

SHORT REPORT

Proposal for a unified CCN nomenclature

D R Brigstock, R Goldschmeding, K-i Katsube, S C-T Lam, L F Lau, K Lyons, C Naus, B Perbal, B Riser, M Takigawa, H Yeger

J Clin Pathol: Mol Pathol 2003;**56**:127–128

A proposal is put forth to unify the nomenclature of the CCN family of secreted, cysteine rich regulatory proteins. In the order of their description in the literature, CCN1 (CYR61), CCN2 (CTGF), CCN3 (NOV), CCN4 (WISP-1), CCN5 (WISP-2), and CCN6 (WISP-3) constitute a family of matricellular proteins that regulate cell adhesion, migration, proliferation, survival, and differentiation, at least in part through integrin mediated mechanisms. This proposal is endorsed by the International CCN Society and will serve to eliminate confusion from the multiple names that have been given to these molecules.

The members of a family of cysteine rich matricellular proteins have recently emerged as multifunctional regulators that control diverse cellular processes and play important roles in vascular and skeletal development.¹ Known as CCN proteins, members of this protein family are characterised by an N-terminal secretory signal, followed by four structural domains with partial sequence identity to insulin-like growth factor binding proteins, von Willebrand factor type C repeat, thrombospondin type 1 repeat, and a C-terminus with sequence similarity to the C-termini of von Willebrand factor, mucin, and slit.^{2–5} The acronym CCN comes from the first three members of the family reported, namely CYR61 (cysteine rich 61), CTGF (connective tissue growth factor), and NOV (nephroblastoma overexpressed).^{6–8} Other members of the CCN family have also been identified, bringing the total number of known CCN proteins to six.⁹

Genes encoding CCN proteins have also been identified in other studies, either based on their differential expression in various tissues, tumour cell lines, or upon induction by growth factors or morphogens. Consequently, a variety of other names have been assigned to these genes that reflected the context of these studies. During the international workshop on the CCN family of genes in Saint-Malo, France, it was recognised that a unified nomenclature will serve to eliminate confusion in the literature.¹ A consensus was reached to propose a unifying nomenclature for the CCN family, numbering them CCN1 through to CCN6 in the order in which they were described in

the literature. Thus, CYR61 will be designated CCN1, CTGF as CCN2, NOV as CCN3, and WISP-1–3 as CCN4–6 (table 1). This proposed nomenclature is endorsed by the International CCN Society (<http://www.ccn-society.jussieu.fr>). The board of the International CCN Society will be acting as a nomenclature committee for the attribution of CCN names to new member genes of the CCN family yet to be discovered.

Authors' affiliations

D R Brigstock, Department of Surgery, Ohio State University, Columbus, Ohio 43205, USA

R Goldschmeding, Pathology H04.312, University Hospital Utrecht, Utrecht, The Netherlands

K-i Katsube, Molecular Pathology, Graduate School of Tokyo Medical and Dental University, Yushima, Tokyo, 113–8549 Japan

S C-T Lam, Department of Pharmacology, University of Illinois, Chicago, IL 60612, USA

L F Lau, Department of Molecular Genetics, University of Illinois, Chicago, IL 60607, USA

K Lyons, Department of Molecular, Cell and Developmental Biology, UCLA, Los Angeles, CA 90095, USA

C Naus, Department of Anatomy, University of British Columbia, Vancouver V6T 1Z3, Canada

B Perbal, Laboratoire d'Oncologie Virale et Moléculaire, Université Paris 7-D, 75005 Paris, France

B Riser, Baxter Healthcare Renal Division, Magaw Park, IL 60085, USA

M Takigawa, Department of Biochemistry and Molecular Dentistry Okayama University Graduate School of Medicine and Dentistry, Okayama, 700–8525, Japan

H Yeger, Department of Pediatric Laboratory Medicine, Hospital for Sick Children, Toronto M5G 1X8, Canada

Correspondence to: Professor B Perbal, Laboratoire d'Oncologie Virale et Moléculaire, Université Paris 7-D, 75005 Paris, France; perbal@ccr.jussieu.fr

Accepted for publication 31 January 2003

REFERENCES

- 1 **Perbal B**, Brigstock DR, Lau LF. Report on the second international workshop on the CCN family of genes. *Mol Pathol* 2003;**56**:80–85.
- 2 **Bork P**. The modular architecture of a new family of growth regulators related to connective tissue growth factor. *FEBS Lett* 1993;**327**:125–30.
- 3 **Brigstock DR**. The connective tissue growth factor/cysteine-rich 61/nephroblastoma overexpressed (CCN) family. *Endocr Rev* 1999;**20**:189–206.
- 4 **Lau LF**, Lam SC-T. The CCN family of angiogenic regulators: the integrin connection. *Exp Cell Res* 1999;**248**:44–57.

Table 1 Nomenclature proposal

Proposed name	Names used previously
CCN1	CYR61 (human, mouse, xenopus), CEF10 (chicken), IGFBP-rP4 (human), BIG-M1 (mouse), CTGF-2, IGFBP10 (human), angiopro
CCN2	CTGF (human, mouse, chicken, xenopus), BIG-M2 (mouse), FISP12 (mouse), IGFBP-rP2 (human), Hcs24 (human), IGFBP8 (human), HBGF-0.8, ecogenin (human)
CCN3	NOV (human, rat, chicken, mouse, quail), IGFBP-rP3 (human), IGFBP9 (human), NOVH (human), NOVm, mNOV (mouse), xNOV (xenopus)
CCN4	WISP-1 (human), ELM-1
CCN5	WISP-2 (human), CTGF-L, CTGF-3, HICP, rCOP-1 (rat)
CCN6	WISP-3 (human)

- 5 **Perbal B.** NOV (nephroblastoma overexpressed) and the CCN family of genes: structural and functional issues. *Mol Pathol* 2001;**54**:57–79.
- 6 **O'Brien TP,** Yang GP, Sanders L, *et al.* Expression of *cyr61*, a growth factor-inducible immediate early gene. *Mol Cell Biol* 1990;**10**:3569–77.
- 7 **Bradham DM,** Igarashi A, Potter RL, *et al.* Connective tissue growth factor: a cysteine-rich mitogen secreted by human vascular endothelial cells is related to the SRC-induced immediate early gene product CEF-10. *J Cell Biol* 1991;**114**:1285–94.
- 8 **Joliet V,** Martinerie C, Dambrine G, *et al.* Proviral rearrangements and overexpression of a new cellular gene (*nov*) in myeloblastosis-associated virus type 1-induced nephroblastomas. *Mol Cell Biol* 1992;**12**:10–21.
- 9 **Pennica D,** Swanson TA, Welsh JW, *et al.* WISP genes are members of the connective tissue growth factor family that are up-regulated in *wnt-1*-transformed cells and aberrantly expressed in human colon tumors. *Proc Natl Acad Sci U S A* 1998;**95**:14717–22.

Reference linking to full text of more than 300 journals

Toll free links

You can access the FULL TEXT of articles cited in *Molecular Pathology* online if the citation is to one of the more than 300 journals hosted by HighWire (<http://highwire.stanford.edu>) without a subscription to that journal. There are also direct links from references to the Medline abstract for other titles.

www.molpath.com