PublisherInfo				
PublisherName	:	BioMed Central		
PublisherLocation	:	London		
PublisherImprintName	:	BioMed Central		

## Skin deep

ArticleInfo		
ArticleID	:	4696
ArticleDOI	:	10.1186/gb-spotlight-20030211-01
ArticleCitationID	:	spotlight-20030211-01
ArticleSequenceNumber	:	48
ArticleCategory	:	Research news
ArticleFirstPage	÷	1
ArticleLastPage	:	2
ArticleHistory	:	RegistrationDate: 2003–2–11OnlineDate: 2003–2–11
ArticleCopyright	:	BioMed Central Ltd2003
ArticleGrants	:	
ArticleContext	:	130594411

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Fundamental differences between human and mouse skin, and differences in cellular transformation, pose a challenge to the development of useful models for studying skin diseases and malignancies. In the February 6 Nature Maya Dajee and colleagues at Stanford University School of Medicine in California describe experiments in normal epidermal cells that demonstrate the roles of oncogenic Ras and NF $\kappa$ B pathways in neoplastic transformation (*Nature* 2003, **421**:639-643). They used an animal model in which normal human skin is grafted onto the back of immunodeficient *scid* mice. They delivered a series of oncogenic genes to human keratinocytes using retroviral infections. Co-expression of oncogenic Ras and a stable repressor mutant of I $\kappa$ B $\alpha$  induced large neoplasms similar to human squamous cell carcinoma (SCC). The tumors displayed several SCC characteristics including an elevated mitotic index. Blocking NF $\kappa$ B activity appears to overcome Ras-induced growth arrest and induces the expression of high levels of the protein kinase CDK4. Dajee *et al.* also demonstrate the importance of the human integrin  $\alpha$ 6 $\beta$ 4 in the skin tumorigenesis process.

## References

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