

Supplementary Table 1. Characteristics and outcomes of biopsied nodules of never-smokers detected at baseline and during follow-up

	Total (n=139)	Detected at baseline (n=104)	Detected during follow- up (n=35)	p-value
Age at diagnosis, mean \pm SD	60.4 \pm 11.9	60.1 \pm 11.7	61.3 \pm 12.6	0.608
Nodule type, n (%)				0.009
Solid	51 (36.7)	30 (28.8)	21 (60.0)	
Part-solid	47 (33.8)	41 (39.4)	6 (17.1)	
Pure GGN	34 (24.5)	27 (26.0)	7 (20.0)	
Cavitary	7 (5.0)	6 (5.8)	1 (2.9)	
Time of planning invasive diagnosis, n (%)				0.070
At first detection of relevant nodule	65 (46.8)	44 (42.3)	21 (60.0)	
During follow-up	74 (53.2)	60 (57.7)	14 (40.0)	
Time from detection to biopsy, months, mean \pm SD	19.8 \pm 30.0	19.7 \pm 30.0	20.2 \pm 30.4	0.922
Nodule size at detection, mm, mean \pm SD	15.4 \pm 8.2	16.2 \pm 8.3	12.7 \pm 7.7	0.029
Nodule size at biopsy, mm, mean \pm SD	17.5 \pm 8.3	18.1 \pm 8.2	15.7 \pm 8.3	0.146
Type of biopsy, n (%)				
Video-assisted thoracic surgery (VATS)	89 (64.0)	71 (68.3)	18 (51.4)	0.073
Percutaneous needle biopsy	38 (27.3)	25 (24.0)	13 (37.1)	0.132
Bronchoscopic biopsy*	12 (8.6)	8 (7.7)	4 (11.4)	0.496
Diagnosed as lung cancer, n (%)	84 (60.4)	74 (71.2)	10 (28.6)	<0.001
Diagnosed as benign (false-positive), n (%)	50 (36.0)	30 (28.8)	20 (57.1)	0.003

SD=standard deviation, GGN=ground glass nodule

*Including EBUS (endobronchial ultrasound) and radial-EBUS guided biopsy

Supplementary Table 2. Subgroup analyses of main outcomes stratified by age and sex

Subgroups	Total	Never-smoker	Ever-smoker	p value
Age <55 (N=25,730)	25,730	11,675	14,055	
Received invasive biopsy, n/N (%)	117 (0.5)	50 (0.4)	67 (0.5)	0.565
Diagnosed as lung cancer, n/N (%)	65 (0.3)	29 (0.2)	36 (0.3)	0.902
Diagnosed as benign (false-positive), n/N (%)	52 (0.2)	21 (0.2)	31 (0.2)	0.469
Any procedure-related complication, n/N (%)	13 (0.1)	4 (0.0)	9 (0.1)	0.290
Age ≥55 (N=11,706)	11,706	6,293	5,413	
Received invasive biopsy, n/N (%)	216 (1.8)	89 (1.4)	127 (2.3)	<0.001
Diagnosed as lung cancer, n/N (%)	142 (1.2)	55 (0.9)	87 (1.6)	<0.001
Diagnosed as benign (false-positive), n/N (%)	67 (0.6)	29 (0.5)	38 (0.7)	0.085
Any procedure-related complication, n/N (%)	42 (0.4)	16 (0.3)	26 (0.5)	0.041
Sex: male (N=23,827)	23,827	5,644	18,183	
Received invasive biopsy, n/N (%)	226 (0.9)	42 (0.7)	184 (1.0)	0.070
Diagnosed as lung cancer, n/N (%)	136 (0.6)	20 (0.4)	116 (0.6)	0.013
Diagnosed as benign (false-positive), n/N (%)	87 (0.4)	21 (0.4)	66 (0.4)	0.921
Any procedure-related complication, n/N (%)	42 (0.2)	7 (0.1)	35 (0.2)	0.284
Sex: female (N=13,609)	13,609	12,324	1,285	
Received invasive biopsy, n/N (%)	107 (0.8)	97 (0.8)	10 (0.8)	0.973
Diagnosed as lung cancer, n/N (%)	71 (0.5)	64 (0.5)	7 (0.5)	0.904
Diagnosed as benign (false-positive), n/N (%)	32 (0.2)	29 (0.2)	3 (0.2)	0.990
Any procedure-related complication, n/N (%)	13 (0.1)	13 (0.1)	0 (0)	0.244

