## **European Respiratory Society Annual Congress 2012**

**Abstract Number: 2688** 

**Publication Number: P1976** 

**Abstract Group:** 6.3. Tobacco, Smoking Control and Health Education

Keyword 1: Smoking Keyword 2: Longitudinal study Keyword 3: Lung cancer / Oncology

**Title:** A pharmacological intervention with varenicline among a lung cancer LDCT screening trial: The MILD experience

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**Body:** Introduction: Low dose CT screening (LDCT) for lung cancer may be a teachable moment for smoking cessation (SC), but no pharmacological intervention has been performed in LDCT trials. Aim: A three-month Varenicline course in a group of patients (Pts) enrolled in the Milan Lung Detection Trial, with biochemical verification of the smoking status. Matherials and Methods: 187 Pts received Varenicline;43% and 32% of them were allocated to 1 vs 2-year LDCT, while 25% to minimal SC advice and no LDCT (Ctrl).Lung function testings (PFTs), exhaled carbon monoxide (CO) and side effects were longitudinally recorded.Pts with a CO ≤6 ppm were considered abstinent.Descriptive statistic as well as parametric and non-parametric tests were performed. Results: Pts were 61±5.2 years old, with a mean CO of 16.3±7.9 ppm,a smoking history of 22.2±21.9 pack/years,a Fagerström test of 7.5±2.2 points and a slight decrease in mean FEV1% (84.1±14.6). Global guit rates were 51.7%, 50.7% and 41.8% on month 1,3,6 respectively; guit rates were equal in LDCT subgroups but they were lower in the subgroup of Pts with FEV1%≥70<80%. Among non-quitters, those in the LDCT active arms, as well as those with a FEV1%≥70<80% showed higher CO values than Ctrl at baseline and along the study. Side effects were presents in 28.4% of Pts and therapy discontinuation happened in 20.4% of cases. Conclusion: A pharmacological intervention within a LDCT trial can lead to rewarding percentages of SC. Mild function impairment poses a higher risk of continuative smoking, while being in an active LDCT arm may result in stronger smoking intensity over time.