919-61100684	0.001 <sup>‡</sup>			5.67	1.00
<i>GTF2H3</i>	12:124118286-124147151	0.957			0.04	0.00
<i>ERCC5</i>	13:103504468-103524748	0.462			-0.61	0.00
<i>CUL4A</i>	13:113863931-113919392	0.390			2.73	1.00
<i>PARP2</i>	14:20811773-20826063	0.190			-0.55	0.00
<i>ERCC4</i>	16:14014014-14046205	0.371		Y	-1.06	0.00
<i>RPA1</i>	17:1733273-1802848	0.480			0.43	0.31
<i>ERCC2</i>	19:45854649-45873845	0.503	suggestive		0.44	0.00
<i>ERCC1</i>	19:45910591-45927177	0.402	suggestive		-0.13	0.00
<i>RBX1</i>	22:41347351-41369019	-			2.68	0.94
<i>CUL4B</i>	X:119658446-119709684	-	-		3.88	1.00

**Legend:** Position – build HG19; Burden – p-values based on the c-alpha test of high-impact variants with AAF < 0.001 (see Methods section), “-” indicates gene not tested (no variants observed), <sup>‡</sup>significant after multiple testing correction p < 0.0022 (=0.05/23), <sup>†</sup>nominally significant p < 0.05; SGS – gene captured by a genome-wide significant or suggestive shared genomic segment, “-” indicates not tested (SGS only looks at autosomal chromosomes); SNV – “Y” indicates a single nucleotide variant with AAF < 0.001, high or moderate deleteriousness, and observed segregating in a high-risk MM pedigree or pathogenic in ClinVar; Intolerance to MS – gene’s intolerance to missense variants based on analysis of ExAC data<sup>41</sup>, signed Z score based on deviation of observed counts from expected, positive Z indicates intolerance to variation; LoF – based on analysis of ExAC data<sup>41</sup>, Loss of Function (LoF) variants include splice donor or acceptor or non-sense SNVs, genes with a probability of LoF Intolerance (pLI) >= 0.9 are considered extremely intolerant to LoF SNVs.