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Up-regulation of focal adhesion kinase in non-small cell lung cancer

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<u>Up-regulation of focal adhesion kinase in non-small cell lung cancer</u>

Previous ArticleC2028T polymorphism in exon 12 and dinucleotide repeat polymorphism in intron 13 of the HIF-1α gene define HIF-1α protein expression in non-small cell lung cancer

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Summary

Focal adhesion kinase (FAK) is a non-receptor tyrosine kinase linked to the integrin and growth factor receptor-signalling pathways that regulates a number of the biological processes involved in neoplastic transformation, invasion and metastases, such as cell adhesion, migration and apoptosis. Its up-regulation might play a role in the tumourigenesis of invasive tumours, but its involvement in human lung cancer tissues has not yet been determined.

We immunohistochemically compared FAK expression and localisation in 60 formalin-fixed and paraffinembedded non-small cell lung cancer (NSCLC) tissues with that in the surrounding non-neoplastic tissue and in a further five microscopically normal lungs. FAK mRNA levels were quantitatively determined by real-time RT-PCR in frozen tissue specimens of all of the tumours and 21 matched non-neoplastic lung parenchymas, and protein expression in 16 homogenates of the matched neoplastic/non-neoplastic specimens was evaluated by Western blotting.

The three different techniques showed that FAK is weakly expressed in non-neoplastic lung parenchyma and up-regulated in NSCLCs. Moreover, Western blotting and real-time RT-PCR indicated a statistically significant correlation between FAK up-regulation and higher disease stages (I + II versus III + IV, p = 0.019 and 0.028, respectively). Our results provide evidence that FAK is up-regulated in NSCLCs, and suggest its potential involvement in lung cancer progression.

Keywords

- FAK
- NSCLC
- <u>Tumourigenesis</u>
- Cell migration
- Cancer progression
- Immunohistochemistry

• Real-time RT-PCR

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References

1. • Aplin A.E.

- Howe A.K.
- Juliano R.L.

Cell adhesion molecules, signal transduction and cell growth.

Curr Opin Cell Biol. 1999; 11: 737-744

View in Article

- <u>Scopus (287)</u>
- PubMed
- Crossref
- Google Scholar
- 2. Schaller M.D.
 - Borgman C.A.
 - Cobb B.S.
 - Vines R.R.
 - Reynolds A.B.
 - Parsons J.T.

pp125FAK a structurally distinctive protein-tyrosine kinase associated with focal adhesions.

Proc Natl Acad Sci USA. 1992; 89: 5192-5196

View in Article

- <u>Scopus (1274)</u>
- PubMed
- Crossref
- Google Scholar
- 3. Cobb B.S.
 - Schaller M.D.
 - Leu T.H.
 - Parsons J.T.

Stable association of pp60src and pp59fyn with the focal adhesion-associated protein tyrosine kinase, pp125FAK.

Mol Cell Biol. 1994; 14: 147-155

- Scopus (481)
- PubMed
- Crossref
- Google Scholar

- 4. Owen J.D.
 - Ruest P.J.
 - Fry D.W.
 - Hanks S.K.

Induced focal adhesion kinase (FAK) expression in FAK-null cells enhances cell spreading and migration requiring both auto- and activation loop phosphorylation sites and inhibits adhesion-dependent tyrosine phosphorylation of Pyk2.

Mol Cell Biol. 1999; 19: 4806-4818

View in Article

- <u>Scopus (337)</u>
- PubMed
- Crossref
- Google Scholar
- 5. Schaller M.D.
 - Otey C.A.
 - Hildebrand J.D.
 - Parsons J.T.

Focal adhesion kinase and paxillin bind to peptides mimicking beta integrin cytoplasmic domains. *J Cell Biol.* 1995; 130: 1181-1187

View in Article

- <u>Scopus (549)</u>
- PubMed
- Crossref
- Google Scholar
- 6. Schlaepfer D.D.
 - Hanks S.K.
 - Hunter T.
 - van der Geer P.

Integrin-mediated signal transduction linked to Ras pathway by GRB2 binding to focal adhesion kinase.

Nature. 1994; 372: 786-791

View in Article

- <u>Scopus (1419)</u>
- <u>PubMed</u>
- Crossref
- Google Scholar
- 7. Cary L.A.
 - Chang J.F.
 - Guan J.L.

Stimulation of cell migration by overexpression of focal adhesion kinase and its association with Src and Fyn.

J Cell Sci. 1996; 109: 1787-1794

View in Article

- PubMed
- Crossref
- Google Scholar
- 8. Sieg D.J.
 - Hauck C.R.
 - Ilic D.
 - Klingbeil C.K.
 - Schaefer E.
 - Damsky C.H.
 - o et al.

FAK integrates growth-factor and integrin signals to promote cell migration.

Nat Cell Biol. 2000; 2: 249-256

View in Article

• Scopus (1038)

- PubMed
- Crossref
- Google Scholar
- 9. Xu L.H.
 - o Owens L.V.
 - Sturge G.C.
 - Yang X.
 - Liu E.T.
 - Craven R.J.
 - o et al.

Attenuation of the expression of the focal adhesion kinase induces apoptosis in tumor cells. *Cell Growth Differ.* 1996; 7: 413-418

View in Article

- PubMed
- Google Scholar
- 10. Schaller M.D.
 - Hildebrand J.D.
 - Shannon J.D.
 - Fox J.W.
 - Vines R.R.
 - Parsons J.T.

Autophosphorylation of the focal adhesion kinase, pp125FAK, directs SH2-dependent binding of pp60src.

Mol Cell Biol. 1994; 14: 1680-1688

View in Article

- <u>Scopus (1107)</u>
- PubMed
- Crossref
- Google Scholar
- 11. Chen H.C.
 - Guan J.L.

Association of focal adhesion kinase with its potential substrate phosphatidylinositol 3-kinase.

Proc Natl Acad Sci USA. 1994; 91: 10148-10152

View in Article

- <u>Scopus (472)</u>
- PubMed
- Crossref
- Google Scholar
- 12. Zhang X.
 - Chattopadhyay A.
 - Ji Q.S.
 - Owen J.D.
 - Ruest P.J.
 - Carpenter G.
 - o et al.

Focal adhesion kinase promotes phospholipase C-gamma1 activity.

Proc Natl Acad Sci USA. 1999; 96: 9021-9026

View in Article

- Scopus (157)
- PubMed
- Crossref
- Google Scholar
- 13. Han D.C.
 - Guan J.L.

Association of focal adhesion kinase with Grb7 and its role in cell migration.

J Biol Chem. 1999; 274: 24425-24430

- Scopus (196)
- PubMed
- Crossref
- Google Scholar
- 14. Hauck C.R.
 - Hsia D.A.
 - Punte X.S.
 - Cheresh D.A.
 - Schlaepfer D.D.

FRNK blocks v-Src stimulated invasion and experimental metastases without effects on cell motility or growth.

EMBO J. 2002; 21: 6289-6302

View in Article

- Scopus (156)
- PubMed
- Crossref
- Google Scholar
- 15. Weiner T.M.
 - Liu E.T.
 - Craven R.J.
 - Cance W.G.

Expression of growth factor receptors, the focal adhesion kinase, and other tyrosine kinases in human soft tissue tumors.

Ann Surg Oncol. 1994; 1: 18-27

View in Article

- <u>Scopus (79)</u>
- <u>PubMed</u>
- Crossref
- Google Scholar
- 16. Gabarra-Niecko V.
 - Schaller M.D.
 - Dunty J.M.

FAK regulates biological processes important for the pathogenesis of cancer.

Cancer Metastasis Rev. 2003; 22: 359-374

View in Article

- Scopus (291)
- PubMed
- Crossref
- Google Scholar
- 17. Cance W.G.
 - Harris J.E.
 - Iacocca M.V.
 - Roche E.
 - Yang X.
 - Chang J.
 - o et al.

Immunohistochemical analyses of focal adhesion kinase expression in benign and malignant human breast and colon tissues: correlation with preinvasive and invasive phenotypes.

Clin Cancer Res. 2000; 6: 2417-2423

- PubMed
- Google Scholar
- 18. Tremblay L.
 - Hauck W.
 - Aprikian A.G.
 - Begin L.R.
 - Chapdelaine A.

• Chevalier S.

Focal adhesion kinase (pp125FAK) expression, activation and association with paxillin and p50CSK in human metastatic prostate carcinoma.

Int J Cancer. 1996; 68: 164-171

View in Article

- <u>Scopus (211)</u>
- PubMed
- Crossref
- Google Scholar
- 19. Owens L.V.
 - Xu L.
 - Dent G.A.
 - Yang X.
 - Sturge G.C.
 - Craven R.J.
 - o et al.

Focal adhesion kinase as a marker of invasive potential differentiated human thyroid cancer. *Ann Surg Oncol.* 1996; 3: 100-105

View in Article

- Scopus (198)
- PubMed
- Crossref
- Google Scholar
- 20. Judson P.L.
 - He X.
 - o Cance W.G.
 - Van Le L.

Overexpression of focal adhesion kinase, a protein tyrosine kinase, in ovarian carcinoma.

Cancer. 1999; 86: 1551-1556

View in Article

- Scopus (171)
- PubMed
- Crossref
- Google Scholar
- 21. Natarajan M.
 - Hecker T.P.
 - Gladson C.L.

FAK signaling in anaplastic astrocytoma and glioblastoma tumors.

Cancer J. 2003; 9: 126-133

View in Article

- Scopus (79)
- <u>PubMed</u>
- Crossref
- Google Scholar
- 22. Weiner T.M.
 - Liu E.T.
 - o Craven R.J.
 - Cance W.G.

Expression of focal adhesion kinase gene and invasive cancer.

Lancet. 1993; 342: 1024-1025

- Scopus (306)
- PubMed
- Abstract
- Google Scholar
- 23. Fujii T.
 - o Koshikawa K.

- Nomoto S.
- Okochi O.
- Kaneko T.
- Inoue S.
- o et al.

Focal adhesion kinase is overexpressed in hepatocellular carcinoma and can be served as an independent prognostic factor.

J Hepatol. 2004; 41: 104-111

View in Article

- <u>Scopus (95)</u>
- PubMed
- Abstract
- Full Text
- Full Text PDF
- Google Scholar
- 24. Recher C.
 - Ysebaert L.
 - Beyne-Rauzy O.
 - Mansat-De Mas V.
 - Ruidavets J.B.
 - Cariven P.

Expression of focal adhesion kinase in acute myeloid leukemia is associated with enhanced blast migration, increased cellularity, and poor prognosis.

Cancer Res. 2004; 64: 3191-3197

View in Article

- <u>Scopus (130)</u>
- PubMed
- Crossref
- Google Scholar
- 25. Kornberg L.
 - Fleigel J.

The effects of inducible overexpression of FAK-related non-kinase (FRNK) on a transformed epithelial cell line.

Anticancer Res. 2003; 23: 91-97

View in Article

- PubMed
- Google Scholar
- 26. Duxbury M.S.
 - Ito H.
 - Benoit E.
 - Zinner M.J.
 - Ashley S.W.
 - Whang E.E.

RNA interference targeting focal adhesion kinase enhances pancreatic adenocarcinoma gemcitabine chemosensitivity.

Biochem Biophys Res Commun. 2003; 311: 786-792

View in Article

- Scopus (135)
- PubMed
- Crossref
- Google Scholar
- 27. Beinke C.
 - Van Beuningen D.
 - o Cordes N.

Ionizing radiation modules of the expression and tyrosine phosphorylation of the focal adhesion-associated proteins focal adhesion kinase (FAK) and its substrates p130cas and paxillin in A549 human lung carcinoma cells *in vitro*.

Int J Radiat Biol. 2003; 79: 721-731

View in Article

- <u>Scopus (40)</u>
- PubMed
- Crossref
- Google Scholar
- 28. Parkin D.M.
 - Pisani P.
 - Ferlay J.

Global cancer statistics.

CA Cancer J Clin. 1999; 49: 33-64

View in Article

- Scopus (1806)
- PubMed
- Crossref
- Google Scholar
- 29. Wingo P.A.
 - Ries L.A.
 - Parker S.L.
 - Heath Jr., C.W.

Long-term cancer patient survival in the United States.

Cancer Epidemiol Biomarkers Prev. 1998; 7: 271-282

View in Article

- PubMed
- Google Scholar
- 30. ∘ Falleni M.
 - Pellegrini C.
 - Marchetti A.
 - o Oprandi B.
 - Buttitta F.
 - Barassi F.
 - o et al.

Survivin gene expression in early-stage non-small cell lung cancer.

J Pathol. 2003; 200: 620-626

View in Article

- Scopus (161)
- PubMed
- Crossref
- Google Scholar
- 31. Golubovskaya V.
 - Kaur A.
 - o Cance W.

Cloning and characterization of the promoter region of human focal adhesion kinase gene: nuclear factor kappa B and p53 binding sites.

Biochim Biophys Acta. 2004; 1678: 111-125

View in Article

- Scopus (140)
- PubMed
- Crossref
- Google Scholar
- 32. Schlaepfer D.D.
 - Mitra S.K.
 - Ilic D.

Control of motile and invasive cell phenotypes by focal adhesion kinase.

Biochim Biophys Acta. 2004; 1692: 77-102

View in Article

• Scopus (358)

- PubMed
- Crossref
- Google Scholar
- 33. ∘ Lark A.L.
 - Livasy C.A.
 - Calvo B.
 - Caskey L.
 - Moore D.T.
 - Yang X.
 - o et al.

Overexpression of focal adhesion kinase in primary colorectal carcinomas and colorectal liver metastases: immunohistochemistry and real-time PCR analyses.

Clin Cancer Res. 2003; 9: 215-222

View in Article

- PubMed
- Google Scholar
- 34. Rovin J.D.
 - Frierson Jr., H.F.
 - Ledinh W.
 - Parsons J.T.
 - Adams R.B.

Expression of focal adhesion kinase in normal and pathologic human prostate tissues.

Prostate. 2002; 53: 124-132

View in Article

- Scopus (67)
- PubMed
- Crossref
- Google Scholar
- 35. Livasy C.A.
 - Moore D.
 - o Cance W.G.
 - Lininger R.A.

Focal adhesion kinase overexpression in endometrial neoplasia.

Appl Immunohistochem Mol Morphol. 2004; 12: 342-345

View in Article

- <u>Scopus (44)</u>
- PubMed
- Crossref
- Google Scholar
- 36. McLean G.W.
 - Avizienyte E.
 - Frame M.C.

Focal adhesion kinase as a potential target in oncology.

Expert Opin Pharmacother. 2003; 4: 227-234

View in Article

- <u>Scopus (54)</u>
- PubMed
- Crossref
- Google Scholar
- 37. Duxbury M.S.
 - Ito H
 - Zinner M.J.
 - Ashley S.W.
 - Whang E.E.

Focal adhesion kinase gene silencing promotes anoikis and suppresses metastasis of human pancreatic adenocarcinoma cells.

Surgery. 2004; 135: 555-562

View in Article

- <u>Scopus (128)</u>
- PubMed
- Abstract
- Full Text
- Full Text PDF
- Google Scholar
- 38. Oktay M.H.
 - Oktay K.
 - Hamele-Bena D.
 - Buyuk A.
 - Koss L.G.

Focal adhesion kinase as a marker of malignant phenotype in breast and cervical carcinomas. *Hum Pathol.* 2003; 34: 240-245

View in Article

- <u>Scopus (107)</u>
- PubMed
- Abstract
- Full Text
- Full Text PDF
- Google Scholar
- 39. Itoh S.
 - Maeda T.
 - Shimada M.
 - Aishima S.
 - Shirabe K.
 - Tanaka S.
 - o et al.

Role of expression of focal adhesion kinase in progression of hepatocellular carcinoma.

Clin Cancer Res. 2004; 10: 2812-2817

View in Article

- Scopus (109)
- PubMed
- Crossref
- Google Scholar
- 40. Ilic D.
 - Kovacic B.
 - McDonagh S.
 - Jin F.
 - Baumbusch C.
 - Gardner D.G.
 - o et al.

Focal adhesion kinase is required for blood vessel morphogenesis.

Circ Res. 2003; 92: 255-257

- <u>Scopus (99)</u>
- PubMed
- Crossref
- Google Scholar
- 41. Haskell H.
 - Natarajan M.
 - Hecker T.P.
 - o Ding Q.
 - Stewart Jr., J.
 - Grammer J.R.
 - et al.

Focal adhesion kinase is expressed in the angiogenic blood vessels of malignant astrocytic tumors *in vivo* and promotes capillary tube formation of brain microvascular endothelial cells.

Clin Cancer Res. 2003; 9: 2157-2165

View in Article

- PubMed
- Google Scholar
- 42. Kornberg L.J.

Focal adhesion kinase and its potential involvement in tumor invasion and metastasis.

Head Neck. 1998; 20: 745-752

View in Article

- Scopus (146)
- PubMed
- Crossref
- Google Scholar
- 43. McLean G.W.
 - Carragher N.O.
 - Avizienyte E.
 - Evans J.
 - Brunton V.G.
 - Frame M.C.

The role of focal-adhesion kinase in cancer—a new therapeutic opportunity.

Nat Rev Cancer. 2005; 5: 505-515

View in Article

- <u>Scopus (821)</u>
- PubMed
- Crossref
- Google Scholar

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