Description of Additional Supplementary Files

File Name: Supplementary Data 1

Description: 40 cSCCs - Patient/Tumour Details

File Name: Supplementary Data 2

Description: Whole exome sequencing (WES) reads yield and coverage across the

40 normal - SCC pairs

File Name: Supplementary Data 3

Description: All identified somatic mutations across 40 cSCC WES samples, with full annotation, aligned reads, VAF and clonality CCFs, subpopulation frequency

estimated by EXPANDS

File Name: Supplementary Data 4

Description: Mutation signatures across all SCC exomes.

File Name: Supplementary Data 5

Description: CNAs and cn-LOH events across the 40 SCC exomes

File Name: Supplementary Data 6

Description: Significantly amplified and deleted regions across the 40 cSCC

tumours based on GISTIC2.0

File Name: Supplementary Data 7

Description: Frequent copy number variations in cSCC

File Name: Supplementary Data 8

Description: Significantly mutated genes of 40 cSCC exomes based on MutSig

analysis (p < 0.05)

File Name: Supplementary Data 9

Description: Significantly mutated genes across 40 cSCC exomes based on

OncodriveFM (q < 0.05)

File Name: Supplementary Data 10

Description: Significantly mutated genes across 40 cSCC exomes based on

OncodriveClust (q < 0.05)

File Name: Supplementary Data 11

Description: Somatic mutations across 40 cSCC WES samples for NOTCH1/2

File Name: Supplementary Data 12

Description: Somatic mutations across 40 cSCC WES samples for TP53 and

CDKN2A

File Name: Supplementary Data 13

Description: Cross Study Comparison of Significantly Mutated Genes

File Name: Supplementary Data 14

Description: Somatic mutations across 40 cSCC WES samples for HRAS,

MAP3K9, PTEN, SF3B1, VPS41 and WHSC1

File Name: Supplementary Data 15

Description: Somatic mutations across 40 cSCC WES samples for FLNB, GLIS3,

CACNA1C, HERC6, TRAPPC9 and MAPK1IP1L

File Name: Supplementary Data 16

Description: Somatic mutations across 40 cSCC WES samples for GRHL2, CLCN3,

TMEM51, ATP1A1, LCLAT1 and CRY1

File Name: Supplementary Data 17

Description: Comparison of 22 SMGs alteration frequencies in immunosupressed

(IS) an immunocompetent (IC) patient samples

File Name: Supplementary Data 18

Description: Mutational signature analysis of mutations in 22 SMGs

File Name: Supplementary Data 19

Description: Tumour purity estimates by EXPANDS and SciClone

File Name: Supplementary Data 20

Description: Estimated number of clones in 35 Exome samples

File Name: Supplementary Data 21

Description: Rank order clonality of 22 SMGs

File Name: Supplementary Data 22 **Description:** Gene expression Data set 1

File Name: Supplementary Data 23

Description: Cross Study Comparison of Significantly Mutated Genes by

Differentiation Status

File Name: Supplementary Data 24 **Description:** Kegg pathway analysis

File Name: Supplementary Data 25

Description: OncodriveFM pathway mutation significance for all profiled KEGG

pathways

File Name: Supplementary Data 26

Description: cSCC Cell Line Patient Details

File Name: Supplementary Data 27

Description: All identified annotated somatic mutations in 15 cell lines

File Name: Supplementary Data 28

Description: Mutation signatures across all cSCC cell lines.

File Name: Supplementary Data 29

Description: Somatic nonsynonymous mutations in the 22 SMGs across cell lines

File Name: Supplementary Data 30 **Description:** Differential expression analysis of SCC cell lines against normal

human keratinocytes