SUPPLEMENTARY FIGURES AND METHODS

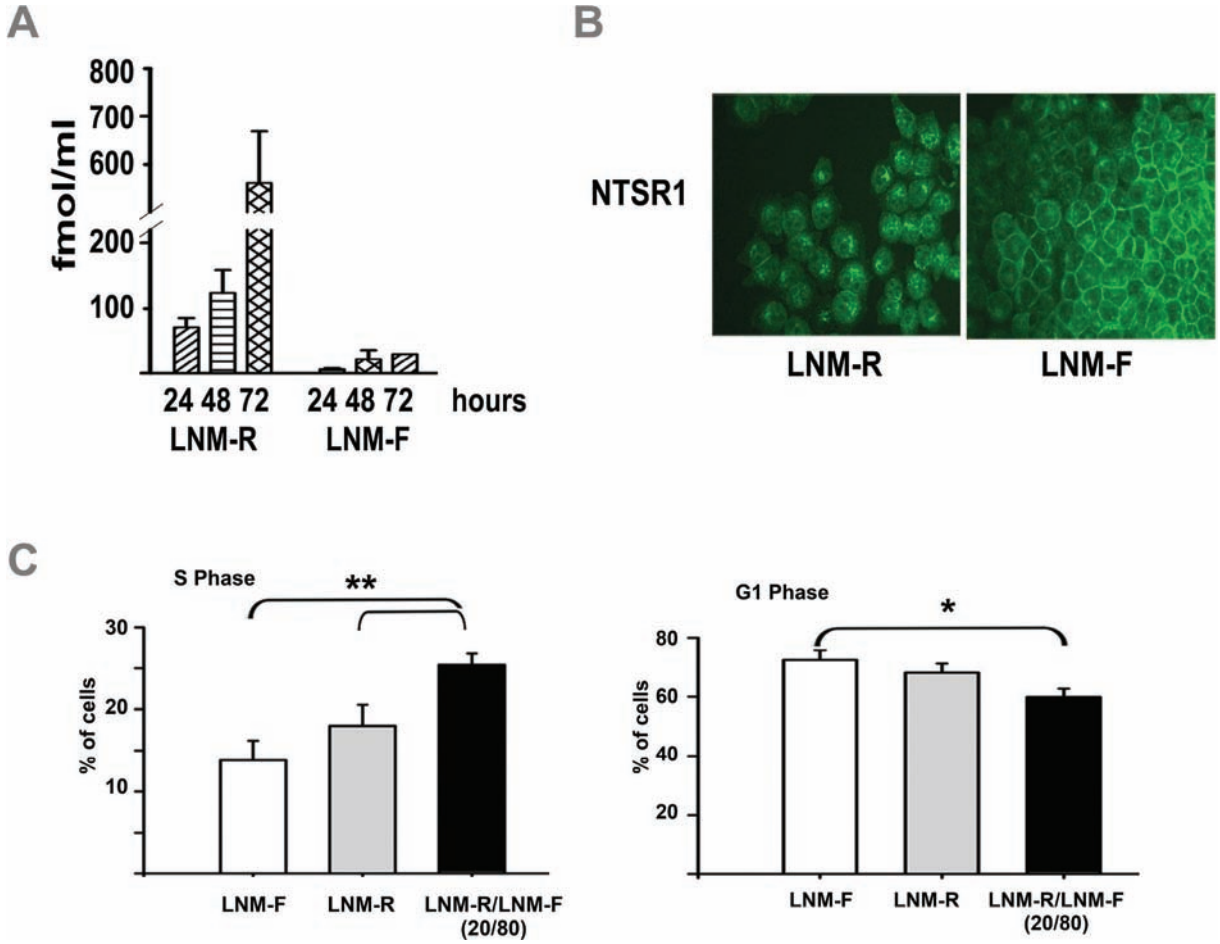

Figure S1: Lung cancer cells characterization. **(A)** NTS radioimmunoassay performed on cellular media of 10^6 LMN-R or LNM-F cells grown for 24, 48, or 72h. Each experiment was performed 3 to 5 times in duplicate. **(B)** Autocrine regulation was demonstrated in LNM-R cells with NTSR1 immunocytochemistry. In LNM-F cells, NTSR1 is localized at the cell surface, demonstrating the naive status of the cells. In contrast, in LNM-R cells, NTSR1 is located in a peri-nuclear area, suggesting an intense internalization of the receptor. **(C)** Cell proliferation was measured by fluorescence activated cell sorting, in LNM-R and LNM-F seeded alone or seeded at the ratio of 20/80 % LNM-R/LNM-F. Left and right represents the % of in S and G1 phase, respectively.

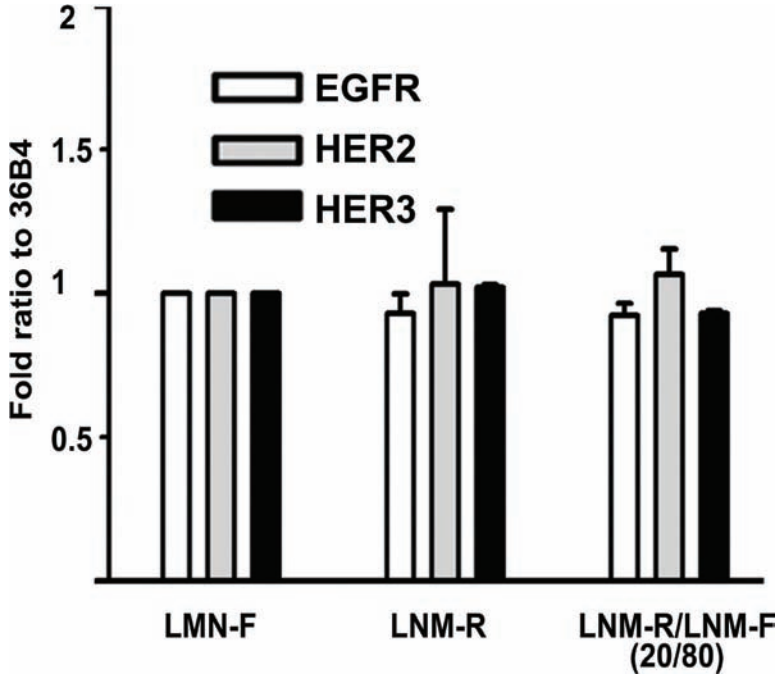

Figure S2: EGFR, HER2, and HER3 Q PCR performed on LNM-F, LNM-R and the mixture (R/F 20/80).

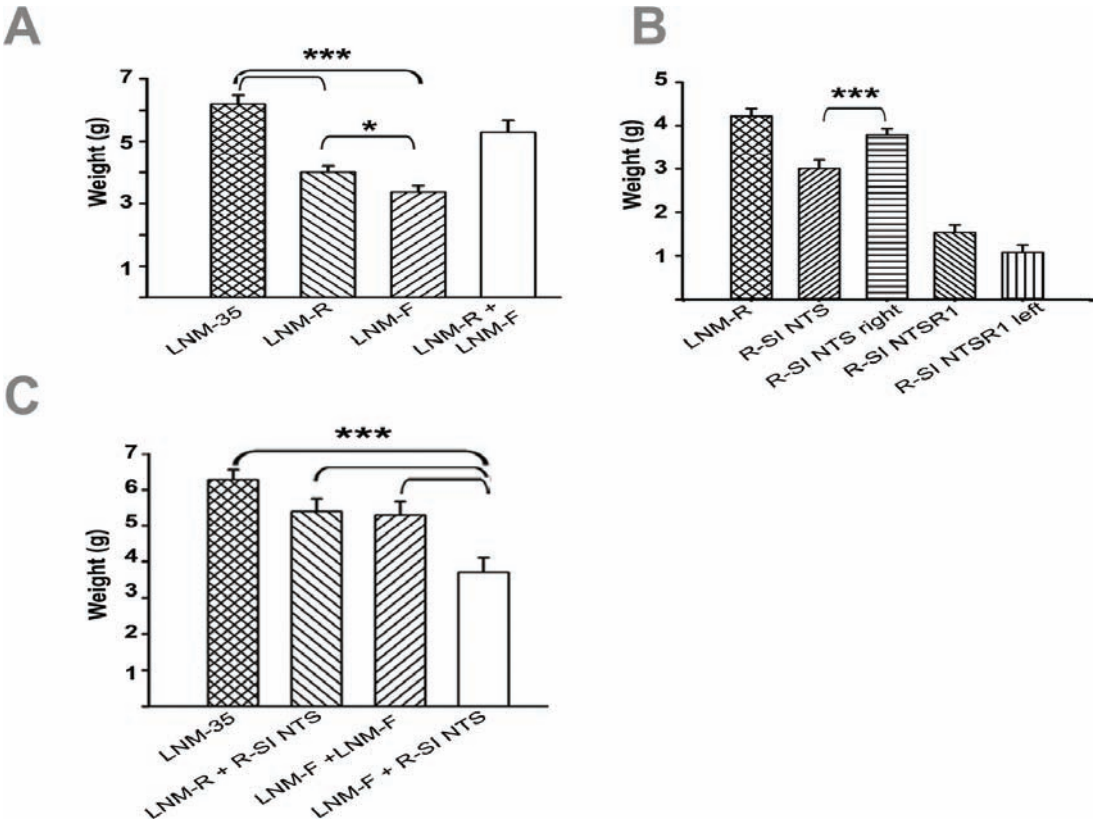

Figure S3: NTS/NTSR1 complex enhances experimental tumor weight. (A) Tumor weights generated by LNM35, LNM-R and LMN-F cells xenografted into nude mice. One million cells from LNM35, LNM-R, LNM-F, or a mixture of LNM-R and LNM-F (50/50) were subcutaneously injected in 24, 36, 34, or 12 nude mice, respectively. **(B)** NTS endocrine regulation and enhancement of tumor weight. One million R-SI NTS cells were injected into the right mice flanks, and one million R-SI NTSR1 cells were injected into the left mice flanks of the same mouse (n=18). In a second set, one million LNM-R cells were injected only into the right mice flanks (n=36). **(C)** Tumor weights generated by mixture of cells expressing or not NTS. One million LNM35 or a 50/50 mixture of R-SI NTS and LNM-R cells or a 50/50 mixture of LNM-R and LNM-F or a 50/50 mixture of R-SI NTS and LNM-F cells were injected in the right mice flanks, 28, 17, 11, and 14 mice were injected, respectively.