Progressive 3q amplification consistently targets SOX2 in preinvasive squamous lung cancer

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- (a) Patient 002. Patient 002 had multiple surveillance bronchoscopies over a 13 month period. The left lower lobe lesion (blue) progressed to cancer after 13 months (black). A left lower lobectomy was performed at month 13 as there was a clinical suspicion of lung cancer , despite no definite radiological evidence. The resection specimen showed widespread high-grade dysplasia and a focal area of invasion, CA1.
- (b) Patient 017. Patient 017 underwent a left upper lobectomy for squamous cell carcinoma prior to entering the study. He was subsequently found to have high-grade dysplasia in the resection margin and had a number of surveillance bronchoscopies. Although there was no biopsy proven evidence of invasion there was a clinical suspicion of cancer so he proceeded to a pneumonectomy (solid bar) at month 15. The pneumonectomy specimen confirmed high-grade preinvasive disease with no focal invasion. Subsequent biopsies at 17 months demonstrated extensive involvement of the trachea.
- (c) Patient 026. Patient 026 entered the surveillance programme after surgical resection of the right middle and lower lobes for SQC two biopsies (026-HG7 and 12) were taken at the first surveillance bronchoscopy, and another (026-HG13) at a subsequent bronchoscopy at month 5.

Supplementary Table 1. Genes within the 3q minimal commonly amplified region (MCAR). Genes, annotated in the reference human genome (Ensembl - NCBI 36; www.ensembl.org) and their location on 3q. Brief notes on the annotated genes are provided. Information was sourced from on-line databases including the Online Mendelian Inheritance in Man (http://www.ncbi.nlm.nih.gov/omim) and references therein.

Gene	Gene Start (bp)	Brief Notes
KCNMB2	180007823	Ca activated potassium channel
ZMAT3	180224221	p53 induced zinc finger transcrition factor, anti-proliferative effects
PIK3CA	<u>180349005</u>	See main text.
KCNMB3	180443259	Calcium-activated potassium channel
<i>ZNF639</i>	<u>180524245</u>	Zinc finger transcription factor
MFN1	<u>180548174</u>	Mediates mitochondrial fusion
GNB4	<u>180599696</u>	GTPase binding protein - mediates signal transduction
ACTL6A	<u>180763402</u>	May modulate transcription factor (including p53 and cMyc) effects
MRPL47	<u>180788951</u>	Mitochondrial protein – unknown function
NDUFB5	<u>180805269</u>	Mitochondrial respiratory chain complex
USP13	<u>180853635</u>	Deubiqitinating enzyme
PEX5L	<u>181000744</u>	Binds peroxisomal targetting signal
TTC14	<u>181802625</u>	Potentially involved in RNA binding
CCDC39	<u>181814491</u>	Unknown function
FXR1	<u>182113146</u>	Potentially involved in RNA binding
DNAJC19	<u>182184200</u>	Component of mitochondrial protein import motor
SOX2	182912416	See main text

Supplementary Figure 1

