

# **Gene Section**

Mini Review

# **CBP (CREB-binding protein)**

## Jean-Loup Huret

Genetics, Dept Medical Information, University of Poitiers, CHU Poitiers Hospital, F-86021 Poitiers, France (JLH)

Published in Atlas Database: July 2000

Online updated version: http://AtlasGeneticsOncology.org/Genes/CBPID42.html

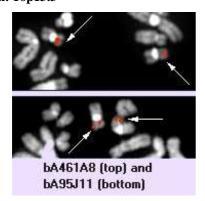
DOI: 10.4267/2042/37634

This work is licensed under a Creative Commons Attribution-Noncommercial-No Derivative Works 2.0 France Licence. © 2000 Atlas of Genetics and Cytogenetics in Oncology and Haematology

## **Identity**

Other names: CREBBP (CREB binding protein (Rubinstein-Taybi syndrome)); RTS (Rubinstein-

Taybi syndrome); RSTS HGNC (Hugo): CREBBP Location: 16p13.3



CBP (16p22) - Courtesy Mariano Rocchi, Resources for Molecular Cytogenetics.

## DNA/RNA

## Description

The gene spans about 190 kb; transcription from centromere to telomere.

## **Transcription**

8.7 kb mRNA, with a 7.3 kb coding sequence.

#### **Protein**

#### Description

2442 amino acids; 265 kDa; from NH2-term, is made

of a CREB-Binding domain, a bromodomain. Cystidine/Histidine-rich domains, and Glutamine-rich domains in COOH-term.

## Expression

Wide expression; expression in the whole embryo as well.

#### Localisation

Nucleus.

#### **Function**

Binds specifically to the DNA-binding protein CREB and connects it to the basal transcriptional machinery: transcription coactivator, with P300; has histone acetyltransferase activity; essential role in embryogenesis, cell differentiation, apoptosis, and proliferation; involved in the regulation of cell cycle during G1/S transition.

#### Homology

P300.

## Implicated in

## t(8;16)(p11;p13)/M4 ANLL --> MOZ/CBP

#### Disease

Acute non lymphocytic leukemia (ANLL) and treatment related ANLL (t-ANLL).

#### **Prognosis**

Poor: remission is obtained in half cases; survival is often less than 1 year.

#### Cytogenetics

+8 as an additional anomalies in half cases.

### Hybrid/Mutated gene

5' MOZ - 3' CBP.

#### **Abnormal protein**

N-term finger motifs and acetyl transferase from MOZ fused to most of CBP, with a breakpoint in 5' of the CREB binding domain of CBP.

## t(11;16)(q23;p13)/t-ANLL --> MLL/CBP

#### Disease

Treatment related ANLL (t-ANLL); should be very close to the t(11;22)(q23;q13).

#### **Prognosis**

Likely to be poor.

#### Hybrid/Mutated gene

5' MLL - 3' CBP.

## **Abnormal protein**

N-term AT hook and DNA methyltransferase from MLL fused to most of CBP; variable breakpoint in CBP: either in 5' of the CREB binding domain (like in the t(8;16)), or just upstream of the bromodomain.

## Rubinstein-taybi syndrome

#### Note

Due to CBP haploinsufficiency.

#### Disease

Autosomal dominant disorder with mental retardation, facial dysmorphia, broad thumbs/halluces, cardiac anomalies, and an increased risk of medulloblastoma, meningioma, and neuroblastoma.

## References

Borrow J, Stanton VP Jr, Andresen JM, Becher R, Behm FG, Chaganti RS, Civin CI, Disteche C, Dubé I, Frischauf AM, Horsman D, Mitelman F, Volinia S, Watmore AE, Housman DE. The translocation t(8;16)(p11;p13) of acute myeloid leukaemia fuses a putative acetyltransferase to the CREB-binding protein. Nat Genet. 1996 Sep;14(1):33-41

Eckner R. p300 and CBP as transcriptional regulators and targets of oncogenic events. Biol Chem. 1996 Nov;377(11):685-8

Giles RH, Petrij F, Dauwerse HG, den Hollander Al, Lushnikova T, van Ommen GJ, Goodman RH, Deaven LL, Doggett NA, Peters DJ, Breuning MH. Construction of a 1.2-Mb contig surrounding, and molecular analysis of, the human CREB-binding protein (CBP/CREBBP) gene on chromosome 16p13.3. Genomics. 1997 May 15;42(1):96-114

Goldman PS, Tran VK, Goodman RH. The multifunctional role of the co-activator CBP in transcriptional regulation. Recent Prog Horm Res. 1997;52:103-19; discussion 119-20

Sobulo OM, Borrow J, Tomek R, Reshmi S, Harden A, Schlegelberger B, Housman D, Doggett NA, Rowley JD, Zeleznik-Le NJ. MLL is fused to CBP, a histone acetyltransferase, in therapy-related acute myeloid leukemia with a t(11;16)(q23;p13.3). Proc Natl Acad Sci U S A. 1997 Aug 5;94(16):8732-7

Taki T, Sako M, Tsuchida M, Hayashi Y. The t(11;16)(q23;p13) translocation in myelodysplastic syndrome fuses the MLL gene to the CBP gene. Blood. 1997 Jun 1;89(11):3945-50

Giordano A, Avantaggiati ML. p300 and CBP: partners for life and death. J Cell Physiol. 1999 Nov;181(2):218-30

Partanen A, Motoyama J, Hui CC. Developmentally regulated expression of the transcriptional cofactors/histone acetyltransferases CBP and p300 during mouse embryogenesis. Int J Dev Biol. 1999 Sep;43(6):487-94

Yuan ZM, Huang Y, Ishiko T, Nakada S, Utsugisawa T, Shioya H, Utsugisawa Y, Shi Y, Weichselbaum R, Kufe D. Function for p300 and not CBP in the apoptotic response to DNA damage. Oncogene. 1999 Oct 7;18(41):5714-7

Ait-Si-Ali S, Polesskaya A, Filleur S, Ferreira R, Duquet A, Robin P, Vervish A, Trouche D, Cabon F, Harel-Bellan A. CBP/p300 histone acetyl-transferase activity is important for the G1/S transition. Oncogene. 2000 May 11;19(20):2430-7

This article should be referenced as such:

Huret JL. CBP (CREB-binding protein). Atlas Genet Cytogenet Oncol Haematol. 2000; 4(3):101-102.