

Can Antipsychotics Prevent the Progression of Non-Small Cell Lung Cancer?

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Background

Despite great advances in diagnostic and therapeutic technologies, lung cancer remains the leading cause of cancer-related mortality worldwide. Non-small cell lung cancer (NSCLC) accounts for approximately 85% of lung cancer cases. Recently, some antipsychotics have been shown to possess anticancer activity.

Aim

Present study aimed to evaluate the possibility of antipsychotics as a therapeutic tools for lung cancer patients.

Methods

1. Meta-analysis of lung cancer with schizophrenia cohort studies
2. Cell proliferation assay
3. Colony formation assay
4. Wound healing assay
5. Western blot analysis
6. Flow cytometry and DNA fragmentation assay
7. Measurement of intracellular Ca^{2+} concentration
8. Tumor xenograft and histological analysis

Figure 1. The association between schizophrenia and lung cancer risk

Study	SIR (95% CI)	Weight
Goldacre 2005	1.18 (0.94-1.45)	0.1239
Barak 2005	0.65 (0.29-1.23)	0.0377
Dalton 2005	0.98 (0.69-1.38)	0.0916
Grinshpoon 2005	1.13 (0.71-1.8)	0.0683
Hippisley-Cox 2007	0.64 (0.44-0.94)	0.0844
Chou 2011	0.55 (0.44-0.69)	0.1218
Lin 2013	0.81 (0.77-0.96)	0.1488
Ji 2013	0.75 (0.68-0.82)	0.1518
Osborn 2013	0.95 (0.65-1.41)	0.0828
Raviv 2014	1.43 (0.98-2.01)	0.0888
Summary	0.86 (0.74-1.01)	

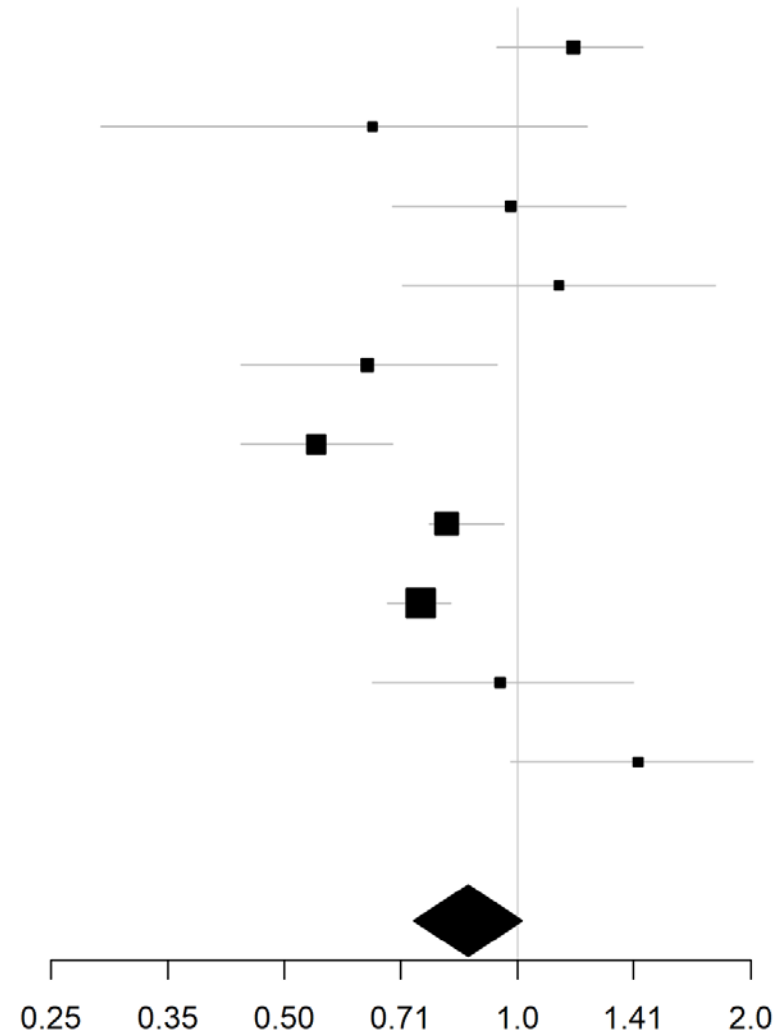


Figure 2. Treatment with TFP and TFP analogs inhibited A549 cell proliferation

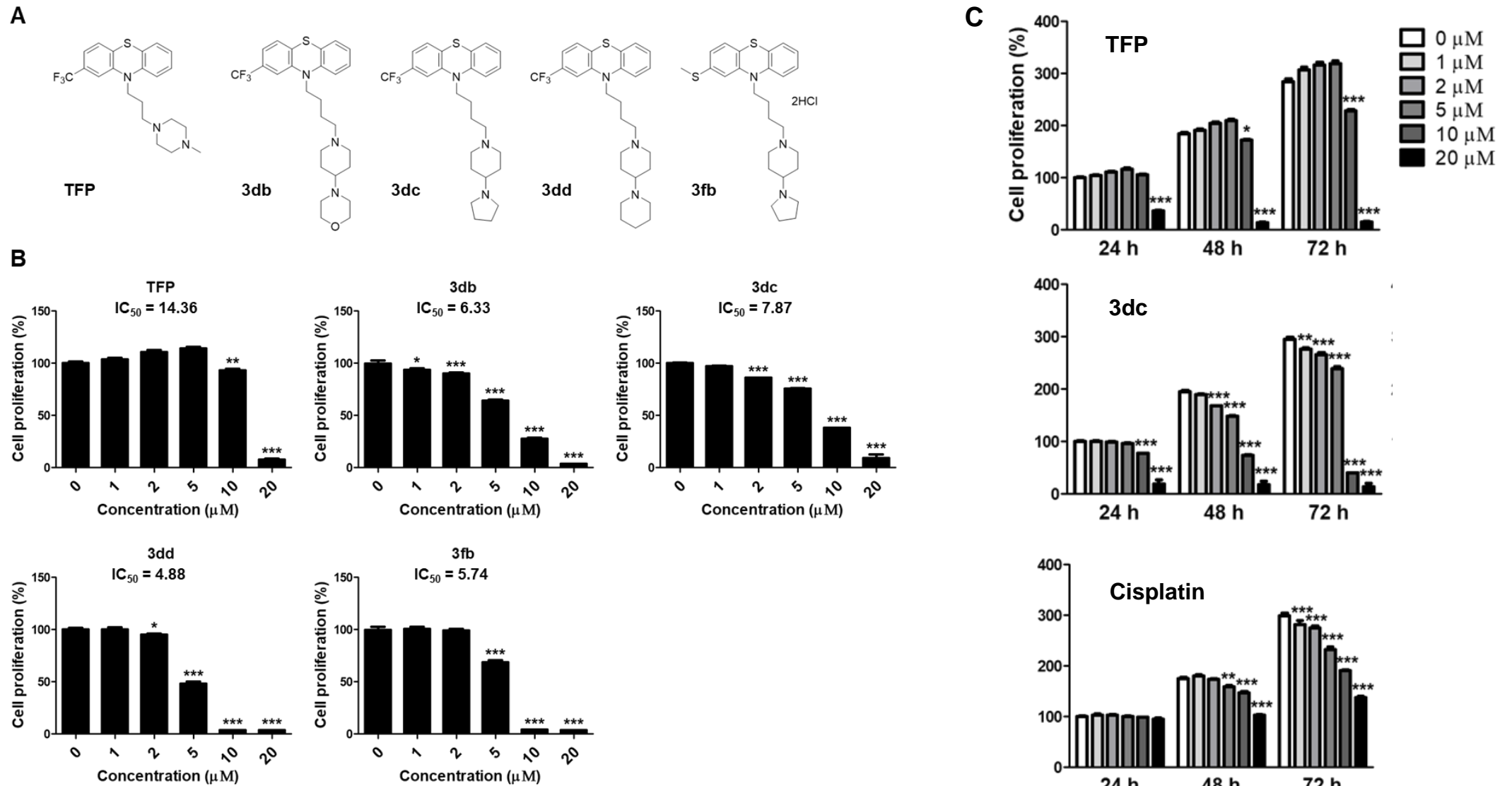


Figure 3. Treatment with TFP and 3dc suppressed colony formation and migration

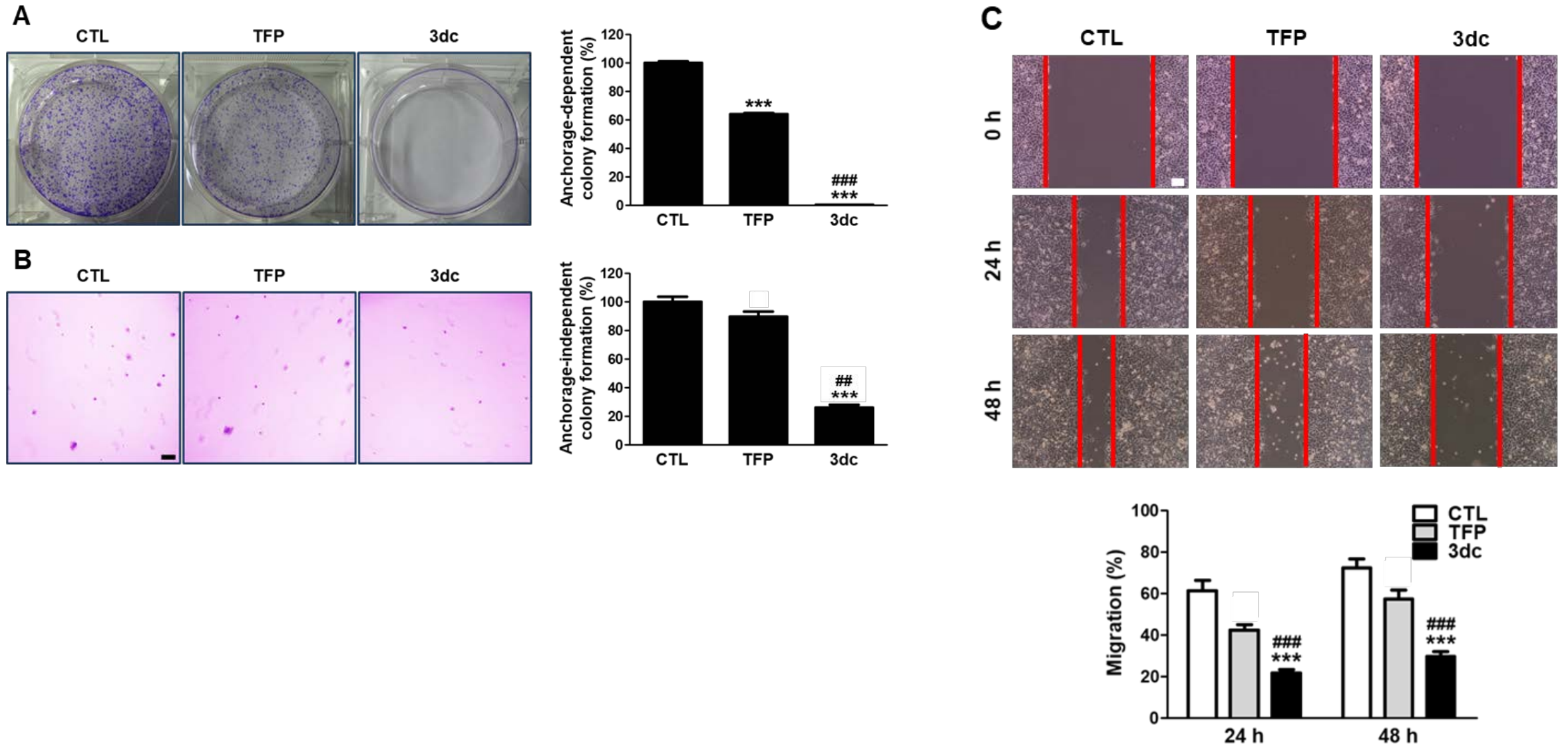


Figure 4. Western blot analysis of apoptosis- and survival-related factors and flow cytometric analysis of TFP and 3dc treatment

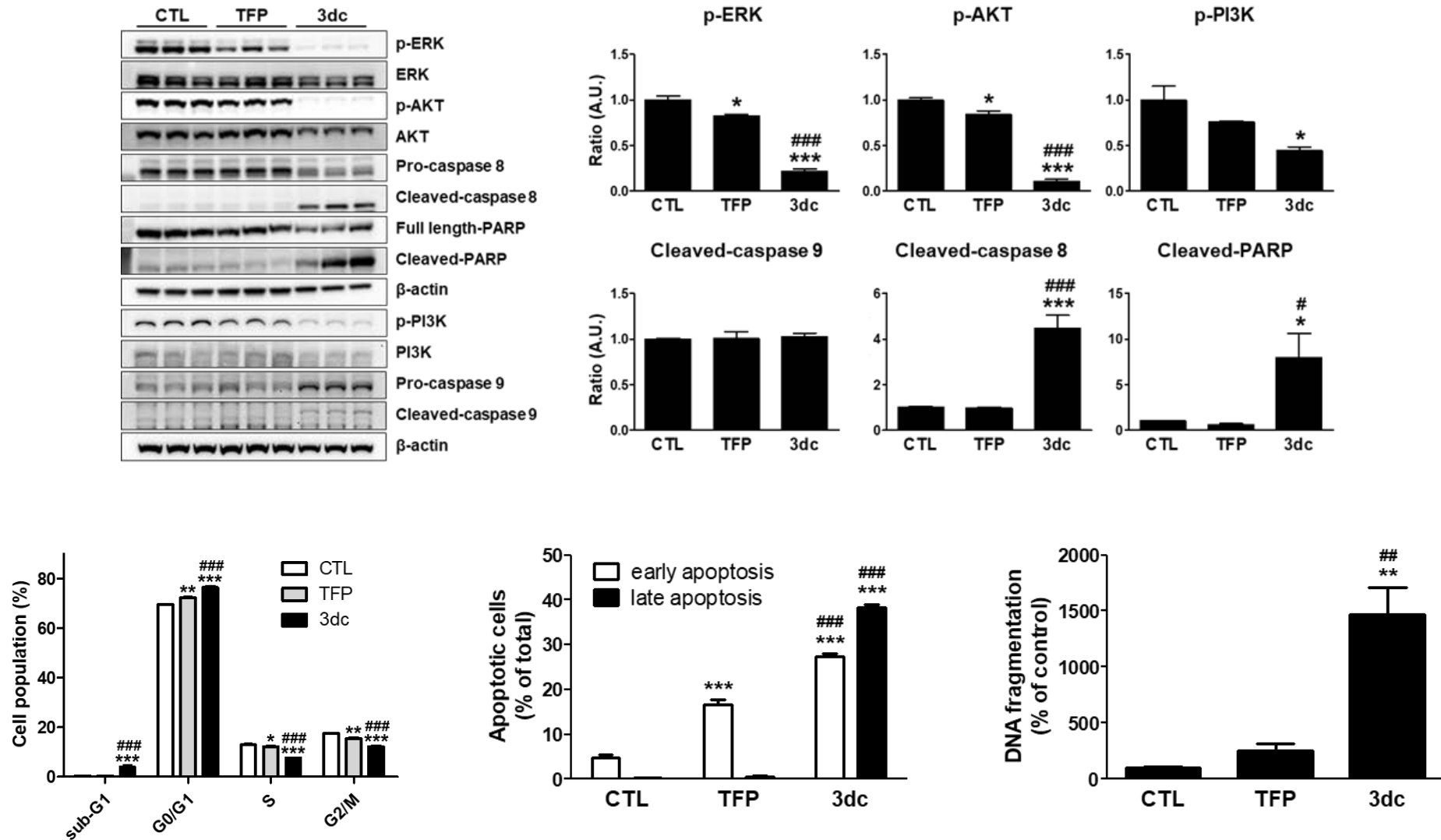


Figure 5. TFP and 3dc treatment increased intracellular Ca^{2+} levels

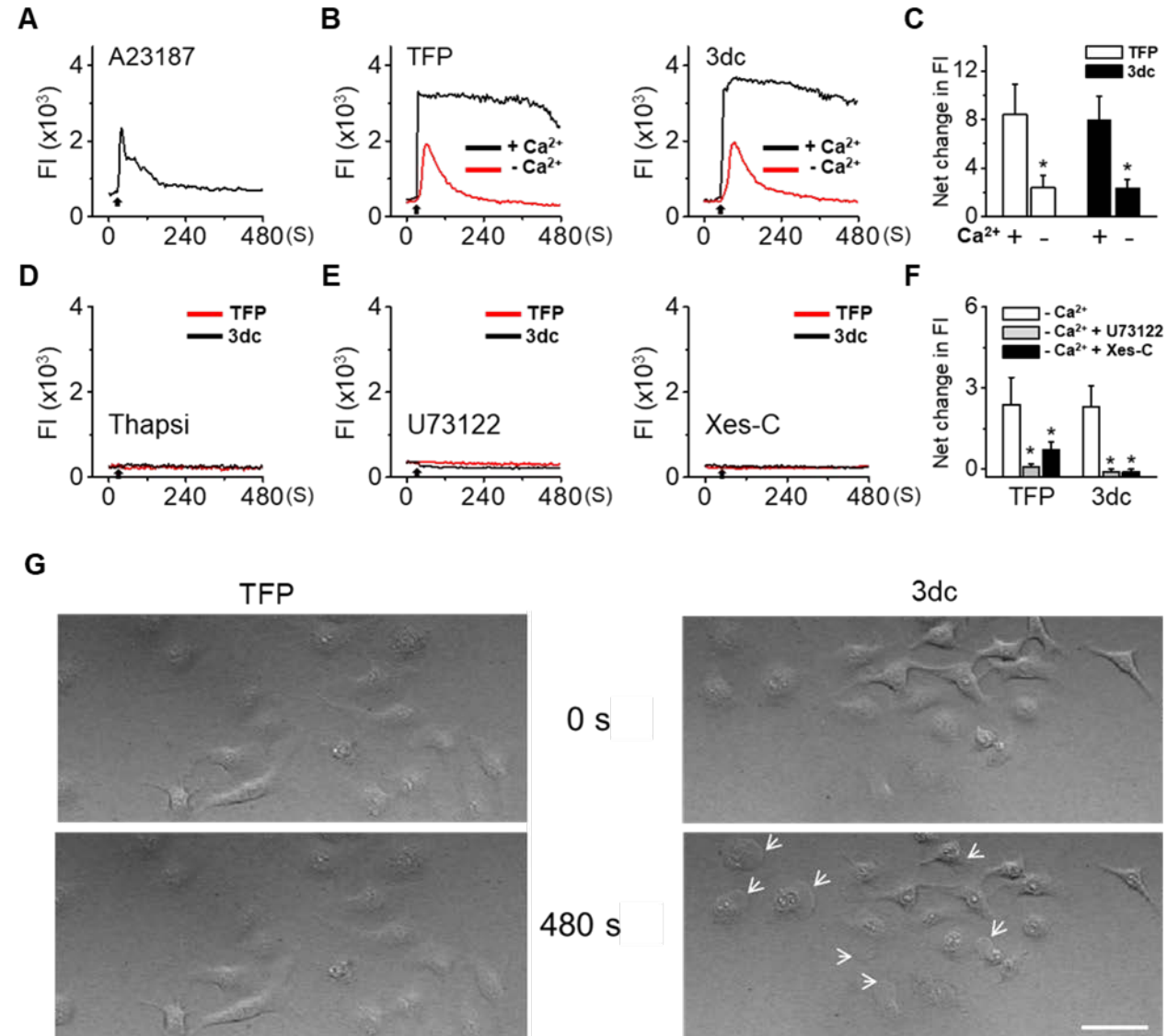
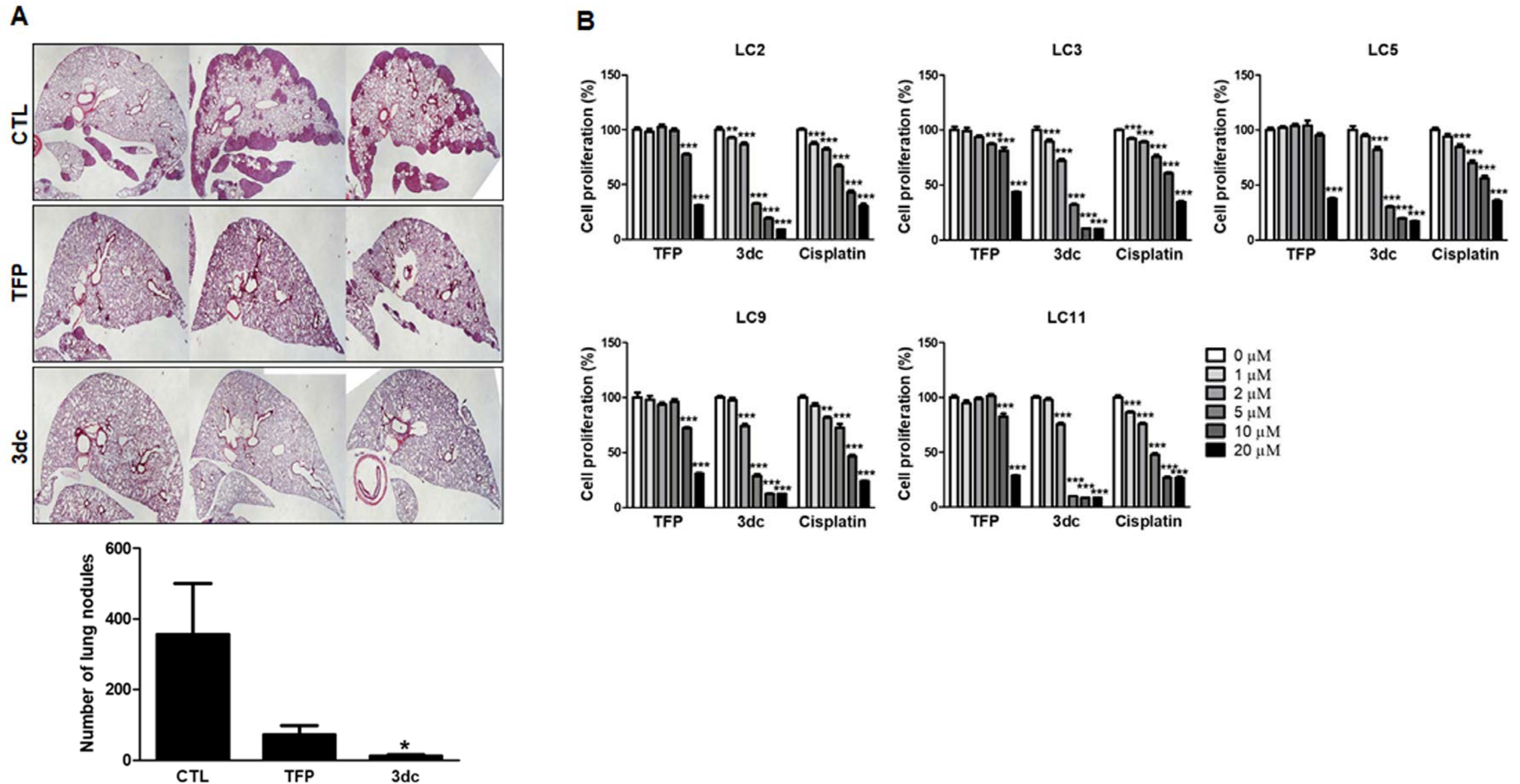


Figure 7. Inhibitory effect of TFP and 3dc on metastasis in A549 cell-derived xenograft model and human primary lung cancer cells



Conclusion

According to our analysis of publicly available clinical data and *in vitro* and *in vivo* experiments, we suggest that some kinds of antipsychotics prevent the progression of NSCLC. Furthermore, this study indicates a synthetic TFP analog that could be a potential therapeutic for lung cancer.