# Package 'curatedOvarianData'

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Type Package

Title Clinically Annotated Data for the Ovarian Cancer Transcriptome

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**Description** The curatedOvarianData package provides data for gene expression analysis in patients with ovarian cancer.

**Depends** R (>= 2.10.0), affy

Imports BiocGenerics

Suggests survival, RUnit, metafor, genefilter, futile.logger, sva,xtable

License Artistic-2.0

URL http://bcb.dfci.harvard.edu/ovariancancer

biocViews ExperimentData, Cancer, Ovarian, RNAExpressionData

## **R** topics documented:

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curatedOvarianData-package

Clinically Annotated Data for the Ovarian Cancer Transcriptome

## Description

The curatedOvarianData package provides manually curated clinical data, uniformly processed expression data, and convenience functions for gene expression analysis in patients with ovarian cancer.

## Details

curatedOvarianData
Package
1.0.5
2013-02-22
Artistic-2.0
R (>= 2.10.0), affy

Please see http://bcb.dfci.harvard.edu/ovariancancer for alterative versions of this package, differing in how redundant probe sets are dealt with. In the curatedOvarianData version, each gene is represented by the gene with maximum mean. In NormalizerVcuratedOvarianData, each gene is represented by the mean of the probesets after removing "noisy" probesets (see the Normalizer function of the Sleipnir library for computational biology), and in FULLVcuratedOvarianData, no collapsing of probe sets is done, but a map is provided to allow the user to do so by their method of choice through featureData(eset).

#### curatedOvarianData-package

In the "Available sample meta-data" sections of each dataset, please refer to the following key.

For "sample\_type": tumor / metastatic / adjacentnormal / healthy / cellline: "healthy" should be only from individuals without cancer, "adjacentnormal" from individuals with cancer, "metastatic" for non-primary tumors.

For "histological\_type": ser=serous / endo=endometrioid / clearcell / mucinous, undifferentiated / other / mix. Other includes sarcomatoid, adenocarcinoma, dysgerminoma.

For "primarysite" and for "arrayedsite": ovlftlother. ov=ovary;ft=fallopian tube

For "summarygrade": low = 1, 2, LMP. High= 3,4,23.

For "summarystage": early = 1,2, 12. late=3,4,23,34.

For "tumorstage": FIGO Stage (I-IV, but coded here as 1-4 to ensure correct ordering in factors). If multiple stages given (eg 34), use the highest.

For "substage": substage (abcd). For cases like ab, bc, use highest given.

For "grade": Grade (1-3): If multiple given, ie 12, 23, use highest given. Most ovarian cancer studies use FIGO grading, with a couple exceptions in this package (Yoshihara and Tothill).

For "pltx": (y/n): patient treated with platin.

For "tax": (y/n): patient treated with taxol.

For "neo": (y/n): patient treated with neoadjuvant treatment.

For "primary\_therapy\_outcome\_success": completeresponselpartialresponselprogressivediseaselstabledisease: response to any kind of therapy (including radiation only).

For "days\_to\_tumor\_recurrence": time to recurrence or last follow-up in days

For "recurrence\_status": recurrence censoring variable (recurrence / norecurrence)

For "days\_to\_death": time to death or last follow-up in days

For "vital\_status": Overall survival censoring variable (living / deceased)

For "os\_binary": dichotomized overall survival variable as defined by study authors (short / long).

For "relapse\_binary": dichotomized relapse variable as defined by study authors (short / long)

For "site\_of\_tumor\_first\_recurrence": (metastasis / locoregional / none / locoregional\_plus\_metastatic). none for no recurrence, na for unknown

For "primary\_therapy\_outcome\_success": (completeresponse / partialresponse / progressivedisease / stabledisease) Response to any kind of therapy (including radiation only).

For "debulking": amount of residual disease (optimal = <1cm, suboptimal=>1cm).

For "percent\_normal\_cells": Estimated percentage of normal cells. An integer 0-100, or can be >70, <70, etc.

For "percent\_stromal\_cells": Estimated percentage of stromal cells. An integer 0-100, or can be >70, <70, etc.

For "percent\_tumor\_cells": Estimated percentage of tumor cells. An integer 0-100, or can be >70, <70, etc.

For "batch": batch variable when available. Hybridization date when Affymetrix CEL files are available.

For "uncurated\_author\_metadata": Original uncurated data, with each field separated by ///.

#### Author(s)

Benjamin F. Ganzfried, Steve Skates, Markus Riester, Victoria Wang, Thomas Risch, Benjamin Haibe-Kains, Curtis Huttenhower, Svitlana Tyekucheva, Jie Ding, Ina Jazic, Michael Birrer, Giovanni Parmigiani, Levi Waldron

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Maintainer: Levi Waldron <levi@jimmy.harvard.edu>

#### Examples

```
##List all datasets:
data(package="curatedOvarianData")
##
##See the actual template used for syntax checking of clinical metadata:
template.file <- system.file("extdata/template_ov.csv", package = "curatedOvarianData")
template <- read.csv(template.file, as.is=TRUE)
head(template)
```

```
E.MTAB.386_eset
```

Angiogenic mRNA and microRNA gene expression signature predicts a novel subtype of serous ovarian cancer.

#### Description

Ovarian cancer is the fifth leading cause of cancer death for women in the U.S. and the seventh most fatal worldwide. Although ovarian cancer is notable for its initial sensitivity to platinum-based therapies, the vast majority of patients eventually develop recurrent cancer and succumb to increasingly platinum-resistant disease. Modern, targeted cancer drugs intervene in cell signaling, and identifying key disease mechanisms and pathways would greatly advance our treatment abilities. In order to shed light on the molecular diversity of ovarian cancer, we performed comprehensive transcriptional profiling on 129 advanced stage, high grade serous ovarian cancers. We implemented a, re-sampling based version of the ISIS class discovery algorithm (rISIS: robust ISIS) and applied it to the entire set of ovarian cancer transcriptional profiles. rISIS identified a previously undescribed patient stratification, further supported by micro-RNA expression profiles, and gene set enrichment analysis found strong biological support for the stratification by extracellular matrix, cell adhesion, and angiogenesis genes. The corresponding "angiogenesis signature" was validated in ten published independent ovarian cancer gene expression datasets and is significantly associated with overall survival. The subtypes we have defined are of potential translational interest as they may be relevant for identifying patients who may benefit from the addition of anti-angiogenic therapies that are now being tested in clinical trials.

#### Usage

data( E.MTAB.386\_eset )

## Format

```
experimentData(eset):
Experiment data
 Experimenter name: Bentink S, Haibe-Kains B, Risch T, Fan J-B, Hirsch MS, Holton K, Rubio R, April C, Ch
 Laboratory: Bentink, Matulonis 2012
 Contact information:
 Title: Angiogenic mRNA and microRNA gene expression signature predicts a novel subtype of serous ovaria
 URL:
 PMIDs: 22348002
 Abstract: A 212 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
      Illumina humanRef-8 v2.0 expression beadchip
   platform_shorttitle:
      Illumina humanRef-8 v2.0
   platform_summary:
      illuminaHumanv2
   platform_manufacturer:
      Illumina
   platform_distribution:
      commercial
   platform_accession:
      GPL6104
Preprocessing: default
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A2M A4GALT ... ZZEF1 (10327 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
unique_patient_ID:
  Length
           Class
                       Mode
     129 character character
sample_type:
tumor
 129
histological_type:
ser
129
primarysite:
ov
129
summarygrade:
high
129
summarystage:
early late
   1 128
tumorstage:
 2 3 4
 1 109 19
substage:
  a b
           c NAs
  5 12 93 19
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                        Max.
  21.00 50.00
               66.00
                        60.71 72.00
                                       95.00
days_to_death:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                        Max.
   3.9 516.9 917.1 1007.0 1401.0 2724.0
vital_status:
deceased living
     73
              56
debulking:
  optimal suboptimal
                          NAs
       98
                  28
                             3
```

uncurated\_author\_metadata: Length Class Mode 129 character character

GSE12418\_eset

*Expression analysis of stage III serous ovarian adenocarcinoma distinguishes a sub-group of survivors.* 

#### Description

It is difficult to predict the clinical outcome for patients with ovarian cancer. However, the use of biomarkers as additional prognostic factors may improve the outcome for these patients. In order to find novel candidate biomarkers, differences in gene expressions were analysed in 54 stage III serous ovarian adenocarcinomas with oligonucleotide microarrays containing 27,000 unique probes. The microarray data was verified with quantitative real-time polymerase chain reaction for the genes TACC1, MUC5B and PRAME. Using hierarchical cluster analysis we detected a subgroup that included 60% of the survivors. The gene expressions in tumours from patients in this sub-group of survivors were compared with the remaining tumours, and 204 genes were found to be differently expressed. We conclude that the sub-group of survivors might represent patients with favourable tumour biology and sensitivity to treatment. A selection of the 204 genes might be used as a predictive model to distinguish patients within and outside of this group. Alternative chemotherapy strategies could then be offered as first-line treatment, which may lead to improvements in the clinical outcome for these patients.

#### Usage

data( GSE12418\_eset )

#### Format

```
experimentData(eset):
Experiment data
Experimenter name: Partheen K, Levan K, Osterberg L, Horvath G.Expression analysis of stage III serous
Laboratory: Partheen, Horvath 2006
Contact information:
Title: Expression analysis of stage III serous ovarian adenocarcinoma distinguishes a sub-group of sur
URL:
PMIDs: 16996261
Abstract: A 177 word abstract is available. Use abstract method.
Information is available on: preprocessing
notes:
    platform_title:
        SWEGENE H_v2.1.1_27k
    platform_shorttitle:
        SWEGENE H_v2.1.1_27k
```

```
platform_summary:
    PartheenMetaData
platform_manufacturer:
    other
platform_distribution:
    non-commercial
platform_accession:
    GPL5886
```

```
Preprocessing: default
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1CF A2LD1 ... ZZZ3 (12656 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 12656 features, 54 samples
Platform type: PartheenMetaData
Binary overall survival summary (definitions of long and short provided by study authors):
long short
  20 34
_____
Available sample meta-data:
_____
alt_sample_name:
  Length Class Mode
      54 character character
sample_type:
tumor
  54
histological_type:
ser
54
primarysite:
ov
54
summarystage:
late
 54
```

```
tumorstage:
 3
54
substage:
b c
19 35
age_at_initial_pathologic_diagnosis:
   Min. 1st Qu. Median
                            Mean 3rd Qu.
                                             Max.
  35.00
          51.25
                   59.50
                           59.56
                                   69.75
                                            84.00
pltx:
У
54
os_binary:
 long short
   20
         34
debulking:
   optimal suboptimal
        13
                    41
uncurated author metadata:
   Length
              Class
                          Mode
       54 character character
```

GSE12470\_eset

Gene expression profiling of advanced-stage serous ovarian cancers distinguishes novel subclasses and implicates ZEB2 in tumor progression and prognosis.

## Description

To elucidate the mechanisms of rapid progression of serous ovarian cancer, gene expression profiles from 43 ovarian cancer tissues comprising eight early stage and 35 advanced stage tissues were carried out using oligonucleotide microarrays of 18,716 genes. By non-negative matrix factorization analysis using 178 genes, which were extracted as stage-specific genes, 35 advanced stage cases were classified into two subclasses with superior (n = 17) and poor (n = 18) outcome evaluated by progression-free survival (log rank test, P = 0.03). Of the 178 stage-specific genes, 112 genes were identified as showing different expression between the two subclasses. Of the 48 genes selected for biological function by gene ontology analysis or Ingenuity Pathway Analysis, five genes (ZEB2, CDH1, LTBP2, COL16A1, and ACTA2) were extracted as candidates for prognostic factors associated with progression-free survival. The relationship between high ZEB2 or low CDH1 expression and shorter progression-free survival was validated by real-time RT-PCR experiments of 37 independent advanced stage cancer samples. ZEB2 expression was negatively correlated with CDH1 expression in advanced stage samples, whereas ZEB2 knockdown in ovarian adenocarcinoma SKOV3 cells resulted in an increase in CDH1 expression. Multivariate analysis showed that high ZEB2 expression was independently associated with poor prognosis. Furthermore, the prognostic effect of E-cadherin encoded by CDH1 was verified using immunohistochemical analysis of an independent advanced stage cancer samples set (n = 74). These findings suggest that the expression of epithelial-mesenchymal transition-related genes such as ZEB2 and CDH1 may play important roles in the invasion process of advanced stage serous ovarian cancer.

#### Usage

```
data( GSE12470_eset )
```

## Format

```
experimentData(eset):
Experiment data
 Experimenter name: Yoshihara K, Tajima A, Komata D, Yamamoto T, Kodama S, Fujiwara H, Suzuki M, Onishi `
 Laboratory: Yoshihara, Tanaka 2009
 Contact information:
 Title: Gene expression profiling of advanced-stage serous ovarian cancers distinguishes novel subclass
 URL:
 PMIDs: 19486012
  Abstract: A 253 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
      Agilent-012097 Human 1A Microarray (V2) G4110B (Feature Number version)
   platform_shorttitle:
      Agilent G4110B
   platform_summary:
      hgug4110b
   platform_manufacturer:
      Agilent
   platform_distribution:
      commercial
   platform_accession:
      GPL887
Preprocessing: default
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1BG A1CF ... ZZZ3 (16880 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

## Details

```
assayData: 16880 features, 53 samples
Platform type: hgug4110b
_____
Available sample meta-data:
-----
alt_sample_name:
  Length
            Class
                       Mode
      53 character character
sample_type:
healthy
         tumor
    10
           43
histological_type:
ser NAs
 43
      10
primarysite:
ov
53
summarystage:
early late NAs
   8
        35
             10
tumorstage:
  1 NAs
  8
      45
uncurated_author_metadata:
  Length
            Class
                      Mode
      53 character character
```

GSE13876\_eset

Survival-related profile, pathways, and transcription factors in ovarian cancer.

## Description

Ovarian cancer has a poor prognosis due to advanced stage at presentation and either intrinsic or acquired resistance to classic cytotoxic drugs such as platinum and taxoids. Recent large clinical trials with different combinations and sequences of classic cytotoxic drugs indicate that further significant improvement in prognosis by this type of drugs is not to be expected. Currently a large

number of drugs, targeting dysregulated molecular pathways in cancer cells have been developed and are introduced in the clinic. A major challenge is to identify those patients who will benefit from drugs targeting these specific dysregulated pathways. The aims of our study were (1) to develop a gene expression profile associated with overall survival in advanced stage serous ovarian cancer, (2) to assess the association of pathways and transcription factors with overall survival, and (3) to validate our identified profile and pathways/transcription factors in an independent set of ovarian cancers. According to a randomized design, profiling of 157 advanced stage serous ovarian cancers was performed in duplicate using approximately 35,000 70-mer oligonucleotide microarrays. A continuous predictor of overall survival was built taking into account well-known issues in microarray analysis, such as multiple testing and overfitting. A functional class scoring analysis was utilized to assess pathways/transcription factors for their association with overall survival. The prognostic value of genes that constitute our overall survival profile was validated on a fully independent, publicly available dataset of 118 well-defined primary serous ovarian cancers. Furthermore, functional class scoring analysis was also performed on this independent dataset to assess the similarities with results from our own dataset. An 86-gene overall survival profile discriminated between patients with unfavorable and favorable prognosis (median survival, 19 versus 41 mo, respectively; permutation p-value of log-rank statistic = 0.015) and maintained its independent prognostic value in multivariate analysis. Genes that composed the overall survival profile were also able to discriminate between the two risk groups in the independent dataset. In our dataset 17/167 pathways and 13/111 transcription factors were associated with overall survival, of which 16 and 12, respectively, were confirmed in the independent dataset. Our study provides new clues to genes, pathways, and transcription factors that contribute to the clinical outcome of serous ovarian cancer and might be exploited in designing new treatment strategies.

#### Usage

data( GSE13876\_eset )

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Crijns AP, Fehrmann RS, de Jong S, Gerbens F, Meersma GJ, Klip HG, Hollema H, Hofstra
 Laboratory: Crijns, van der Zee 2009
 Contact information:
 Title: Survival-related profile, pathways, and transcription factors in ovarian cancer.
 URL:
 PMIDs: 19192944
 Abstract: A 371 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
      Operon human v3 ~35K 70-mer two-color oligonucleotide microarrays
  platform_shorttitle:
     Operon v3 two-color
  platform_summary:
     OperonHumanV3
  platform_manufacturer:
```

other platform\_distribution: non-commercial platform\_accession: GPL7759

Preprocessing: default
featureData(eset):
An object of class AnnotatedDataFrame
 featureNames: A1BG A1CF ... ZZZ3 (20505 total)
 varLabels: probeset gene
 varMetadata: labelDescription

```
assayData: 20505 features, 157 samples
Platform type: OperonHumanV3
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
       n.max n.start events median 0.95LCL 0.95UCL
records
157.00 157.00 157.00 113.00
                               2.05 1.56 2.71
_____
Available sample meta-data:
_____
alt_sample_name:
151 NAs
  1 156
unique_patient_ID:
  Min. 1st Qu. Median
                     Mean 3rd Qu.
                                      Max.
     1
           40 79
                        79
                              118
                                      157
sample_type:
tumor
 157
histological_type:
ser
157
primarysite:
ov
157
summarygrade:
```

```
high
      low NAs
  85
       59
             13
summarystage:
late
157
grade:
        2
   1
              3
                   4 NAs
  14
       45
             82
                   3
                       13
age_at_initial_pathologic_diagnosis:
   Min. 1st Qu.
                  Median
                             Mean 3rd Qu.
                                               Max.
                   60.00
  21.00
          50.00
                            57.95
                                     67.00
                                              84.00
days_to_death:
   Min. 1st Qu.
                  Median
                             Mean 3rd Qu.
                                               Max.
     30
             360
                      630
                             1100
                                      1470
                                               7020
vital_status:
deceased
            living
     113
                44
uncurated_author_metadata:
   Length
               Class
                           Mode
      157 character character
```

GSE14764\_eset

A prognostic gene expression index in ovarian cancer - validation across different independent data sets.

#### Description

Ovarian carcinoma has the highest mortality rate among gynaecological malignancies. In this project, we investigated the hypothesis that molecular markers are able to predict outcome of ovarian cancer independently of classical clinical predictors, and that these molecular markers can be validated using independent data sets. We applied a semi-supervised method for prediction of patient survival. Microarrays from a cohort of 80 ovarian carcinomas (TOC cohort) were used for the development of a predictive model, which was then evaluated in an entirely independent cohort of 118 carcinomas (Duke cohort). A 300-gene ovarian prognostic index (OPI) was generated and validated in a leave-one-out approach in the TOC cohort (Kaplan-Meier analysis, p = 0.0087). In a second validation step, the prognostic power of the OPI was confirmed in an independent data set (Duke cohort, p = 0.0063). In multivariate analysis, the OPI was independent of the post-operative residual tumour, the main clinico-pathological prognostic parameter with an adjusted hazard ratio of 6.4 (TOC cohort, CI 1.8-23.5, p = 0.0049) and 1.9 (Duke cohort, CI 1.2-3.0, p = 0.0068). We constructed a combined score of molecular data (OPI) and clinical parameters (residual tumour), which

was able to define patient groups with highly significant differences in survival. The integrated analysis of gene expression data as well as residual tumour can be used for optimized assessment of the prognosis of platinum-taxol-treated ovarian cancer. As traditional treatment options are limited, this analysis may be able to optimize clinical management and to identify those patients who would be candidates for new therapeutic strategies.

## Usage

```
data( GSE14764_eset )
```

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Denkert C, Budczies J, Darb-Esfahani S, Gy??rffy B et al. A prognostic gene expressi
 Laboratory: Denkert, Lage 2009
  Contact information:
 Title: A prognostic gene expression index in ovarian cancer - validation across different independent of
  URL:
  PMIDs: 19294737
  Abstract: A 254 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133A] Affymetrix Human Genome U133A Array
   platform_shorttitle:
      Affymetrix HG-U133A
   platform_summary:
      hgu133a
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL96
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1CF A2M ... ZZZ3 (12981 total)
  varLabels: probeset gene
```

```
varMetadata: labelDescription
```

#### Details

assayData: 12981 features, 80 samples Platform type: hgu133a

```
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
        n.max n.start events median 0.95LCL 0.95UCL
records
 80.00 80.00 80.00 21.00 4.52
                                    4.19
                                             NA
_____
Available sample meta-data:
_____
alt_sample_name:
  Min. 1st Qu. Median Mean 3rd Qu.
                                     Max.
  1.00 20.75 40.50 40.50 60.25
                                    80.00
sample_type:
tumor
  80
histological_type:
  Length
          Class
                     Mode
      80 character character
primarysite:
ov
80
summarygrade:
high low
 54 26
summarystage:
early late
   9 71
tumorstage:
1 2 3 4
8 1 69 2
substage:
  a b
         c NAs
      6 32 38
  4
grade:
1 2 3
3 23 54
recurrence_status:
norecurrence recurrence
                             NAs
```

50 26 4 days\_to\_death: Min. 1st Qu. Median Mean 3rd Qu. Max. 210 660 1050 1011 1328 2190 vital\_status: deceased living 21 59 batch: 2004-09-29 2004-09-30 2004-10-01 2005-01-21 2005-01-25 2005-01-26 2005-01-28 1 2 6 4 7 8 10 2005-03-02 2006-07-26 2006-07-27 2006-07-28 2006-08-11 2006-08-18 2006-08-19 6 4 6 4 10 3 4 2006-08-21 5 uncurated\_author\_metadata: Length Class Mode 80 character character

GSE17260\_eset

Gene expression profile for predicting survival in advanced-stage serous ovarian cancer across two independent datasets.

#### Description

Advanced-stage ovarian cancer patients are generally treated with platinum/taxane-based chemotherapy after primary debulking surgery. However, there is a wide range of outcomes for individual patients. Therefore, the clinicopathological factors alone are insufficient for predicting prognosis. Our aim is to identify a progression-free survival (PFS)-related molecular profile for predicting survival of patients with advanced-stage serous ovarian cancer. Advanced-stage serous ovarian cancer tissues from 110 Japanese patients who underwent primary surgery and platinum/taxane-based chemotherapy were profiled using oligonucleotide microarrays. We selected 88 PFS-related genes by a univariate Cox model (p<0.01) and generated the prognostic index based on 88 PFS-related genes after adjustment of regression coefficients of the respective genes by ridge regression Cox model using 10-fold cross-validation. The prognostic index was independently associated with PFS time compared to other clinical factors in multivariate analysis [hazard ratio (HR), 3.72; 95% confidence interval (CI), 2.66-5.43; p<0.0001]. In an external dataset, multivariate analysis revealed that this prognostic index was significantly correlated with PFS time (HR, 1.54; 95% CI, 1.20-1.98; p = 0.0008). Furthermore, the correlation between the prognostic index and overall survival time was confirmed in the two independent external datasets (log rank test, p = 0.0010 and 0.0008). The prognostic ability of our index based on the 88-gene expression profile in ridge regression Cox hazard model was shown to be independent of other clinical factors in predicting cancer prognosis across two distinct datasets. Further study will be necessary to improve predictive accuracy of the

prognostic index toward clinical application for evaluation of the risk of recurrence in patients with advanced-stage serous ovarian cancer.

#### Usage

data( GSE17260\_eset )

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Yoshihara K, Tajima A, Yahata T, Kodama S, Fujiwara H, Suzuki M, Onishi Y, Hatae M, S
 Laboratory: Yoshihara, Tanaka 2010
 Contact information:
 Title: Gene expression profile for predicting survival in advanced-stage serous ovarian cancer across
 URL:
  PMIDs: 20300634
  Abstract: A 257 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      Agilent-012391 Whole Human Genome Oligo Microarray G4112A
   platform_shorttitle:
      Agilent G4112A
   platform_summary:
      hgug4112a
   platform_manufacturer:
      Agilent
   platform_distribution:
      commercial
   platform_accession:
      GPL6848
Preprocessing: default
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1CF A2LD1 ... ZZZ3 (19358 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 19358 features, 110 samples
Platform type: hgug4112a
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
```

records n.max n.start events median 0.95LCL 0.95UCL 110.00 110.00 110.00 46.00 4.44 4.03 NA \_\_\_\_\_ Available sample meta-data: \_\_\_\_\_ alt\_sample\_name: Length Class Mode 110 character character sample\_type: tumor 110 histological\_type: ser 110 primarysite: ov 110 summarygrade: high low 43 67 summarystage: late 110 tumorstage: 3 4 93 17 substage: a b c NAs 6 18 69 17 grade: 1 2 3 26 41 43 pltx: у 110 tax:

```
у
110
days_to_tumor_recurrence:
   Min. 1st Qu. Median
                            Mean 3rd Qu.
                                             Max.
   30.0
          285.0
                  510.0
                           673.9
                                 870.0 2250.0
recurrence_status:
norecurrence
               recurrence
          34
                        76
days_to_death:
   Min. 1st Qu.
                 Median
                            Mean 3rd Qu.
                                             Max.
     30
            660
                    915
                            1086
                                    1530
                                             2430
vital_status:
deceased
           living
      46
               64
debulking:
   optimal suboptimal
        57
                    53
uncurated_author_metadata:
   Length
              Class
                          Mode
      110 character character
```

GSE18520\_eset

A gene signature predictive for outcome in advanced ovarian cancer identifies a survival factor: microfibril-associated glycoprotein 2.

#### Description

Advanced stage papillary serous tumors of the ovary are responsible for the majority of ovarian cancer deaths, yet the molecular determinants modulating patient survival are poorly characterized. Here, we identify and validate a prognostic gene expression signature correlating with survival in a series of microdissected serous ovarian tumors. Independent evaluation confirmed the association of a prognostic gene microfibril-associated glycoprotein 2 (MAGP2) with poor prognosis, whereas in vitro mechanistic analyses demonstrated its ability to prolong tumor cell survival and stimulate endothelial cell motility and survival via the alpha(V)beta(3) integrin receptor. Increased MAGP2 expression correlated with microvessel density suggesting a proangiogenic role in vivo. Thus, MAGP2 may serve as a survival-associated target.

#### Usage

data( GSE18520\_eset )

#### GSE18520\_eset

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Mok SC, Bonome T, Vathipadiekal V, Bell A, Johnson ME, Wong KK, Park DC, Hao K, Yip D
 Laboratory: Mok, Birrer 2009
 Contact information:
 Title: A gene signature predictive for outcome in advanced ovarian cancer identifies a survival factor
 URL:
 PMIDs: 19962670
 Abstract: A 110 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
   platform_shorttitle:
      Affymetrix HG-U133Plus2
   platform_summary:
      hgu133plus2
   platform_manufacturer:
      Affymetrix | Operon
   platform_distribution:
      commercial | non-commercial
   platform_accession:
      GPL570|GPL9216
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
```

```
An object of class AnnotatedDataFrame
featureNames: A1BG A1CF ... ZZZ3 (19093 total)
varLabels: probeset gene
varMetadata: labelDescription
```

#### Details

```
assayData: 19093 features, 63 samples
Platform type: hgu133plus2
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
```

10 observations deleted due to missingness records n.max n.start events median 0.95LCL 0.95UCL 53.00 53.00 53.00 41.00 2.05 1.48 3.70

-----

Available sample meta-data:

```
alt_sample_name:
  Min. 1st Qu. Median
                       Mean 3rd Qu.
                                       Max.
 312.0 395.0 694.0 893.3 1040.0 2237.0
sample_type:
healthy tumor
    10
            53
histological_type:
ser NAs
 53 10
primarysite:
ov
63
summarygrade:
high NAs
 53 10
summarystage:
late NAs
 53 10
tumorstage:
  3 NAs
 53 10
grade:
  3 NAs
 53 10
days_to_death:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                        Max.
                                               NAs
   150
          450 630
                        1212 1440
                                        4500
                                             10
vital_status:
                 NAs
deceased living
     41
             12
                    10
debulking:
optimal
    63
percent_normal_cells:
0
63
```

```
percent_stromal_cells:
0
63
percent_tumor_cells:
100
63
batch:
2004-03-12 2004-04-08 2004-04-09 2004-07-20 2004-08-12 2004-08-13 2004-09-30
        20
                     6
                                9
                                           11
                                                       10
                                                                    1
                                                                               6
uncurated_author_metadata:
   Length
              Class
                          Mode
       63 character character
```

GSE19829.GPL570\_eset Gene expression profile of BRCAness that correlates with responsiveness to chemotherapy and with outcome in patients with epithelial ovarian cancer.

#### Description

To define a gene expression profile of BRCAness that correlates with chemotherapy response and outcome in epithelial ovarian cancer (EOC). A publicly available microarray data set including 61 patients with EOC with either sporadic disease or BRCA(1/2) germline mutations was used for development of the BRCAness profile. Correlation with platinum responsiveness was assessed in platinum-sensitive and platinum-resistant tumor biopsy specimens from six patients with BRCA germline mutations. Association with poly-ADP ribose polymerase (PARP) inhibitor responsiveness and with radiation-induced RAD51 foci formation (a surrogate of homologous recombination) was assessed in Capan-1 cell line clones. The BRCAness profile was validated in 70 patients enriched for sporadic disease to assess its association with outcome. The BRCAness profile accurately predicted platinum responsiveness in eight out of 10 patient-derived tumor specimens, and between PARP-inhibitor sensitivity and resistance in four out of four Capan-1 clones. [corrected] When applied to the 70 patients with sporadic disease, patients with the BRCA-like (BL) profile had improved disease-free survival (34 months v 15 months; log-rank P = .013) and overall survival (72 months v 41 months; log-rank P = .006) compared with patients with a non-BRCA-like (NBL) profile, respectively. The BRCAness profile maintained independent prognostic value in multivariate analysis, which controlled for other known clinical prognostic factors. The BRCAness profile correlates with responsiveness to platinum and PARP inhibitors and identifies a subset of sporadic patients with improved outcome. Additional evaluation of this profile as a predictive tool in patients with sporadic EOC is warranted.

#### Usage

data( GSE19829.GPL570\_eset )

## Format

```
experimentData(eset):
Experiment data
 Experimenter name: Konstantinopoulos PA, Spentzos D, Karlan BY, Taniguchi T et al. Gene expression pro-
 Laboratory: Konstantinopoulos, Cannistra 2010 hgu133plus2
 Contact information:
 Title: Gene expression profile of BRCAness that correlates with responsiveness to chemotherapy and wit
 URL:
 PMIDs: 20547991
 Abstract: A 241 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
   platform_shorttitle:
      Affymetrix HG-U133Plus2
   platform_summary:
      hgu133plus2
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL570|GPL8300
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1BG A1CF ... ZZZ3 (19093 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

## Details

```
assayData: 19093 features, 28 samples
Platform type: hgu133plus2
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
records n.max n.start events median 0.95LCL 0.95UCL
28.00 28.00 28.00 17.00 3.95 2.71 NA
_______Available sample meta-data:
_______
```

alt\_sample\_name: Mode Length Class 28 character character sample\_type: tumor 28 primarysite: ov 28 days\_to\_death: Min. 1st Qu. Mean 3rd Qu. Median Max. 150 540 1050 1291 1688 3450 vital\_status: deceased living 17 11 batch: 2009-08-14 28 uncurated\_author\_metadata: Length Class Mode 28 character character

GSE19829.GPL8300\_eset Gene expression profile of BRCAness that correlates with responsiveness to chemotherapy and with outcome in patients with epithelial ovarian cancer.

## Description

To define a gene expression profile of BRCAness that correlates with chemotherapy response and outcome in epithelial ovarian cancer (EOC). A publicly available microarray data set including 61 patients with EOC with either sporadic disease or BRCA(1/2) germline mutations was used for development of the BRCAness profile. Correlation with platinum responsiveness was assessed in platinum-sensitive and platinum-resistant tumor biopsy specimens from six patients with BRCA germline mutations. Association with poly-ADP ribose polymerase (PARP) inhibitor responsiveness and with radiation-induced RAD51 foci formation (a surrogate of homologous recombination) was assessed in Capan-1 cell line clones. The BRCAness profile was validated in 70 patients enriched for sporadic disease to assess its association with outcome. The BRCAness profile accurately predicted platinum responsiveness in eight out of 10 patient-derived tumor specimens, and between PARP-inhibitor sensitivity and resistance in four out of four Capan-1 clones. [corrected] When

applied to the 70 patients with sporadic disease, patients with the BRCA-like (BL) profile had improved disease-free survival (34 months v 15 months; log-rank P = .013) and overall survival (72 months v 41 months; log-rank P = .006) compared with patients with a non-BRCA-like (NBL) profile, respectively. The BRCAness profile maintained independent prognostic value in multivariate analysis, which controlled for other known clinical prognostic factors. The BRCAness profile correlates with responsiveness to platinum and PARP inhibitors and identifies a subset of sporadic patients with improved outcome. Additional evaluation of this profile as a predictive tool in patients with sporadic EOC is warranted.

#### Usage

```
data( GSE19829.GPL8300_eset )
```

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Konstantinopoulos PA, Spentzos D, Karlan BY, Taniguchi T et al. Gene expression pro-
 Laboratory: Konstantinopoulos, Cannistra 2010 hgu95
 Contact information:
 Title: Gene expression profile of BRCAness that correlates with responsiveness to chemotherapy and wit
 URL:
 PMIDs: 20547991
  Abstract: A 241 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
      [HG_U95Av2] Affymetrix Human Genome U95 Version 2 Array
   platform_shorttitle:
      Affymetrix HG_U95Av2
   platform_summary:
      hgu95av2
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL570|GPL8300
Preprocessing: rma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: AADAC AAK1 ... ZZZ3 (8957 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

#### Details

```
assayData: 8957 features, 42 samples
Platform type: hgu95av2
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
         n.max n.start events median 0.95LCL 0.95UCL
records
 42.00
         42.00 42.00
                      23.00
                                3.78
                                        2.79
                                                 NA
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
                       Mode
            Class
      42 character character
sample_type:
tumor
  42
primarysite:
οv
42
days_to_death:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                        Max.
  30.0 727.5 1155.0 1089.0 1485.0 2040.0
vital_status:
deceased
        living
     23
             19
batch:
2001-09-14 2001-12-14 2002-08-20 2003-09-09 2003-09-18
        7
                  4
                        14
                                    13
                                                 4
uncurated_author_metadata:
  Length
            Class
                       Mode
      42 character character
```

GSE20565\_eset

A genomic and transcriptomic approach for a differential diagnosis between primary and secondary ovarian carcinomas in patients with a previous history of breast cancer.

#### Description

The distinction between primary and secondary ovarian tumors may be challenging for pathologists. The purpose of the present work was to develop genomic and transcriptomic tools to further refine the pathological diagnosis of ovarian tumors after a previous history of breast cancer. Sixteen paired breast-ovary tumors from patients with a former diagnosis of breast cancer were collected. The genomic profiles of paired tumors were analyzed using the Affymetrix GeneChip Mapping 50 K Xba Array or Genome-Wide Human SNP Array 6.0 (for one pair), and the data were normalized with ITALICS (ITerative and Alternative normaLIzation and Copy number calling for affymetrix Snp arrays) algorithm or Partek Genomic Suite, respectively. The transcriptome of paired samples was analyzed using Affymetrix GeneChip Human Genome U133 Plus 2.0 Arrays, and the data were normalized with gc-Robust Multi-array Average (gcRMA) algorithm. A hierarchical clustering of these samples was performed, combined with a dataset of well-identified primary and secondary ovarian tumors. In 12 of the 16 paired tumors analyzed, the comparison of genomic profiles confirmed the pathological diagnosis of primary ovarian tumor (n = 5) or metastasis of breast cancer (n = 7). Among four cases with uncertain pathological diagnosis, genomic profiles were clearly distinct between the ovarian and breast tumors in two pairs, thus indicating primary ovarian carcinomas, and showed common patterns in the two others, indicating metastases from breast cancer. In all pairs, the result of the transcriptomic analysis was concordant with that of the genomic analysis. In patients with ovarian carcinoma and a previous history of breast cancer, SNP array analysis can be used to distinguish primary and secondary ovarian tumors. Transcriptomic analysis may be used when primary breast tissue specimen is not available.

## Usage

data( GSE20565\_eset )

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Meyniel JP, Cottu PH, Decraene C, Stern MH, Couturier J, Lebigot I, Nicolas A, Weber
 Laboratory: Meyniel, Sastre-Garau 2010
 Contact information:
 Title: A genomic and transcriptomic approach for a differential diagnosis between primary and secondar
 URL:
 PMIDs: 20492709
 Abstract: A 277 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
  platform_shorttitle:
     Affymetrix HG-U133Plus2
  platform_summary:
```

```
hgu133plus2
```

```
platform_manufacturer:
```

```
Affymetrix
```

```
platform_distribution:
    commercial
platform_accession:
    GPL570|GPL2005|GPL6801
```

```
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1BG A1CF ... ZZZ3 (19093 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 19093 features, 140 samples
Platform type: hgu133plus2
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
           Class
                     Mode
     140 character character
sample_type:
tumor
 140
histological_type:
  Length
           Class
                     Mode
     140 character character
primarysite:
other
       ov
       96
  44
summarygrade:
high low NAs
 63
     33
         44
summarystage:
early late NAs
  27
      67
             46
tumorstage:
           3
  1
       2
             4 NAs
 18
       9 52 15 46
```

```
substage:
            c NAs
  а
      b
 14
      10
           55 61
grade:
       2
            3 NAs
  1
  6
      27
           63
               44
batch:
2006-06-01 2006-06-27 2006-06-28 2006-06-29 2006-06-30 2006-07-20 2008-03-06
       21
                  18
                            37 20
                                                 36
                                                            7
                                                                        1
uncurated_author_metadata:
            Class
  Length
                       Mode
     140 character character
```

GSE2109\_eset IGC Expression Project for Oncology

#### Description

EXpression Project for Oncology, International Genomics Consortium, www.intgen.org

#### Usage

data( GSE2109\_eset )

## Format

```
experimentData(eset):
Experiment data
 Experimenter name: EXpression Project for Oncology, International Genomics Consortium, www.intgen.org
 Laboratory: exp0, IGC 2005
  Contact information:
  Title: IGC EXpression Project for Oncology
 URL:
  PMIDs: PMID unknown
  Abstract: A 8 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
   platform_shorttitle:
      Affymetrix HG-U133Plus2
   platform_summary:
```

```
hgu133plus2
platform_manufacturer:
   Affymetrix
platform_distribution:
   commercial
platform_accession:
   GPL570
```

```
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1BG A1CF ... ZZZ3 (19093 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 19093 features, 204 samples
Platform type: hgu133plus2
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
            Class
                     Mode
     204 character character
sample_type:
tumor
 204
histological_type:
         Class
  Length
                     Mode
     204 character character
primarysite:
other
      ov NAs
  23
      178
             3
summarygrade:
high low NAs
 91 31 82
summarystage:
early late NAs
  37
       87
             80
tumorstage:
```

4 NAs substage: а b c NAs grade: 4 NAs age\_at\_initial\_pathologic\_diagnosis: Min. 1st Qu. Median Mean 3rd Qu. Max. 25.00 45.00 55.00 58.82 65.00 85.00 batch: 2004-12-03 2004-12-04 2004-12-07 2005-02-11 2005-03-03 2005-03-10 2005-03-11 2005-03-15 2005-03-16 2005-03-17 2005-03-19 2005-03-22 2005-04-13 2005-04-26 2005-04-29 2005-05-10 2005-06-01 2005-06-03 2005-06-08 2005-06-17 2005-08-05 2005-08-09 2005-08-11 2005-09-07 2005-09-09 2005-09-13 2005-11-02 2005-11-04 2005-11-15 2005-11-18 2005-12-02 2006-01-24 2006-01-26 2006-02-07 2006-02-28 2006-03-06 2006-03-14 2006-04-18 2006-04-20 2006-05-16 2006-06-08 2006-07-26 2006-07-28 2006-09-12 2006-09-14 2006-10-10 2006-10-24 2006-10-31 2006-11-09 2006-11-21 2006-11-30 2006-12-07 2007-01-12 2007-02-09 2007-03-07 2007-03-09 2007-03-15 2007-05-01 2007-05-03 2007-05-15 2007-05-18 2007-05-30 2007-06-12 2007-07-27 2007-09-05 2007-09-07 2007-09-11 2007-09-12 2008-02-15 2008-02-21 2008-02-27 2008-03-04 2008-05-13 2008-05-16 2008-05-23 uncurated\_author\_metadata: Length Class Mode 204 character character

```
GSE26712_eset
```

A gene signature predicting for survival in suboptimally debulked patients with ovarian cancer.

#### Description

Despite the existence of morphologically indistinguishable disease, patients with advanced ovarian tumors display a broad range of survival end points. We hypothesize that gene expression profiling can identify a prognostic signature accounting for these distinct clinical outcomes. To resolve survival-associated loci, gene expression profiling was completed for an extensive set of 185 (90 optimal/95 suboptimal) primary ovarian tumors using the Affymetrix human U133A microarray. Cox regression analysis identified probe sets associated with survival in optimally and suboptimally debulked tumor sets at a P value of <0.01. Leave-one-out cross-validation was applied to each tumor cohort and confirmed by a permutation test. External validation was conducted by applying the gene signature to a publicly available array database of expression profiles of advanced stage suboptimally debulked tumors. The prognostic signature successfully classified the tumors according to survival for suboptimally (P = 0.0179) but not optimally debulked (P = 0.144) patients. The suboptimal gene signature was validated using the independent set of tumors (odds ratio, 8.75; P = 0.0146). To elucidate signaling events amenable to the apeutic intervention in suboptimally debulked patients, pathway analysis was completed for the top 57 survival-associated probe sets. For suboptimally debulked patients, confirmation of the predictive gene signature supports the existence of a clinically relevant predictor, as well as the possibility of novel therapeutic opportunities. Ultimately, the prognostic classifier defined for suboptimally debulked tumors may aid in the classification and enhancement of patient outcome for this high-risk population.

#### Usage

data( GSE26712\_eset )

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Bonome T, Levine DA, Shih J, Randonovich M, Pise-Masison CA, Bogomolniy F, Ozbun L,
 Laboratory: Bonome, Birrer 2008
 Contact information:
 Title: A gene signature predicting for survival in suboptimally debulked patients with ovarian cancer.
 URL:
 PMIDs: 18593951
 Abstract: A 238 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
      [HG-U133A] Affymetrix Human Genome U133A Array
  platform_shorttitle:
     Affymetrix HG-U133A
  platform_summary:
```

```
hgu133a
platform_manufacturer:
Affymetrix
platform_distribution:
```

commercial

```
platform_accession:
    GPL96
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1CF A2M ... ZZZ3 (12981 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 12981 features, 195 samples
Platform type: hgu133a
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
  10 observations deleted due to missingness
records n.max n.start events median 0.95LCL 0.95UCL
185.00 185.00 185.00 129.00
                                3.83
                                       3.24
                                              4.83
_____
Available sample meta-data:
_____
alt_sample_name:
                      Mode
  Length
            Class
     195 character character
sample_type:
healthy tumor
    10
          185
histological_type:
ser NAs
185 10
primarysite:
ov
195
summarygrade:
high NAs
185
     10
summarystage:
late NAs
185
     10
```

```
tumorstage:
  3 4 NAs
146 36 13
substage:
  b c NAs
  9 137 49
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median
                     Mean 3rd Qu.
                                    Max.
                                            NAs
 26.00 52.00 63.00 61.54 70.00
                                  84.00
                                           13
recurrence_status:
norecurrence recurrence
        42
                 153
days_to_death:
  Min. 1st Qu. Median Mean 3rd Qu.
                                            NAs
                                     Max.
  21.9 660.6 1164.0 1429.0 1880.0 4982.0
                                          10
vital_status:
deceased living
                NAs
    129
          56
                  10
debulking:
  optimal suboptimal
                        NAs
      90
           95
                        10
percent_normal_cells:
20-
195
percent_stromal_cells:
20-
195
percent_tumor_cells:
80+
195
batch:
2003-11-04 2003-11-05 2003-11-06 2003-11-07 2003-11-20 2003-11-21 2003-12-16
                         9
      14 16
                                  6 10 15 17
2003-12-23 2003-12-24 2004-04-20 2004-04-21 2004-04-27 2004-09-28 2005-07-27
                11
                          20
                               17
                                             9
                                                                15
      12
                                                     14
2006-11-09
      10
```

uncurated\_author\_metadata: Length Class Mode 195 character character

GSE30009\_eset

Multidrug resistance-linked gene signature predicts overall survival of patients with primary ovarian serous carcinoma.

#### Description

This study assesses the ability of multidrug resistance (MDR)-associated gene expression patterns to predict survival in patients with newly diagnosed carcinoma of the ovary. The scope of this research differs substantially from that of previous reports, as a very large set of genes was evaluated whose expression has been shown to affect response to chemotherapy. We applied a customized TaqMan low density array, a highly sensitive and specific assay, to study the expression profiles of 380 MDR-linked genes in 80 tumor specimens collected at initial surgery to debulk primary serous carcinoma. The RNA expression profiles of these drug resistance genes were correlated with clinical outcomes.Leave-one-out cross-validation was used to estimate the ability of MDR gene expression to predict survival. Although gene expression alone does not predict overall survival (OS; P = 0.06), four covariates (age, stage, CA125 level, and surgical debulking) do (P = 0.03). When gene expression was added to the covariates, we found an 11-gene signature that provides a major improvement in OS prediction (log-rank statistic P < 0.003). The predictive power of this 11-gene signature was confirmed by dividing high- and low-risk patient groups, as defined by their clinical covariates, into four specific risk groups on the basis of expression levels. This study reveals an 11-gene signature that allows a more precise prognosis for patients with serous cancer of the ovary treated with carboplatin- and paclitaxel-based therapy. These 11 new targets offer opportunities for new therapies to improve clinical outcome in ovarian cancer.

#### Usage

data( GSE30009\_eset )

#### Format

```
experimentData(eset):
Experiment data
Experimenter name: Gillet JP, Calcagno AM, Varma S, Davidson B et al. Multidrug resistance-linked gene
Laboratory: Gillet, Gottesman 2012
```

Contact information:

Title: Multidrug resistance-linked gene signature predicts overall survival of patients with primary c URL:

PMIDs: 22492981

Abstract: A 244 word abstract is available. Use abstract method. Information is available on: preprocessing

```
notes:
   platform_title:
      TaqMan qRT-PCR Homo sapiens Low-Density Array 380
   platform_shorttitle:
      TaqMan qRT-PCR
   platform_summary:
      NA
   platform_manufacturer:
      TagMan
   platform_distribution:
      custom
   platform_accession:
      GPL13728
Preprocessing: default
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: ABCA1 ABCA10 ... XRCC6 (359 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 359 features, 103 samples
Platform type: NA
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
records
       n.max n.start events median 0.95LCL 0.95UCL
103.00 103.00 103.00 57.00 3.42 2.92
                                             5.34
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
         Class
                      Mode
     103 character character
sample_type:
tumor
 103
histological_type:
clearcell
             ser
       1
              102
summarygrade:
```

```
high low NAs
 92
        9
             2
summarystage:
late
103
tumorstage:
3 4
82 21
substage:
  b
       c NAs
   2
       60
            41
grade:
  1
        2
             3 NAs
        5
   4
            92
                  2
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median
                           Mean 3rd Qu.
                                            Max.
  30.00 56.00
                61.00
                          62.45
                                 71.50
                                           87.00
days_to_death:
  Min. 1st Qu. Median
                           Mean 3rd Qu.
                                            Max.
     24
            598
                   1053
                           1156
                                   1568
                                            4748
vital_status:
deceased
           living
      57
               46
debulking:
  optimal suboptimal
        81
                   22
uncurated_author_metadata:
  Length
             Class
                         Mode
      103 character character
```

```
GSE30161_eset
```

Multi-gene expression predictors of single drug responses to adjuvant chemotherapy in ovarian carcinoma: predicting platinum resistance.

## Description

Despite advances in radical surgery and chemotherapy delivery, ovarian cancer is the most lethal gynecologic malignancy. Standard therapy includes treatment with platinum-based combination

chemotherapies yet there is no biomarker model to predict their responses to these agents. We here have developed and independently tested our multi-gene molecular predictors for forecasting patients' responses to individual drugs on a cohort of 55 ovarian cancer patients. To independently validate these molecular predictors, we performed microarray profiling on FFPE tumor samples of 55 ovarian cancer patients (UVA-55) treated with platinum-based adjuvant chemotherapy. Genomewide chemosensitivity biomarkers were initially discovered from the in vitro drug activities and genomic expression data for carboplatin and paclitaxel, respectively. Multivariate predictors were trained with the cell line data and then evaluated with a historical patient cohort. For the UVA-55 cohort, the carboplatin, taxol, and combination predictors significantly stratified responder patients and non-responder patients (p = 0.019, 0.04, 0.014) with sensitivity = 91%, 96%, 93 and NPV = 57%, 67%, 67% in pathologic clinical response. The combination predictor also demonstrated a significant survival difference between predicted responders and non-responders with a median survival of 55.4 months vs. 32.1 months. Thus, COXEN single- and combination-drug predictors successfully stratified platinum resistance and taxane response in an independent cohort of ovarian cancer patients based on their FFPE tumor samples.

#### Usage

data( GSE30161\_eset )

## Format

```
experimentData(eset):
Experiment data
 Experimenter name: Ferriss JS, Kim Y, Duska L, Birrer M, Levine DA, Moskaluk C, Theodorescu D, Lee JK
 Laboratory: Ferriss, Lee 2012
 Contact information:
 Title: Multi-gene expression predictors of single drug responses to adjuvant chemotherapy in ovarian c
 URL:
 PMIDs: 22348014
 Abstract: A 215 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
  platform_shorttitle:
     Affymetrix HG-U133Plus2
  platform_summary:
     hgu133plus2
  platform_manufacturer:
     Affymetrix
  platform_distribution:
      commercial
  platform_accession:
     GPL570
Preprocessing: frma
featureData(eset):
```

```
An object of class AnnotatedDataFrame
featureNames: A1BG A1CF ... ZZZ3 (19093 total)
varLabels: probeset gene
varMetadata: labelDescription
```

```
assayData: 19093 features, 58 samples
Platform type: hgu133plus2
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
records
        n.max n.start events median 0.95LCL 0.95UCL
 58.00 58.00 58.00 36.00 4.19 2.70 6.17
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
         Class
                      Mode
      58 character character
sample_type:
tumor
  58
histological_type:
  Length
         Class
                      Mode
      58 character character
summarygrade:
high low NAs
 33
     21
           4
summarystage:
late
 58
tumorstage:
3 4
53 5
substage:
a b c
9 11 38
grade:
```

```
1 2
        3 NAs
  2 19 33 4
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median Mean 3rd Qu.
                                  Max.
 38.00 53.50 62.00 62.57 72.00 85.00
pltx:
У
58
tax:
n y
4 54
neo:
n
58
days_to_tumor_recurrence:
  Min. 1st Qu. Median Mean 3rd Qu. Max.
  12.0 255.2 386.0 742.1 768.2 4208.0
recurrence_status:
norecurrence recurrence
                          NAs
        6 48
                           4
days_to_death:
  Min. 1st Qu. Median Mean 3rd Qu.
                                   Max.
  49.0 585.2 1010.0 1375.0 2131.0 4208.0
vital_status:
deceased living
    36
            22
debulking:
  optimal suboptimal NAs
      26 30
                      2
batch:
2009-10-07 2009-10-08 2009-10-09 2009-10-20
      28
         18 8 4
uncurated_author_metadata:
  Length Class Mode
     58 character character
```

GSE32062.GPL6480\_eset

High-risk ovarian cancer based on 126-gene expression signature is uniquely characterized by downregulation of antigen presentation pathway.

## Description

High-grade serous ovarian cancers are heterogeneous not only in terms of clinical outcome but also at the molecular level. Our aim was to establish a novel risk classification system based on a gene expression signature for predicting overall survival, leading to suggesting novel therapeutic strategies for high-risk patients. In this large-scale cross-platform study of six microarray data sets consisting of 1,054 ovarian cancer patients, we developed a gene expression signature for predicting overall survival by applying elastic net and 10-fold cross-validation to a Japanese data set A (n = 260) and evaluated the signature in five other data sets. Subsequently, we investigated differences in the biological characteristics between high- and low-risk ovarian cancer groups. An elastic net analysis identified a 126-gene expression signature for predicting overall survival in patients with ovarian cancer using the Japanese data set A (multivariate analysis, P = 4?? 10(-20)). We validated its predictive ability with five other data sets using multivariate analysis (Tothill's data set, P =1 ?? 10(-5); Bonome's data set, P = 0.0033; Dressman's data set, P = 0.0016; TCGA data set, P = 0.0027; Japanese data set B, P = 0.021). Through gene ontology and pathway analyses, we identified a significant reduction in expression of immune-response-related genes, especially on the antigen presentation pathway, in high-risk ovarian cancer patients. This risk classification based on the 126-gene expression signature is an accurate predictor of clinical outcome in patients with advanced stage high-grade serous ovarian cancer and has the potential to develop new therapeutic strategies for high-grade serous ovarian cancer patients.

## Usage

data( GSE32062.GPL6480\_eset )

#### Format

```
experimentData(eset):
Experiment data
Experimenter name: Yoshihara K, Tsunoda T, Shigemizu D, Fujiwara H et al. High-risk ovarian cancer base
Laboratory: Yoshihara, Tanaka 2012
Contact information:
Title: High-risk ovarian cancer based on 126-gene expression signature is uniquely characterized by do
URL:
PMIDs: 22241791
Abstract: A 255 word abstract is available. Use abstract method.
Information is available on: preprocessing
notes:
    platform_title:
        Agilent-014850 Whole Human Genome Microarray 4x44K G4112F (Probe Name vers
ion)
```

```
platform_shorttitle:
    Agilent G4112F
platform_summary:
    hgug4112a
platform_manufacturer:
    Agilent
platform_distribution:
    commercial
platform_accession:
    GPL6480
```

```
Preprocessing: default
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1CF A2LD1 ... ZZZ3 (19358 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 19358 features, 260 samples
Platform type: hgug4112a
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
        n.max n.start events median 0.95LCL 0.95UCL
records
260.00 260.00 260.00 121.00
                               4.93
                                      4.11
                                            6.58
 ------
Available sample meta-data:
_____
alt_sample_name:
  Length
           Class
                      Mode
     260 character character
sample_type:
tumor
  260
histological_type:
ser
260
summarygrade:
high low
129 131
```

```
summarystage:
late
260
tumorstage:
  3
    4
204 56
substage:
   а
        b
             c NAs
   4
       20
          180
                 56
grade:
  2 3
131 129
pltx:
 у
260
tax:
 у
260
days_to_death:
  Min. 1st Qu. Median
                           Mean 3rd Qu.
                                            Max.
     30
            810
                   1245
                           1344
                                   1710
                                            3840
vital_status:
deceased
           living
     121
              139
debulking:
   optimal suboptimal
       103
                  157
uncurated_author_metadata:
   Length
              Class
                         Mode
      260 character character
```

GSE32063\_eset

High-risk ovarian cancer based on 126-gene expression signature is uniquely characterized by downregulation of antigen presentation pathway.

#### Description

High-grade serous ovarian cancers are heterogeneous not only in terms of clinical outcome but also at the molecular level. Our aim was to establish a novel risk classification system based on a gene expression signature for predicting overall survival, leading to suggesting novel therapeutic strategies for high-risk patients. In this large-scale cross-platform study of six microarray data sets consisting of 1,054 ovarian cancer patients, we developed a gene expression signature for predicting overall survival by applying elastic net and 10-fold cross-validation to a Japanese data set A (n =260) and evaluated the signature in five other data sets. Subsequently, we investigated differences in the biological characteristics between high- and low-risk ovarian cancer groups. An elastic net analysis identified a 126-gene expression signature for predicting overall survival in patients with ovarian cancer using the Japanese data set A (multivariate analysis, P = 4?? 10(-20)). We validated its predictive ability with five other data sets using multivariate analysis (Tothill's data set, P =1 ?? 10(-5); Bonome's data set, P = 0.0033; Dressman's data set, P = 0.0016; TCGA data set, P = 0.0027; Japanese data set B, P = 0.021). Through gene ontology and pathway analyses, we identified a significant reduction in expression of immune-response-related genes, especially on the antigen presentation pathway, in high-risk ovarian cancer patients. This risk classification based on the 126-gene expression signature is an accurate predictor of clinical outcome in patients with advanced stage high-grade serous ovarian cancer and has the potential to develop new therapeutic strategies for high-grade serous ovarian cancer patients.

#### Usage

data( GSE32063\_eset )

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Yoshihara K, Tsunoda T, Shigemizu D, Fujiwara H et al. High-risk ovarian cancer base
 Laboratory: Yoshihara, Tanaka 2012
 Contact information:
 Title: High-risk ovarian cancer based on 126-gene expression signature is uniquely characterized by do
 URL:
 PMIDs: 22241791
 Abstract: A 255 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
     Agilent-014850 Whole Human Genome Microarray 4x44K G4112F (Probe Name vers
ion)
  platform_shorttitle:
     Agilent G4112F
  platform_summary:
     hgug4112a
  platform_manufacturer:
     Agilent
  platform_distribution:
```

```
commercial
platform_accession:
    GPL6480
Preprocessing: default
featureData(eset):
An object of class AnnotatedDataFrame
   featureNames: A1CF A2LD1 ... ZZZ3 (19358 total)
   varLabels: probeset gene
   varMetadata: labelDescription
```

```
assayData: 19358 features, 40 samples
Platform type: hgug4112a
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
records n.max n.start events median 0.95LCL 0.95UCL
 40.00 40.00 40.00 22.00
                               4.44
                                      3.29
                                                NA
_____
Available sample meta-data:
_____
alt_sample_name:
                      Mode
  Length
            Class
      40 character character
sample_type:
tumor
  40
histological_type:
ser
40
summarygrade:
high low
 17 23
summarystage:
late
 40
tumorstage:
3 4
31 9
```

```
substage:
   b
        c NAs
   3
       28
              9
grade:
2 3
23 17
pltx:
у
40
tax:
у
40
days_to_death:
                             Mean 3rd Qu.
   Min. 1st Qu.
                  Median
                                              Max.
    210
            705
                    1155
                             1346
                                     1792
                                              3330
vital_status:
deceased
           living
      22
                18
debulking:
   optimal suboptimal
        19
                    21
uncurated_author_metadata:
   Length
               Class
                          Mode
       40 character character
```

GSE6008\_eset

Lysophosphatidic acid-induced transcriptional profile represents serous epithelial ovarian carcinoma and worsened prognosis.

## Description

Lysophosphatidic acid (LPA) governs a number of physiologic and pathophysiological processes. Malignant ascites fluid is rich in LPA, and LPA receptors are aberrantly expressed by ovarian cancer cells, implicating LPA in the initiation and progression of ovarian cancer. However, there is an absence of systematic data critically analyzing the transcriptional changes induced by LPA in ovarian cancer. In this study, gene expression profiling was used to examine LPA-mediated transcription by exogenously adding LPA to human epithelial ovarian cancer cells for 24 h to mimic long-term stimulation in the tumor microenvironment. The resultant transcriptional profile comprised a 39-gene

signature that closely correlated to serous epithelial ovarian carcinoma. Hierarchical clustering of ovarian cancer patient specimens demonstrated that the signature is associated with worsened prognosis. Patients with LPA-signature-positive ovarian tumors have reduced disease-specific and progression-free survival times. They have a higher frequency of stage IIIc serous carcinoma and a greater proportion is deceased. Among the 39-gene signature, a group of seven genes associated with cell adhesion recapitulated the results. Out of those seven, claudin-1, an adhesion molecule and phenotypic epithelial marker, is the only independent biomarker of serous epithelial ovarian carcinoma. Knockdown of claudin-1 expression in ovarian cancer cells reduces LPA-mediated cellular adhesion, enhances suspended cells and reduces LPA-mediated migration. The data suggest that transcriptional events mediated by LPA in the tumor microenvironment influence tumor progression through modulation of cell adhesion molecules like claudin-1 and, for the first time, report an LPA-mediated expression signature in ovarian cancer that predicts a worse prognosis.

#### Usage

data( GSE6008\_eset )

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Murph MM, Liu W, Yu S, Lu Y, Hall H, Hennessy BT, Lahad J, Schaner M, Helland A, Kris
 Laboratory: Murph, Mills 2009
 Contact information:
 Title: Lysophosphatidic acid-induced transcriptional profile represents serous epithelial ovarian car
 URL:
 PMIDs: 19440550
  Abstract: A 247 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133A] Affymetrix Human Genome U133A Array
   platform_shorttitle:
      Affymetrix HG-U133A
   platform_summary:
      hgu133a
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL96
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1CF A2M ... ZZZ3 (12981 total)
  varLabels: probeset gene
```

# GSE6008\_eset

varMetadata: labelDescription

## Details

```
assayData: 12981 features, 103 samples
Platform type: hgu133a
-----
Available sample meta-data:
_____
alt_sample_name:
  Length Class Mode
    103 character character
sample_type:
healthy tumor
    4
          99
histological_type:
clearcell endo mucinous ser NAs
8 37 13 41 4
primarysite:
ov
103
summarygrade:
high low NAs
 38 36 29
summarystage:
early late NAs
  42 53 8
tumorstage:
 1 2 3 4 NAs
 35 11 44 9 4
substage:
        c d NAs
 a b
 19
    2 54 1 27
grade:
 1 2 3 NAs
 19 17 38 29
batch:
2002-04-03 2002-04-04 2002-04-09 2002-04-10 2002-04-12 2002-08-13 2002-08-15
```

```
2
         3
                     8
                                9
                                                        3
                                                                   4
                                                                               4
2002-08-22 2002-08-23 2002-08-27 2002-08-28 2002-08-29 2002-08-30 2002-09-11
         8
                     8
                                5
                                            6
                                                       16
                                                                  14
                                                                               9
2006-01-27
         4
uncurated_author_metadata:
              Class
   Length
                          Mode
      103 character character
```

GSE6822\_eset

Classification of ovarian tumor samples

## Description

Ouellet V, Provencher DM, Maugard CM, Le Page C, Ren F, Lussier C, Novak J, Ge B, Hudson TJ, Tonin PN, Mes-Masson A-M: Discrimination between serous low malignant potential and invasive epithelial ovarian tumors using molecular profiling. Oncogene 2005, 24:4672-4687.

## Usage

data( GSE6822\_eset )

## Format

```
experimentData(eset):
Experiment data
 Experimenter name: Ouellet V, Provencher DM, Maugard CM, Le Page C, Ren F, Lussier C, Novak J, Ge B, Hud
 Laboratory: Ouellet, Mes-Masson 2005
  Contact information:
  Title: Classification of ovarian tumor samples
  URL:
  PMIDs: PMID unknown
  Abstract: A 40 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [Hu6800] Affymetrix Human Full Length HuGeneFL Array
   platform_shorttitle:
      Affymetrix Hu6800
   platform_summary:
      hu6800
   platform_manufacturer:
      Affymetrix
   platform_distribution:
```

## GSE6822\_eset

```
commercial
platform_accession:
    GPL80
Preprocessing: rma
featureData(eset):
An object of class AnnotatedDataFrame
    featureNames: A2M AADAC ... ZYX (5231 total)
    varLabels: probeset gene
    varMetadata: labelDescription
```

## Details

```
assayData: 5231 features, 66 samples
Platform type: hu6800
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
        Class
                   Mode
     66 character character
sample_type:
tumor
  66
histological_type:
  Length Class
                   Mode
     66 character character
primarysite:
ov
66
summarygrade:
high low NAs
 40 15 11
grade:
  1 2 3 NAs
  1 14 40 11
batch:
2000-12-21 2001-05-03 2001-05-29 2001-06-12 2001-09-25 2001-09-26 2001-09-27
     1 1 3 3 1 5
                                                     8
2002-02-14 2002-04-17 2002-04-18 2002-07-18 2002-07-24 2002-10-20 2002-10-30
      4
         1 9
                          7
                                   4
                                             10
                                                         5
```

```
2002-11-01 2002-11-13
2 2
uncurated_author_metadata:
Length Class Mode
66 character character
```

GSE9891\_eset

Novel molecular subtypes of serous and endometrioid ovarian cancer linked to clinical outcome.

## Description

The study aim to identify novel molecular subtypes of ovarian cancer by gene expression profiling with linkage to clinical and pathologic features. Microarray gene expression profiling was done on 285 serous and endometrioid tumors of the ovary, peritoneum, and fallopian tube. K-means clustering was applied to identify robust molecular subtypes. Statistical analysis identified differentially expressed genes, pathways, and gene ontologies. Laser capture microdissection, pathology review, and immunohistochemistry validated the array-based findings. Patient survival within kmeans groups was evaluated using Cox proportional hazards models. Class prediction validated k-means groups in an independent dataset. A semisupervised survival analysis of the array data was used to compare against unsupervised clustering results. Optimal clustering of array data identified six molecular subtypes. Two subtypes represented predominantly serous low malignant potential and low-grade endometrioid subtypes, respectively. The remaining four subtypes represented higher grade and advanced stage cancers of serous and endometrioid morphology. A novel subtype of high-grade serous cancers reflected a mesenchymal cell type, characterized by overexpression of N-cadherin and P-cadherin and low expression of differentiation markers, including CA125 and MUC1. A poor prognosis subtype was defined by a reactive stroma gene expression signature, correlating with extensive desmoplasia in such samples. A similar poor prognosis signature could be found using a semisupervised analysis. Each subtype displayed distinct levels and patterns of immune cell infiltration. Class prediction identified similar subtypes in an independent ovarian dataset with similar prognostic trends.Gene expression profiling identified molecular subtypes of ovarian cancer of biological and clinical importance.

#### Usage

data( GSE9891\_eset )

#### Format

experimentData(eset): Experiment data Experimenter name: Tothill RW, Tinker AV, George J, Brown R, Fox SB, Lade S, Johnson DS, Trivett MK, Ete Laboratory: Tothill, Bowtell 2008 Contact information: Title Newlandscher Ausgebruchen auf generation of the second seco

Title: Novel molecular subtypes of serous and endometrioid ovarian cancer linked to clinical outcome.

```
URL:
  PMIDs: 18698038
  Abstract: A 243 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
   platform_shorttitle:
      Affymetrix HG-U133Plus2
   platform_summary:
      hgu133plus2
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL570
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1BG A1CF ... ZZZ3 (19093 total)
  varLabels: probeset gene
```

```
assayData: 19093 features, 285 samples
Platform type: hgu133plus2
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
  7 observations deleted due to missingness
records n.max n.start events median 0.95LCL 0.95UCL
278.00 278.00 278.00 113.00
                               3.95
                                      3.53
                                              5.01
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
           Class
                      Mode
     285 character character
sample_type:
tumor
  285
```

varMetadata: labelDescription

```
histological_type:
endo other ser
  20 1
           264
primarysite:
  ft other
          ov
  8 34
         243
arrayedsite:
  ft other
           ٥v
  2 83
           200
summarygrade:
high low NAs
163 116 6
summarystage:
early late NAs
 42 240 3
tumorstage:
 1 2 3 4 NAs
 24 18 218 22 3
substage:
 a b cNAs
 26
    19 212 28
grade:
 1 2 3 NAs
 19 97 163 6
age_at_initial_pathologic_diagnosis:
                                 Max. NAs
80.00 3
  Min. 1st Qu. Median Mean 3rd Qu.
 22.00 53.00 59.00 59.62 68.00
pltx:
 n yNAs
 39 243 3
tax:
n yNAs
 87 195 3
neo:
n yNAs
264 18 3
```

days\_to\_tumor\_recurrence: Min. 1st Qu. Median Mean 3rd Qu. NAs Max. 0.0 300.0 450.0 618.9 810.0 4980.0 recurrence\_status: norecurrence recurrence NAs days\_to\_death: Min. 1st Qu. Median Mean 3rd Qu. Max. NAs 0.0 547.5 855.0 955.1 1252.0 6420.0 vital\_status: deceased living NAs debulking: optimal suboptimal NAs batch: 2004-12-03 2004-12-23 2005-01-12 2005-01-17 2005-01-24 2005-01-31 2005-02-21 2005-03-17 2005-05-05 2005-05-09 2005-05-25 2005-05-27 2005-05-30 2005-06-02 2005-06-06 2005-06-08 2005-06-16 2005-06-17 2005-06-24 2005-07-06 2005-07-15 2005-07-20 2005-07-29 2005-08-03 2005-08-05 2005-08-18 2005-08-24 2005-08-26 2005-09-09 2005-09-14 2005-09-16 2005-09-21 2005-10-05 2005-10-26 2005-10-28 2005-11-04 2005-11-09 2005-11-11 2005-11-23 2005-12-15 2005-12-21 2006-01-20 2006-01-31 2006-02-08 2006-02-28 2006-04-05 2006-04-06 2006-04-12 2006-04-13 2006-04-28 2006-05-03 2006-06-06 2006-06-07 2006-06-22 2006-07-07 2006-07-19 uncurated\_author\_metadata: Mode Length Class 285 character character

PMID15897565\_eset

Patterns of gene expression that characterize long-term survival in advanced stage serous ovarian cancers.

## Description

A better understanding of the underlying biology of invasive serous ovarian cancer is critical for the development of early detection strategies and new therapeutics. The objective of this study was to define gene expression patterns associated with favorable survival.RNA from 65 serous ovarian cancers was analyzed using Affymetrix U133A microarrays. This included 54 stage III/IV cases (30 short-term survivors who lived <3 years and 24 long-term survivors who lived >7 years) and 11 stage I/II cases. Genes were screened on the basis of their level of and variability in expression, leaving 7,821 for use in developing a predictive model for survival. A composite predictive model was developed that combines Bayesian classification tree and multivariate discriminant models. Leave-one-out cross-validation was used to select and evaluate models.Patterns of genes were identified that distinguish short-term and long-term ovarian cancer survivors. The expression model developed for advanced stage disease classified all 11 early-stage ovarian cancers as long-term survivors. The MAL gene, which has been shown to confer resistance to cancer therapy, was most highly overexpressed in short-term survivors (3-fold compared with long-term survivors, and 29fold compared with early-stage cases). These results suggest that gene expression patterns underlie differences in outcome, and an examination of the genes that provide this discrimination reveals that many are implicated in processes that define the malignant phenotype.Differences in survival of advanced ovarian cancers are reflected by distinct patterns of gene expression. This biological distinction is further emphasized by the finding that early-stage cancers share expression patterns with the advanced stage long-term survivors, suggesting a shared favorable biology.

## Usage

data( PMID15897565\_eset )

# Format

experimentData(eset):											
Experiment data											
<pre>Experimenter name: Berchuck A, Iversen ES, Lancaster JM, Pittman J, Luo J, Lee P, Murphy S, Dres Laboratory: Berchuck, Marks 2005 Contact information: Title: Patterns of gene expression that characterize long-term survival in advanced stage serou URL:</pre>											
Abstract: A 258 word abstract is available. Use abstract method.											
Information is available on: preprocessing											
notes:											
platform_title:											
[HG-U133A] Affymetrix Human Genome U133A Array											
platform_shorttitle:											
Affymetrix HG-U133A											
platform_summary:											
hgu133a											
platform_manufacturer:											
Affymetrix											
platform_distribution:											

# PMID15897565\_eset

```
commercial
platform_accession:
    GPL96
warnings:
    These samples are a subset of PMID17290060.
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1CF A2M ... ZZZ3 (12981 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

# Details

```
assayData: 12981 features, 63 samples
Platform type: hgu133a
Binary overall survival summary (definitions of long and short provided by study authors):
```

```
long short NAs
  24 28 11
_____
Available sample meta-data:
-----
alt_sample_name:
  Min. 1st Qu. Median
                    Mean 3rd Qu.
                                   Max.
  1761 1828 1907
                     2001
                            2032
                                   2536
sample_type:
tumor
  63
histological_type:
ser
63
primarysite:
ov
63
summarygrade:
high low NAs
 25
    37 1
summarystage:
early late
```

```
11
         52
tumorstage:
1 2 3 4
7
  4 4 8 4
grade:
        2
             3
                  4 NAs
   1
   2
       35
            24
                  1
                       1
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median
                           Mean 3rd Qu.
                                            Max.
  33.00
        52.50
                  59.00
                          59.21 67.00
                                           79.00
os_binary:
long short
             NAs
   24
         28
               11
debulking:
   optimal suboptimal
                             NAs
        24
                   28
                               11
batch:
2002-09-20 2002-10-23 2002-11-12 2002-12-16 2002-12-21 2003-01-03 2003-05-30
                    9
                               10
                                           1
                                                       3
                                                                            13
        15
                                                                 11
2003-07-02
         1
uncurated_author_metadata:
              Class
  Length
                         Mode
       63 character character
```

PMID17290060\_eset

An integrated genomic-based approach to individualized treatment of patients with advanced-stage ovarian cancer.

# Description

The purpose of this study was to develop an integrated genomic-based approach to personalized treatment of patients with advanced-stage ovarian cancer. We have used gene expression profiles to identify patients likely to be resistant to primary platinum-based chemotherapy and also to identify alternate targeted therapeutic options for patients with de novo platinum-resistant disease. A gene expression model that predicts response to platinum-based therapy was developed using a training set of 83 advanced-stage serous ovarian cancers and tested on a 36-sample external validation set. In parallel, expression signatures that define the status of oncogenic signaling pathways were evaluated in 119 primary ovarian cancers and 12 ovarian cancer cell lines. In an effort to increase

chemotherapy sensitivity, pathways shown to be activated in platinum-resistant cancers were subject to targeted therapy in ovarian cancer cell lines.Gene expression profiles identified patients with ovarian cancer likely to be resistant to primary platinum-based chemotherapy with greater than 80% accuracy. In patients with platinum-resistant disease, we identified expression signatures consistent with activation of Src and Rb/E2F pathways, components of which were successfully targeted to increase response in ovarian cancer cell lines.We have defined a strategy for treatment of patients with advanced-stage ovarian cancer that uses therapeutic stratification based on predictions of response to chemotherapy, coupled with prediction of oncogenic pathway deregulation, as a method to direct the use of targeted agents.

#### Usage

```
data( PMID17290060_eset )
```

## Format

```
experimentData(eset):
Experiment data
Experimenter name: Dressman HK, Berchuck A, Chan G, Zhai J, Bild A, Sayer R, Cragun J, Clarke J, Whitake
Laboratory: Dressman, Lancaster 2007
Contact information:
Title: An integrated genomic-based approach to individualized treatment of patients with advanced-stag
URL:
PMIDs: 17290060
```

```
Abstract: A 223 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133A] Affymetrix Human Genome U133A Array
   platform_shorttitle:
      Affymetrix HG-U133A
   platform_summary:
      hgu133a
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL96
   warnings:
      This paper has been retracted.
Preprocessing: frma
```

```
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1CF A2M ... ZZZ3 (12981 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 12981 features, 117 samples
Platform type: hgu133a
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
records n.max n.start events median 0.95LCL 0.95UCL
117.00 117.00 117.00 67.00 5.26 2.79 7.48
_____
Available sample meta-data:
_____
alt_sample_name:
  Length Class
                    Mode
    117 character character
sample_type:
tumor
 117
histological_type:
ser
117
primarysite:
ov
117
summarygrade:
high low NAs
 57 57 3
summarystage:
early late NAs
  1 115 1
tumorstage:
  2 3 4 NAs
  1 98 17 1
grade:
  1
      2
         3 4 NAs
  4 53 56 1 3
days_to_death:
```

```
Min. 1st Qu.
                 Median
                            Mean 3rd Qu.
                                             Max.
     30
            510
                    1020
                            1496
                                     2220
                                             5550
vital_status:
deceased
           living
      67
                50
primary_therapy_outcome_success:
  completeresponse progressivedisease
                 85
                                     32
debulking:
   optimal suboptimal
                    54
        63
batch:
2002-09-20 2002-10-23 2002-11-12 2002-12-16 2002-12-21 2003-01-03 2003-05-30
        10
                     8
                                9
                                            1
                                                        3
                                                                   11
                                                                              10
2004-03-09 2004-03-16 2004-04-20 2004-05-18 2004-05-21 2004-05-27 2004-06-22
        16
                     6
                                5
                                           15
                                                        7
                                                                    7
                                                                               1
2004-06-23
         8
uncurated_author_metadata:
                          Mode
   Length
              Class
      117 character character
```

PMID19318476\_eset Microarray analysis of early stage serous ovarian cancers shows profiles predictive of favorable outcome.

#### Description

Although few women with advanced serous ovarian cancer are cured, detection of the disease at an early stage is associated with a much higher likelihood of survival. We previously used gene expression array analysis to distinguish subsets of advanced cancers based on disease outcome. In the present study, we report on gene expression of early-stage cancers and validate our prognostic model for advanced-stage cancers. Frozen specimens from 39 stage I/II, 42 stage III/IV, and 20 low malignant potential cancers were obtained from four different sites. A linear discriminant model was used to predict survival based upon array data. We validated the late-stage survival model and show that three of the most differentially expressed genes continue to be predictive of outcome. Most early-stage cancers (38 of 39 invasive, 15 of 20 low malignant potential) were classified as long-term survivors (median probabilities 0.97 and 0.86). MAL, the most differentially expressed gene, was further validated at the protein level and found to be an independent predictor of poor survival in an unselected group of advanced serous cancers (P = 0.0004). These data suggest that serous ovarian cancers detected at an early stage generally have a favorable underlying biology

#### PMID19318476\_eset

similar to advanced-stage cases that are long-term survivors. Conversely, most late-stage ovarian cancers seem to have a more virulent biology. This insight suggests that if screening approaches are to succeed it will be necessary to develop approaches that are able to detect these virulent cancers at an early stage.

### Usage

data( PMID19318476\_eset )

## Format

```
experimentData(eset):
Experiment data
 Experimenter name: Berchuck A, Iversen ES, Luo J, Clarke JP, Horne H, Levine DA, Boyd J, Alonso MA, Seco
 Laboratory: Berchuck, Lancaster 2009
 Contact information:
 Title: Microarray analysis of early stage serous ovarian cancers shows profiles predictive of favorabl
 URL:
 PMIDs: 19318476
 Abstract: A 241 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
      [HG-U133A] Affymetrix Human Genome U133A Array
   platform_shorttitle:
      Affymetrix HG-U133A
   platform_summary:
      hgu133a
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL96
   warnings:
      These samples are a subset of PMID17290060.
Preprocessing: frma
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: A1CF A2M ... ZZZ3 (12981 total)
  varLabels: probeset gene
```

# Details

assayData: 12981 features, 42 samples

varMetadata: labelDescription

```
Platform type: hgu133a
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
records
        n.max n.start events median 0.95LCL 0.95UCL
 42.00 42.00 42.00 22.00 2.79 2.30
                                            NA
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
         Class
                    Mode
     42 character character
sample_type:
tumor
  42
histological_type:
ser
42
summarygrade:
high low NAs
 24
    17 1
summarystage:
early late NAs
   2 39 1
tumorstage:
  1
      2 3 4 NAs
    1 29 10 1
  1
substage:
  а
    b
         c NAs
     1 29 11
  1
grade:
     2
  1
         3 NAs
  2
    15
         24 1
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median Mean 3rd Qu.
                                   Max.
                                          NAs
 33.00 55.00 62.00 61.46 70.00 81.00
                                            1
```

recurrence\_status:

```
norecurrence
               recurrence
                        36
           6
days_to_death:
  Min. 1st Qu.
                 Median
                            Mean 3rd Qu.
                                             Max.
   30.0
          367.5
                  825.0 1105.0 1050.0
                                         3420.0
vital_status:
deceased
           living
      22
               20
debulking:
   optimal suboptimal
                             NAs
        20
                   21
                                1
batch:
2004-03-09 2004-03-16 2004-04-20 2004-05-18 2004-05-21 2004-05-27 2004-06-22
                    3
        14
                                4
                                           8
                                                       6
                                                                  5
                                                                              1
2004-06-23
         1
uncurated_author_metadata:
   Length
              Class
                         Mode
       42 character character
```

TCGA.mirna.8x15kv2\_eset

Integrated genomic analyses of ovarian carcinoma.

# Description

A catalogue of molecular aberrations that cause ovarian cancer is critical for developing and deploying therapies that will improve patients' lives. The Cancer Genome Atlas project has analysed messenger RNA expression, microRNA expression, promoter methylation and DNA copy number in 489 high-grade serous ovarian adenocarcinomas and the DNA sequences of exons from coding genes in 316 of these tumours. Here we report that high-grade serous ovarian cancer is characterized by TP53 mutations in almost all tumours (96%); low prevalence but statistically recurrent somatic mutations in nine further genes including NF1, BRCA1, BRCA2, RB1 and CDK12; 113 significant focal DNA copy number aberrations; and promoter methylation events involving 168 genes. Analyses delineated four ovarian cancer transcriptional subtypes, three microRNA subtypes, four promoter methylation subtypes and a transcriptional signature associated with survival duration, and shed new light on the impact that tumours with BRCA1/2 (BRCA1 or BRCA2) and CCNE1 aberrations have on survival. Pathway analyses suggested that homologous recombination is defective in about half of the tumours analysed, and that NOTCH and FOXM1 signalling are involved in serous ovarian cancer pathophysiology.

# Usage

data( TCGA.mirna.8x15kv2\_eset )

# Format

```
experimentData(eset):
Experiment data
 Experimenter name: Integrated genomic analyses of ovarian carcinoma. Nature 2011, 474:609-615.
  Laboratory: Cancer Genome Atlas Research Network 2011
  Contact information:
  Title: Integrated genomic analyses of ovarian carcinoma.
  URL:
  PMIDs: 21720365
  Abstract: A 179 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [miRNA-8x15k2] Agilent Human miRNA G4470B
   platform_shorttitle:
      Agilent miRNA-8x15k2 G4470B
   platform_summary:
      NA
   platform_manufacturer:
      Agilent
   platform_distribution:
      commercial
   platform_accession:
      NA
Preprocessing: default
featureData(eset):
An object of class AnnotatedDataFrame
  featureNames: ebv-miR-BART1-3p ebv-miR-BART1-5p ... kshv-miR-K12-9*
    (799 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

#### Details

```
assayData: 799 features, 554 samples
Platform type: NA
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
```

10 observations deleted due to missingness records n.max n.start events median 0.95LCL 0.95UCL

4.03

544.00 544.00 544.00 286.00 3.71 3.42 \_\_\_\_\_ Available sample meta-data: \_\_\_\_\_ alt\_sample\_name: Length Class Mode 554 character character unique\_patient\_ID: Length Class Mode 554 character character sample\_type: tumor 554 histological\_type: ser 554 primarysite: other ov 550 4 summarygrade: high low NAs 474 68 12 summarystage: early late NAs 39 511 4 tumorstage: 1 2 3 4 NAs 16 23 426 85 4 substage: b c NAs 31 434 89 grade: 1 2 3 4 NAs 6 62 473 1 12 age\_at\_initial\_pathologic\_diagnosis: Min. 1st Qu. Median Mean 3rd Qu. Max.

26.00 51.00 59.00 59.81 69.00 89.00 pltx: n y NAs 19 478 57 tax: y NAs n 43 454 57 neo: n NAs 497 57 days\_to\_tumor\_recurrence: Min. 1st Qu. Median Mean 3rd Qu. Max. NAs 8.0 235.2 436.0 618.7 797.0 5480.0 44 recurrence\_status: norecurrence recurrence 268 286 days\_to\_death: Min. 1st Qu. Median Mean 3rd Qu. Max. NAs 8.0 346.0 867.5 997.7 1446.0 5480.0 10 vital\_status: deceased living NAs 286 7 261 site\_of\_tumor\_first\_recurrence: locoregional locoregional\_plus\_metastatic 145 3 metastasis NAs 138 268 primary\_therapy\_outcome\_success: completeresponse partialresponse progressivedisease stabledisease 308 63 41 30 NAs 112 debulking: optimal suboptimal NAs 359 137 58 percent\_normal\_cells: Min. 1st Qu. Median Mean 3rd Qu. NAs Max.

```
0.000
          0.000
                   0.000
                            2.375
                                    0.000 55.000
                                                        10
percent_stromal_cells:
   Min. 1st Qu.
                 Median
                            Mean 3rd Qu.
                                              Max.
                                                      NAs
   0.00
           5.00
                   10.00
                            12.78
                                    19.00
                                             70.00
                                                        16
percent_tumor_cells:
                 Median
   Min. 1st Qu.
                            Mean 3rd Qu.
                                                      NAs
                                              Max.
   0.00
          75.00
                   85.00
                            80.72
                                    90.00
                                           100.00
                                                        13
uncurated_author_metadata:
   Length
              Class
                          Mode
      554 character character
```

TCGA\_eset

Integrated genomic analyses of ovarian carcinoma.

#### Description

A catalogue of molecular aberrations that cause ovarian cancer is critical for developing and deploying therapies that will improve patients' lives. The Cancer Genome Atlas project has analysed messenger RNA expression, microRNA expression, promoter methylation and DNA copy number in 489 high-grade serous ovarian adenocarcinomas and the DNA sequences of exons from coding genes in 316 of these tumours. Here we report that high-grade serous ovarian cancer is characterized by TP53 mutations in almost all tumours (96%); low prevalence but statistically recurrent somatic mutations in nine further genes including NF1, BRCA1, BRCA2, RB1 and CDK12; 113 significant focal DNA copy number aberrations; and promoter methylation events involving 168 genes. Analyses delineated four ovarian cancer transcriptional subtypes, three microRNA subtypes, four promoter methylation subtypes and a transcriptional signature associated with survival duration, and shed new light on the impact that tumours with BRCA1/2 (BRCA1 or BRCA2) and CCNE1 aberrations have on survival. Pathway analyses suggested that homologous recombination is defective in about half of the tumours analysed, and that NOTCH and FOXM1 signalling are involved in serous ovarian cancer pathophysiology.

## Usage

data( TCGA\_eset )

## Format

experimentData(eset): Experiment data Experimenter name: Integrated genomic analyses of ovarian carcinoma. Nature 2011, 474:609-615. Laboratory: Cancer Genome Atlas Research Network 2011 Contact information: Title: Integrated genomic analyses of ovarian carcinoma.

```
URL:
 PMIDs: 21720365
  Abstract: A 179 word abstract is available. Use abstract method.
  Information is available on: preprocessing
  notes:
  platform_title:
      [HT_HG-U133A] Affymetrix HT Human Genome U133A Array
   platform_shorttitle:
      Affymetrix HT_HG-U133A
   platform_summary:
      hthgu133a
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL3921
   warnings:
     The following samples are likely from specimens also used in GSE26712: TCG
A.13.0725, TCGA.13.0885, TCGA.13.0887, TCGA.13.0890, TCGA.13.0886, TCGA.13
.0714, TCGA.13.0727, TCGA.13.1817, TCGA.13.1499, TCGA.13.0883
Preprocessing: rma
featureData(eset):
An object of class AnnotatedDataFrame
```

```
An object of class AnnotatedDataFrame
featureNames: A1CF A2M ... ZZZ3 (12981 total)
varLabels: probeset gene
varMetadata: labelDescription
```

```
assayData: 12981 features, 578 samples
Platform type: hthgu133a
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
```

21 observations deleted due to missingness records n.max n.start events median 0.95LCL 0.95UCL 557.00 557.00 557.00 290.00 3.73 3.45 4.06

```
------
```

```
Available sample meta-data:
```

alt\_sample\_name: Length Class Mode 578 character character

```
unique_patient_ID:
           Class
  Length
                     Mode
     578 character character
sample_type:
adjacentnormal
                   tumor
           8
                      570
histological_type:
ser NAs
568 10
primarysite:
other ov NAs
   4 564 10
summarygrade:
high low NAs
480 75 23
summarystage:
early late NAs
  43 520
          15
tumorstage:
  1 2 3
             4 NAs
 16
     27 436
             84 15
substage:
  b c NAs
 31 448
         99
grade:
  1 2 3
               4 NAs
  6 69 479
             1 23
age_at_initial_pathologic_diagnosis:
                                         NAs
  Min. 1st Qu. Median
                     Mean 3rd Qu.
                                    Max.
 26.00 51.00 59.00
                     59.70 68.25
                                    89.00
                                         10
pltx:
  n
     y NAs
 19 492 67
tax:
    y NAs
  n
 43 468 67
```

neo: n NAs 511 67 days\_to\_tumor\_recurrence: Min. 1st Qu. Median Mean 3rd Qu. Max. NAs 238.2 623.7 812.0 5480.0 56 8.0 443.5 recurrence\_status: norecurrence recurrence 279 299 days\_to\_death: Min. 1st Qu. Median Mean 3rd Qu. Max. NAs 8 349 881 1010 1446 5480 21 vital\_status: deceased living NAs 290 270 18 site\_of\_tumor\_first\_recurrence: locoregional locoregional\_plus\_metastatic 153 3 metastasis NAs 143 279 primary\_therapy\_outcome\_success: completeresponse partialresponse progressivedisease stabledisease 318 30 65 41 NAs 124 debulking: optimal suboptimal NAs 367 140 71 percent\_normal\_cells: Min. 1st Qu. Median Mean 3rd Qu. Max. NAs 0.000 0.000 0.000 2.385 0.000 55.000 19 percent\_stromal\_cells: Min. 1st Ou. Median Mean 3rd Qu. NAs Max. 0.00 5.00 10.00 12.85 20.00 70.00 25 percent\_tumor\_cells: Min. 1st Qu. Median Mean 3rd Qu. NAs Max. 0.00 75.00 85.00 80.64 90.00 100.00 22

batch:														
9	11	12	13	14	15	17	18	19	21	22	24	27	40 NAs	
45	37	45	47	46	22	47	47	47	46	47	46	6	49	1

 ${\tt uncurated\_author\_metadata:}$ 

Length Class Mode 578 character character

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