# 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, logging, sva, xtable, futile.logger, BiocStyle

License Artistic-2.0

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

biocViews ExperimentData, RNASeqData, CancerData, OvarianCancerData, MicroarrayData

NeedsCompilation no

## **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

Package:curatedOvarianDataType:PackageVersion:1.14.0Date:2015-03-05License:Artistic-2.0Depends: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).

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
Laboratory: Bentink, Matulonis 2012
Contact information:
Title: Angiogenic mRNA and microRNA gene expression signature predicts a novel subtype of serous
URL:
PMIDs: 22348002
Abstract: A 212 word abstract is available. Use 'abstract' method.
Information is available on: preprocessing
notes:
platform_title:
```

4

```
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
   platform_technology:
      in situ oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
```

```
featureNames: A2M A4GALT ... ZZEF1 (10357 total)
varLabels: probeset gene
varMetadata: labelDescription
```

```
assayData: 10357 features, 129 samples
Platform type: illuminaHumanv2
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
129.00 129.00 129.00 73.00 3.51 2.68 4.13
_____
Available sample meta-data:
_____
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
        128
    1
tumorstage:
  2
      3
          4
  1 109
        19
substage:
        b
             c NA's
   а
   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
                             NA's
        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

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#### GSE12418\_eset

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 s
  Laboratory: Partheen, Horvath 2006
  Contact information:
 Title: Expression analysis of stage III serous ovarian adenocarcinoma distinguishes a sub-group of
  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
   platform_technology:
      spotted oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1CF A2M ... ZZZ3 (12681 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

#### Details

```
assayData: 12681 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
```

8

#### 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 )

```
experimentData(eset):
Experiment data
Experimenter name: Yoshihara K, Tajima A, Komata D, Yamamoto T, Kodama S, Fujiwara H, Suzuki M, On
Laboratory: Yoshihara, Tanaka 2009
Contact information:
Title: Gene expression profiling of advanced-stage serous ovarian cancers distinguishes novel sul
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

platform_technology:

in situ oligonucleotide

Preprocessing: default

featureData(eset):

An object of class 'AnnotatedDataFrame'

featureNames: AIBG_AICE____7773__(16889_tota
```

```
featureNames: A1BG A1CF ... ZZZ3 (16889 total)
varLabels: probeset gene
varMetadata: labelDescription
```

#### Details

```
assayData: 16889 features, 53 samples
Platform type: hgug4110b
-----
Available sample meta-data:
_____
alt_sample_name:
                   Mode
  Length Class
     53 character character
sample_type:
healthy tumor
    10
         43
histological_type:
ser NA's
 43
    10
primarysite:
ov
53
summarystage:
early late NA's
   8
      35
            10
tumorstage:
  1 NA's
  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, H 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
   platform_technology:
      spotted oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1CF ... ZZZ3 (20577 total)
  varLabels: probeset gene
 varMetadata: labelDescription
```

```
assayData: 20577 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 NA's
  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 NA's
  85
       59
            13
summarystage:
late
 157
grade:
   1
        2
             3
                   4 NA's
  14
       45
            82
                   3
                       13
age_at_initial_pathologic_diagnosis:
   Min. 1st Qu. Median
                            Mean 3rd Qu.
                                             Max.
  21.00
          50.00
                  60.00
                           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 )

```
experimentData(eset):
Experiment data
 Experimenter name: Denkert C, Budczies J, Darb-Esfahani S, Gy??rffy B et al. A prognostic gene ex
  Laboratory: Denkert, Lage 2009
  Contact information:
 Title: A prognostic gene expression index in ovarian cancer - validation across different indeper
  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
   platform_technology:
      in situ oligonucleotide
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1CF A2M ... ZZZ3 (13104 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

## GSE14764\_eset

```
assayData: 13104 features, 80 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
 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 NA's
  4
      6
          32 38
grade:
1 2 3
3 23 54
recurrence_status:
                             NA's
norecurrence recurrence
              26
        50
                                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:
   Length
              Class
                          Mode
       80 character character
uncurated_author_metadata:
                          Mode
   Length
              Class
       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 )

#### GSE17260\_eset

```
experimentData(eset):
   Experiment data
     Experimenter name: Yoshihara K, Tajima A, Yahata T, Kodama S, Fujiwara H, Suzuki M, Onishi Y, Hata
     Laboratory: Yoshihara, Tanaka 2010
     Contact information:
     Title: Gene expression profile for predicting survival in advanced-stage serous ovarian cancer ad
     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
      platform_technology:
         in situ oligonucleotide
   Preprocessing: default
   featureData(eset):
   An object of class 'AnnotatedDataFrame'
     featureNames: A1BG A1BG-AS1 ... ZZZ3 (20106 total)
     varLabels: probeset gene
     varMetadata: labelDescription
Details
```

```
assayData: 20106 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 cNA's
  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.
```

#### GSE18520\_eset

```
30
             660
                     915
                            1086
                                     1530
                                             2430
vital_status:
           living
deceased
      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 )
```

```
experimentData(eset):
Experiment data
Experimenter name: Mok SC, Bonome T, Vathipadiekal V, Bell A, Johnson ME, Wong KK, Park DC, Hao K,
Laboratory: Mok, Birrer 2009
Contact information:
Title: A gene signature predictive for outcome in advanced ovarian cancer identifies a survival f
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
   platform_distribution:
      commercial
   platform_accession:
      GPL570
   platform_technology:
      in situ oligonucleotide
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total)
  varLabels: probeset gene
```

varMetadata: labelDescription

```
assayData: 19816 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 NA's
 53 10
primarysite:
ov
63
summarygrade:
```

```
high NA's
 53
     10
summarystage:
late NA's
 53
     10
tumorstage:
  3 NA's
 53
     10
grade:
   3 NA's
 53 10
days_to_death:
  Min. 1st Qu. Median
                          Mean 3rd Qu.
                                         Max.
                                                 NA's
   150
           450
                          1212 1440
                                         4500
                                                  10
                630
vital_status:
deceased living
                     NA's
     41
                      10
             12
debulking:
  optimal suboptimal
                           NA's
       28
                  11
                             24
percent_normal_cells:
 0
63
percent_stromal_cells:
0
63
percent_tumor_cells:
100
 63
batch:
  Length
             Class
                        Mode
      63 character character
uncurated_author_metadata:
            Class
  Length
                       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 expressio
 Laboratory: Konstantinopoulos, Cannistra 2010 hgu133plus2
 Contact information:
 Title: Gene expression profile of BRCAness that correlates with responsiveness to chemotherapy and
  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
   platform_technology:
```

22

in situ oligonucleotide

```
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total)
varLabels: probeset gene
varMetadata: labelDescription
```

```
assayData: 19816 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:
  Length Class
                    Mode
      28 character character
sample_type:
tumor
  28
primarysite:
ov
28
days_to_death:
  Min. 1st Qu. Median Mean 3rd Qu.
                                     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 )

```
experimentData(eset):
Experiment data
 Experimenter name: Konstantinopoulos PA, Spentzos D, Karlan BY, Taniguchi T et al. Gene expression
 Laboratory: Konstantinopoulos, Cannistra 2010 hgu95
  Contact information:
 Title: Gene expression profile of BRCAness that correlates with responsiveness to chemotherapy and
  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:
        GPL8300
    platform_technology:
        in situ oligonucleotide
Preprocessing: rma
featureData(eset):
An object of class 'AnnotatedDataFrame'
    featureNames: AADAC AAK1 ... ZZZ3 (8995 total)
```

varLabels: probeset gene
varMetadata: labelDescription

```
assayData: 8995 features, 42 samples
Platform type: hgu95av2
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 23.00 3.78 2.79
                                             NA
_____
Available sample meta-data:
_____
alt_sample_name:
  Length Class
                    Mode
      42 character character
sample_type:
tumor
  42
primarysite:
ov
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, Laboratory: Meyniel, Sastre-Garau 2010 Contact information: Title: A genomic and transcriptomic approach for a differential diagnosis between primary and sec URL: PMIDs: 20492709

Abstract: A 277 word abstract is available. Use 'abstract' method.

## GSE20565\_eset

```
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
   platform_technology:
      in situ oligonucleotide
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total)
 varLabels: probeset gene
```

## Details

```
assayData: 19816 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 NA's
 63 33 44
```

varMetadata: labelDescription

```
summarystage:
early late NA's
        67
              46
   27
tumorstage:
            3
                 4 NA's
  1
       2
 18
       9
           52
               15 46
substage:
  а
       b
            c NA's
 14
      10
           55 61
grade:
       2
            3 NA's
   1
   6
      27
           63 44
batch:
  Length
             Class
                        Mode
     140 character character
uncurated_author_metadata:
  Length
           Class
                        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 )

```
experimentData(eset):
Experiment data
Experimenter name: EXpression Project for Oncology, International Genomics Consortium, www.intge
Laboratory: expO, 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
   platform_technology:
      in situ oligonucleotide
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total)
 varLabels: probeset gene
```

#### Details

```
assayData: 19816 features, 204 samples
Platform type: hgu133plus2
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
         Class
                   Mode
     204 character character
sample_type:
   benign borderline
                      tumor
       2
            8
                       194
histological_type:
  Length Class
                    Mode
     204 character character
primarysite:
      ov NA's
other
  23
      178
            3
summarygrade:
high low NA's
 91
    31 82
summarystage:
early late NA's
     87
            80
  37
```

varMetadata: labelDescription

```
tumorstage:
              3
                   4 NA's
   1
        2
  20
       14
             58
                  18
                        94
substage:
              c NA's
        b
   а
  17
       22
             79
                  86
grade:
        2
              3
                   4 NA's
   1
  11
       20
             83
                        82
                   8
age_at_initial_pathologic_diagnosis:
   Min. 1st Qu.
                  Median
                             Mean 3rd Qu.
                                              Max.
          45.00
  25.00
                   55.00
                            58.82
                                     65.00
                                             85.00
batch:
   Length
               Class
                           Mode
      204 character character
uncurated_author_metadata:
   Length
               Class
                           Mode
      204 character character
```

GSE26193\_eset

miR-141 and miR-200a act on ovarian tumorigenesis by controlling oxidative stress response.

## Description

Although there is evidence that redox regulation has an essential role in malignancies, its impact on tumor prognosis remains unclear. Here we show crosstalk between oxidative stress and the miR-200 family of microRNAs that affects tumorigenesis and chemosensitivity. miR-141 and miR-200a target p38?? and modulate the oxidative stress response. Enhanced expression of these microR-NAs mimics p38?? deficiency and increases tumor growth in mouse models, but it also improves the response to chemotherapeutic agents. High-grade human ovarian adenocarcinomas that accumulate miR-200a have low concentrations of p38?? and an associated oxidative stress signature. The miR200a-dependent stress signature correlates with improved survival of patients in response to treatment. Therefore, the role of miR-200a in stress could be a predictive marker for clinical outcome in ovarian cancer. In addition, although oxidative stress promotes tumor growth, it also sensitizes tumors to treatment, which could account for the limited success of antioxidants in clinical trials.

## Usage

data( GSE26193\_eset )

30

#### GSE26193\_eset

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Mateescu B, Batista L, Mariani O, Meyniel J, Cottu PH, Sastre-Garau X, Mechta-
  Laboratory: Mateescu, Mechta-Grigoriou 2011
  Contact information:
 Title: miR-141 and miR-200a act on ovarian tumorigenesis by controlling oxidative stress response
  URL:
  PMIDs: 22101765
  Abstract: A 149 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
   platform_technology:
      in situ oligonucleotide
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 19816 features, 107 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
107.00 107.00 107.00 76.00 3.05 2.50 4.56
-------
Available sample meta-data:
-------
alt_sample_name:
Length Class Mode
```

```
107 character character
sample_type:
tumor
 107
histological_type:
clearcell endo mucinous other ser
6 8 8 6 79
summarygrade:
high low
 67 40
summarystage:
early late
  31 76
tumorstage:
1 2 3 4
20 11 59 17
substage:
  a b cNA's
 16 12 62 17
grade:
1 2 3
7 33 67
days_to_tumor_recurrence:
  Min. 1st Qu. Median Mean 3rd Qu. Max.
3.0 340.5 584.0 1108.0 1525.0 7386.0
recurrence_status:
norecurrence recurrence
                    80
         27
days_to_death:
  Min. 1st Qu. Median Mean 3rd Qu.
                                        Max.
     3 668 1096 1520 2220
                                        7386
vital_status:
deceased living
     76
          31
batch:
          Class
  Length
                   Mode
     107 character character
uncurated_author_metadata:
  Length Class Mode
```

#### 107 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 appendix 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 )

```
experimentData(eset):
Experiment data
 Experimenter name: Bonome T, Levine DA, Shih J, Randonovich M, Pise-Masison CA, Bogomolniy F, Ozb
 Laboratory: Bonome, Birrer 2008
 Contact information:
 Title: A gene signature predicting for survival in suboptimally debulked patients with ovarian ca
  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
platform_technology:
    in situ oligonucleotide
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1CF A2M ... ZZZ3 (13104 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 13104 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:
  Length
          Class
                      Mode
     195 character character
sample_type:
healthy tumor
    10
          185
histological_type:
ser NA's
185 10
primarysite:
ov
195
summarygrade:
high NA's
185 10
```

```
summarystage:
late NA's
185 10
tumorstage:
  3 4 NA's
146 36 13
substage:
  b c NA's
  9 137 49
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median Mean 3rd Qu.
                                      Max.
                                              NA's
 26.00 52.00 63.00 61.54 70.00 84.00
                                               13
recurrence_status:
norecurrence recurrence
               153
        42
days_to_death:
  Min. 1st Qu. Median Mean 3rd Qu.
                                              NA's
                                       Max.
  21.9 660.6 1164.0 1429.0 1880.0 4982.0
                                               10
vital_status:
                  NA's
deceased living
    129
             56
                    10
debulking:
  optimal suboptimal
                        NA's
       90
                 95
                          10
percent_normal_cells:
20-
195
percent_stromal_cells:
20-
195
percent_tumor_cells:
80+
195
batch:
  Length
          Class
                     Mode
     195 character character
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 )

```
experimentData(eset):
Experiment data
 Experimenter name: Gillet JP, Calcagno AM, Varma S, Davidson B et al. Multidrug resistance-linked
 Laboratory: Gillet, Gottesman 2012
  Contact information:
 Title: Multidrug resistance-linked gene signature predicts overall survival of patients with pri
 URL:
 PMIDs: 22492981
  Abstract: A 244 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      TagMan gRT-PCR Homo sapiens Low-Density Array 380
   platform_shorttitle:
      TaqMan qRT-PCR
   platform_summary:
      NA
   platform_manufacturer:
      TaqMan
   platform_distribution:
```

## GSE30009\_eset

```
custom
platform_accession:
    GPL13728
platform_technology:
    qRT-PCR
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: ABCA1 ABCA10 ... XRCC6 (359 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

# Details

```
assayData: 359 features, 103 samples
Platform type: NA
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
103.00 103.00 103.00 57.00 3.42 2.92 5.34
_____
Available sample meta-data:
_____
alt_sample_name:
                    Mode
  Length
          Class
     103 character character
sample_type:
borderline
             tumor
        1
              102
histological_type:
clearcell
              ser
              102
       1
summarygrade:
high low NA's
 92
      9
          2
summarystage:
late
103
tumorstage:
3 4
82 21
substage:
```

```
b
        c NA's
   2
       60
             41
grade:
        2
   1
              3 NA's
        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.

```
GSE30161_eset
```

## 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
  Laboratory: Ferriss, Lee 2012
  Contact information:
 Title: Multi-gene expression predictors of single drug responses to adjuvant chemotherapy in ovar
  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
   platform_technology:
      in situ oligonucleotide
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

## Details

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 NA's
 33 21 4
summarystage:
late
 58
tumorstage:
3 4
53 5
substage:
a b c
9 11 38
grade:
  1 2
         3 NA's
  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:
                                    NA's
norecurrence
                recurrence
                                       4
           6
                        48
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
                              NA's
                                 2
        26
                    30
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 cance
  Laboratory: Yoshihara, Tanaka 2012
  Contact information:
 Title: High-risk ovarian cancer based on 126-gene expression signature is uniquely characterized
  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
   platform_technology:
      in situ oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1BG-AS1 ... ZZZ3 (20106 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

## Details

```
assayData: 20106 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:
  a b cNA's
  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 cance Laboratory: Yoshihara, Tanaka 2012 Contact information: Title: High-risk ovarian cancer based on 126-gene expression signature is uniquely characterized 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
   platform_technology:
      in situ oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1BG-AS1 ... ZZZ3 (20106 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 20106 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:
  Length Class
                    Mode
      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:
       c NA's
  b
   3
       28
             9
grade:
 2 3
23 17
pltx:
у
40
tax:
у
40
days_to_death:
  Min. 1st Qu. Median
                           Mean 3rd Qu.
                                           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
```

GSE44104\_eset

*COL11A1 promotes tumor progression and predicts poor clinical outcome in ovarian cancer.* 

# Description

Biomarkers that predict disease progression might assist the development of better therapeutic strategies for aggressive cancers, such as ovarian cancer. Here, we investigated the role of collagen type XI alpha 1 (COL11A1) in cell invasiveness and tumor formation and the prognostic

impact of COL11A1 expression in ovarian cancer. Microarray analysis suggested that COL11A1 is a disease progression-associated gene that is linked to ovarian cancer recurrence and poor survival. Small interference RNA-mediated specific reduction in COL11A1 protein levels suppressed the invasive ability and oncogenic potential of ovarian cancer cells and decreased tumor formation and lung colonization in mouse xenografts. A combination of experimental approaches, including realtime RT-PCR, casein zymography and chromatin immunoprecipitation (ChIP) assays, showed that COL11A1 knockdown attenuated MMP3 expression and suppressed binding of Ets-1 to its putative MMP3 promoter-binding site, suggesting that the Ets-1-MMP3 axis is upregulated by COL11A1. Transforming growth factor (TGF)-beta (TGF-??1) treatment triggers the activation of smad2 signaling cascades, leading to activation of COL11A1 and MMP3. Pharmacological inhibition of MMP3 abrogated the TGF-??1-triggered, COL11A1-dependent cell invasiveness. Furthermore, the NF-YA-binding site on the COL11A1 promoter was identified as the major determinant of TGF-??1-dependent COL11A1 activation. Analysis of 88 ovarian cancer patients indicated that high COL11A1 mRNA levels are associated with advanced disease stage. The 5-year recurrence-free and overall survival rates were significantly lower (P=0.006 and P=0.018, respectively) among patients with high expression levels of tissue COL11A1 mRNA compared with those with low expression. We conclude that COL11A1 may promote tumor aggressiveness via the TGF-??1-MMP3 axis and that COL11A1 expression can predict clinical outcome in ovarian cancer patients.

## Usage

data( GSE44104\_eset )

#### Format

```
experimentData(eset):
Experiment data
  Experimenter name: Wu Y, Chang T, Huang Y, Huang H, Chou C
  Laboratory: Wu, Chou 2013
  Contact information:
 Title: COL11A1 promotes tumor progression and predicts poor clinical outcome in ovarian cancer.
  URL:
  PMIDs: 23934190
  Abstract: A 260 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
   platform_technology:
      in situ oligonucleotide
Preprocessing: frma
```

```
featureData(eset):
An object of class 'AnnotatedDataFrame'
featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total)
varLabels: probeset gene
varMetadata: labelDescription
```

```
assayData: 19816 features, 60 samples
Platform type: hgu133plus2
Binary overall survival summary (definitions of long and short provided by study authors):
long short
  44 16
_____
Available sample meta-data:
_____
alt_sample_name:
  Length Class
                     Mode
      60 character character
sample_type:
tumor
  60
histological_type:
clearcell endo mucinous
                             ser
     12
             11 9
                                28
summarystage:
early late
  25
      35
tumorstage:
1 2 3 4
17 8 30 5
recurrence_status:
norecurrence recurrence
        40
               20
os_binary:
long short
  44 16
relapse_binary:
long short
  40
       20
batch:
```

#### GSE49997\_eset

2010-09-07 2010-09-08 2010-10-14 2010-12-10 2010-12-14 20 2 18 16 4 uncurated\_author\_metadata: Length Class Mode 60 character character

GSE49997\_eset

Validating the impact of a molecular subtype in ovarian cancer on outcomes: a study of the OVCAD Consortium.

### Description

Most patients with epithelial ovarian cancer (EOC) are diagnosed at advanced stage and have a poor prognosis. However, a small proportion of these patients will survive, whereas others will die very quickly. Clinicopathological factors do not allow precise identification of these subgroups. Thus, we have validated a molecular subclassification as new prognostic factor in EOC. One hundred and ninety-four patients with Stage II-IV EOC were characterized by whole-genome expression profiling of tumor tissues and were classified using a published 112 gene set, derived from an International Federation of Gynecology and Obstetrics (FIGO) stage-directed supervised classification approach. The 194 tumor samples were classified into two subclasses comprising 95 (Subclass 1) and 99 (Subclass 2) tumors. All nine FIGO II tumors were grouped in Subclass 1 (P = 0.001). Subclass 2 (54% of advanced-stage tumors) was significantly correlated with peritoneal carcinomatosis and non-optimal debulking. Patients with Subclass 2 tumors had a worse overall survival for both serous and non-serous histological subtypes, as revealed by univariate analysis (hazard ratios [HR] of 3.17 and 17.11, respectively; P ??? 0.001) and in models corrected for relevant clinicopathologic parameters (HR 2.87 and 12.42, respectively; P ??? 0.023). Significance analysis of microarrays revealed 2082 genes that were differentially expressed in advanced-grade serous tumors of both subclasses and the focal adhesion pathway as the most deregulated pathway. In the present validation study, we have shown that, in advanced-stage serous ovarian cancer, two approximately equally large molecular subtypes exist, independent of classical clinocopathological parameters and presenting with highly different whole-genome expression profiles and a markedly different overall survival. Similar results were obtained in a small cohort of patients with non-serous tumors.?? 2012 Japanese Cancer Association.

#### Usage

```
data( GSE49997_eset )
```

### Format

experimentData(eset): Experiment data Experimenter name: Pils D1, Hager G, Tong D, Aust S, Heinze G, Kohl M, Schuster E, Wolf A, Sehouli Laboratory: Pils, Zeilinger 2012 Contact information: Title: Validating the impact of a molecular subtype in ovarian cancer on outcomes: a study of the URL: PMIDs: 22497737

```
Abstract: A 276 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
 notes:
   platform_title:
      ABI Human Genome Survey Microarray Version 2
   platform_shorttitle:
      ABI Human Genome
   platform_summary:
   platform_manufacturer:
      Applied Biosystems
   platform_distribution:
      commercial
   platform_accession:
      GPL2986
   platform_technology:
      in situ oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1CF ... ZZZ3 (16048 total)
 varLabels: probeset gene
 varMetadata: labelDescription
```

```
assayData: 16048 features, 204 samples
Platform type:
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
194.00 194.00 194.00 57.00
                               NA
                                      3.67
_____
Available sample meta-data:
_____
alt_sample_name:
  Length Class
                     Mode
     204 character character
sample_type:
tumor
 204
histological_type:
other ser NA's
      171 10
  23
```

NA

```
summarygrade:
high low NA's
143 50 11
summarystage:
early late NA's
      185
   9
            10
tumorstage:
  2 3
           4 NA's
  9 154
         31 10
grade:
  2 3 NA's
 50 143 11
age_at_initial_pathologic_diagnosis:
                                              NA's
  Min. 1st Qu. Median
                      Mean 3rd Qu.
                                      Max.
 26.00 50.00 57.00 57.66 67.00 85.00
                                              10
days_to_tumor_recurrence:
  Min. 1st Qu. Median
                       Mean 3rd Qu.
                                              NA's
                                       Max.
  30.0 335.0 487.0 580.1 722.5 1461.0
                                                10
recurrence_status:
norecurrence recurrence
                              NA's
        70
                   124
                               10
days_to_death:
  Min. 1st Qu. Median
                      Mean 3rd Qu.
                                              NA's
                                       Max.
  30.0 517.0 745.5 782.9 1027.0 1491.0
                                               10
vital_status:
deceased living
                   NA's
     57
           137
                    10
debulking:
  optimal suboptimal
                         NA's
      137
                 57
                          10
uncurated_author_metadata:
          Class
  Length
                      Mode
     204 character character
```

GSE51088\_eset

*POSTN/TGFBI-associated stromal signature predicts poor prognosis in serous epithelial ovarian cancer.* 

### Description

To identify molecular prognosticators and therapeutic targets for high-grade serous epithelial ovarian cancers (EOCs) using genetic analyses driven by biologic features of EOC pathogenesis.Ovarian tissue samples (n = 172; 122 serous EOCs, 30 other EOCs, 20 normal/benign) collected prospectively from sequential patients undergoing gynecologic surgery were analyzed using RNA expression microarrays. Samples were classified based on expression of genes with potential relevance in ovarian cancer. Gene sets were defined using Rosetta Similarity Search Tool (ROAST) and analysis of variance (ANOVA). Gene copy number variations were identified by array comparative genomic hybridization. No distinct subgroups of EOC could be identified by unsupervised clustering, however, analyses based on genes correlated with periostin (POSTN) and estrogen receptoralpha (ESR1) yielded distinct subgroups. When 95 high-grade serous EOCs were grouped by genes based on ANOVA comparing ESR1/WT1 and POSTN/TGFBI samples, overall survival (OS) was significantly shorter for 43 patients with tumors expressing genes associated with POSTN/TGFBI compared to 52 patients with tumors expressing genes associated with ESR1/WT1 (median 30 versus 49 months, respectively; P = 0.022). Several targets with therapeutic potential were identified within each subgroup. BRCA germline mutations were more frequent in the ESR1/WT1 subgroup. Proliferation-associated genes and TP53 status (mutated or wild-type) did not correlate with survival. Findings were validated using independent ovarian cancer datasets. Two distinct molecular subgroups of high-grade serous EOCs based on POSTN/TGFBI and ESR1/WT1 expressions were identified with significantly different OS. Specific differentially expressed genes between these subgroups provide potential prognostic and therapeutic targets. Copyright ?? 2013 Elsevier Inc. All rights reserved.

## Usage

data( GSE51088\_eset )

### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Karlan BY, Dering J, Walsh C, Orsulic S, Lester J, Anderson LA, Ginther CL, Fe
 Laboratory: Karlan, Slamon 2014
  Contact information:
 Title: POSTN/TGFBI-associated stromal signature predicts poor prognosis in serous epithelial ova
 URL:
 PMIDs: 24368280
 Abstract: A 250 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      Agilent-012097 Human 1A Microarray (V2) G4110B (Probe Name version)
   platform_shorttitle:
      Agilent G4110B
   platform_summary:
      hgug4110b
   platform_manufacturer:
      Agilent
   platform_distribution:
      commercial
```

## GSE51088\_eset

```
GPL7264
platform_technology:
    in situ oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1CF A2M ... ZZZ3 (8211 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

## Details

```
assayData: 8211 features, 172 samples
Platform type: hgug4110b
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
  20 observations deleted due to missingness
records n.max n.start events median 0.95LCL 0.95UCL
152.00 152.00 152.00 112.00 4.13 3.50 4.92
_____
Available sample meta-data:
_____
alt_sample_name:
  Length Class Mode
    172 character character
sample_type:
   benign borderline
                     healthy metastatic
                                         tumor
       5
           12
                     15
                            17
                                           123
histological_type:
         Class
                     Mode
  Length
     172 character character
summarygrade:
high low NA's
119
    30
         23
summarystage:
early late NA's
  31 120
            21
tumorstage:
  1 2 3 4 NA's
      9 103 17 21
 22
substage:
         c NA's
  a b
```

```
17
       22
            94
                 39
grade:
        1
             2
                  3 NA's
   0
   8
        8
            14 119
                      23
age_at_initial_pathologic_diagnosis:
   Min. 1st Qu. Median
                           Mean 3rd Qu.
                                           Max.
   26.0
           49.0
                   57.5
                           58.6
                                   68.0
                                           91.0
neo:
 n
172
recurrence_status:
                                  NA's
norecurrence recurrence
          36
                      111
                                    25
days_to_death:
  Min. 1st Qu. Median
                           Mean 3rd Qu.
                                           Max.
                                                   NA's
     30
            791
                   1491
                           1835
                                   2344
                                           7001
                                                     20
vital_status:
                      NA's
deceased
          living
     112
               40
                        20
percent_normal_cells:
 30- NA's
 140
     32
percent_stromal_cells:
 30- NA's
 140
      32
percent_tumor_cells:
 70+ NA's
 140
       32
uncurated_author_metadata:
   Length
           Class
                         Mode
      172 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
  Laboratory: Murph, Mills 2009
  Contact information:
 Title: Lysophosphatidic acid-induced transcriptional profile represents serous epithelial ovaria
  URI :
  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
   platform_technology:
      in situ oligonucleotide
Preprocessing: frma
featureData(eset):
```

```
An object of class 'AnnotatedDataFrame'
featureNames: A1CF A2M ... ZZZ3 (13104 total)
varLabels: probeset gene
varMetadata: labelDescription
```

```
assayData: 13104 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
                                     NA's
            37 13
                             41
     8
                                       4
primarysite:
ov
103
summarygrade:
high low NA's
 38 36 29
summarystage:
early late NA's
     53 8
  42
tumorstage:
 1 2
          3
              4 NA's
 35 11
         44
              9 4
substage:
  a b
              d NA's
          С
 19
      2
         54
              1 27
grade:
     2
         3 NA's
  1
 19
    17
         38 29
batch:
                    Mode
  Length
           Class
    103 character character
```

uncurated\_author\_metadata: Length Class 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
  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:
      commercial
   platform_accession:
      GPL80
   platform_technology:
      in situ oligonucleotide
Preprocessing: rma
featureData(eset):
An object of class 'AnnotatedDataFrame'
```

```
featureNames: A2M AADAC ... ZYX (5251 total)
varLabels: probeset gene
varMetadata: labelDescription
```

```
assayData: 5251 features, 66 samples
Platform type: hu6800
_____
Available sample meta-data:
_____
alt_sample_name:
  Length Class Mode
     66 character character
sample_type:
borderline cellline tumor
       6
                        58
         2
histological_type:
  Length Class
                  Mode
     66 character character
primarysite:
ov
66
summarygrade:
high low NA's
 40 15 11
grade:
  1 2 3 NA's
  1 14 40 11
batch:
  Length Class Mode
     66 character character
uncurated_author_metadata:
  Length Class Mode
     66 character character
```

GSE8842\_eset

Analysis of gene expression in early-stage ovarian cancer.

#### GSE8842\_eset

#### Description

Gene expression profile was analyzed in 68 stage I and 15 borderline ovarian cancers to determine if different clinical features of stage I ovarian cancer such as histotype, grade, and survival are related to differential gene expression. Tumors were obtained directly at surgery and immediately frozen in liquid nitrogen until analysis. Glass arrays containing 16,000 genes were used in a dualcolor assay labeling protocol.Unsupervised analysis identified eight major patient partitions, one of which was statistically associated to overall survival, grading, and histotype and another with grading and histotype. Supervised analysis allowed detection of gene profiles clearly associated to histotype or to degree of differentiation. No difference was found between borderline and grade 1 tumors. As to recurrence, a subset of genes able to differentiate relapsers from nonrelapsers was identified. Among these, cyclin E and minichromosome maintenance protein 5 were found particularly relevant, as their expression was inversely correlated to progression-free survival (P = 0.00033 and 0.017, respectively). Specific molecular signatures define different histotypes and prognosis of stage I ovarian cancer. Mucinous and clear cells histotypes can be distinguished from the others regardless of tumor grade. Cyclin E and minichromosome maintenance protein 5, whose expression was found previously to be related to a bad prognosis of advanced ovarian cancer, appear to be potential prognostic markers in stage I ovarian cancer too, independent of other pathologic and clinical variables.

## Usage

data( GSE8842\_eset )

#### Format

```
experimentData(eset):
Experiment data
 Experimenter name: Marchini S, Mariani P, Chiorino G, Marrazzo E, Bonomi R, Fruscio R, Clivio L,
 Laboratory: Marchini, D'Incalci 2008
  Contact information:
  Title: Analysis of gene expression in early-stage ovarian cancer.
 URL:
 PMIDs: 19047114
 Abstract: A 225 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      Agilent Human 1 cDNA Microarray (G4100A)
   platform_shorttitle:
      Agilent G4100A cDNA
   platform_summary:
      hgug4100a
   platform_manufacturer:
      Agilent
   platform_distribution:
      custom-commerical
   platform_accession:
      GPL5689
   platform_technology:
      spotted DNA/cDNA
```

```
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
featureNames: A2M AADAC ... ZYX (6536 total)
varLabels: probeset gene
varMetadata: labelDescription
```

```
assayData: 6536 features, 83 samples
Platform type: hgug4100a
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
    83
           83
                 83
                        15
                                 NA
                                        12
                                               NA
 _____
Available sample meta-data:
_____
alt_sample_name:
  Length Class
                      Mode
      83 character character
sample_type:
borderline
             tumor
      15
                 68
histological_type:
  Length
         Class
                      Mode
      83 character character
primarysite:
ov
83
summarygrade:
high low NA's
 35 33 15
summarystage:
early
  83
tumorstage:
1
83
substage:
a b c
25 5 53
```

```
grade:
   1
        2
             3 NA's
  13
       20
            35
                 15
age_at_initial_pathologic_diagnosis:
   Min. 1st Qu. Median
                            Mean 3rd Qu.
                                             Max.
          43.00
                  50.00
  21.00
                           51.25
                                   61.00
                                            87.00
recurrence_status:
norecurrence
               recurrence
          62
                        21
days_to_death:
   Min. 1st Qu.
                 Median
                            Mean 3rd Qu.
                                             Max.
      0
           1192
                    2248
                            2273
                                     3048
                                             5824
vital_status:
deceased
           living
      15
               68
uncurated_author_metadata:
   Length
              Class
                          Mode
       83 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 N Laboratory: Tothill, Bowtell 2008 Contact information: Title: Novel molecular subtypes of serous and endometrioid ovarian cancer linked to clinical outc 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 platform\_technology: in situ oligonucleotide Preprocessing: frma featureData(eset): An object of class 'AnnotatedDataFrame' featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total) varLabels: probeset gene

## Details

```
assayData: 19816 features, 285 samples
Platform type: hgu133plus2
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
```

varMetadata: labelDescription

\_\_\_\_\_

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

## GSE9891\_eset

```
Available sample meta-data:
_____
alt_sample_name:
  Length Class
                  Mode
    285 character character
sample_type:
borderline tumor
   18
            267
histological_type:
endo other ser
  20 1 264
primarysite:
  ft other ov
  8 34 243
arrayedsite:
  ft other ov
  2 83 200
summarygrade:
high low NA's
163 116 6
summarystage:
early late NA's
 42 240 3
tumorstage:
 1 2 3 4 NA's
 24 18 218 22 3
substage:
 a b cNA's
 26 19 212 28
grade:
 1 2 3 NA's
 19 97 163 6
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median Mean 3rd Qu. Max. NA's
 22.00 53.00 59.00 59.62 68.00 80.00
                                      3
pltx:
 n yNA's
 39 243 3
tax:
```

```
y NA's
   n
      195
  87
              3
neo:
        y NA's
   n
 264
       18
              3
days_to_tumor_recurrence:
   Min. 1st Qu.
                 Median
                            Mean 3rd Qu.
                                              Max.
                                                      NA's
    0.0
          300.0
                   450.0
                           618.9
                                    810.0
                                            4980.0
                                                         10
recurrence_status:
norecurrence
                                    NA's
                recurrence
          94
                       188
                                       3
days_to_death:
   Min. 1st Qu.
                  Median
                            Mean 3rd Qu.
                                                      NA's
                                              Max.
    0.0
          547.5
                   855.0
                           955.1 1252.0
                                           6420.0
                                                          7
vital_status:
                       NA's
deceased
           living
     113
               169
                          3
debulking:
   optimal suboptimal
                              NA's
       160
                    88
                                37
batch:
   Length
               Class
                          Mode
      285 character character
uncurated_author_metadata:
   Length
               Class
                          Mode
      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 29-fold 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 )
```

varMetadata: labelDescription

#### Format

experimentData(eset): Experiment data Experimenter name: Berchuck A, Iversen ES, Lancaster JM, Pittman J, Luo J, Lee P, Murphy S, Dress Laboratory: Berchuck, Marks 2005 Contact information: Title: Patterns of gene expression that characterize long-term survival in advanced stage serous URL: PMIDs: 15897565

```
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:
      commercial
   platform_accession:
      GPL96
   platform_technology:
      in situ oligonucleotide
   warnings:
      These samples are a subset of PMID17290060.
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1CF A2M ... ZZZ3 (13104 total)
  varLabels: probeset gene
```

```
assayData: 13104 features, 63 samples
Platform type: hgu133a
Binary overall survival summary (definitions of long and short provided by study authors):
long short NA's
  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 NA's
 25
    37 1
summarystage:
early late
  11
       52
tumorstage:
1 2 3 4
7 4 48 4
grade:
  1
       2
           3
               4 NA's
  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 NA's
  24 28 11
```

#### PMID17290060\_eset

debulking: optimal suboptimal NA's 24 28 11 batch: Length Class Mode 63 character character uncurated\_author\_metadata: Length Class 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, W Laboratory: Dressman, Lancaster 2007 Contact information: Title: An integrated genomic-based approach to individualized treatment of patients with advanced 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
   platform_technology:
      in situ oligonucleotide
   warnings:
      This paper has been retracted.
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1CF A2M ... ZZZ3 (13104 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

```
assayData: 13104 features, 117 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
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 NA's
 57
     57
            3
summarystage:
early late NA's
   1 115
              1
tumorstage:
   2
     3
            4 NA's
   1
       98
           17
                 1
grade:
   1
       2
            3
                 4 NA's
   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
        63
                  54
batch:
  Length
             Class
                        Mode
     117 character character
uncurated_author_metadata:
  Length
             Class
                        Mode
     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 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 M/
 Laboratory: Berchuck, Lancaster 2009
  Contact information:
 Title: Microarray analysis of early stage serous ovarian cancers shows profiles predictive of fav
 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
   platform_technology:
      in situ oligonucleotide
   warnings:
```

## PMID19318476\_eset

These samples are a subset of PMID17290060. Preprocessing: frma featureData(eset): An object of class 'AnnotatedDataFrame' featureNames: A1CF A2M ... ZZZ3 (13104 total)

varLabels: probeset gene
varMetadata: labelDescription

Details

```
assayData: 13104 features, 42 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
 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 NA's
 24 17 1
summarystage:
early late NA's
   2
      39
            1
tumorstage:
  1 2 3
             4 NA's
  1
      1
          29
             10 1
substage:
  a b
          c NA's
  1
      1
          29 11
```

grade:

1

```
2
             3 NA's
   1
       15
   2
            24
                   1
age_at_initial_pathologic_diagnosis:
                            Mean 3rd Qu.
   Min. 1st Qu.
                 Median
                                                      NA's
                                             Max.
  33.00
          55.00
                   62.00
                           61.46
                                    70.00
                                            81.00
recurrence_status:
norecurrence
                recurrence
           6
                        36
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
                20
      22
debulking:
   optimal suboptimal
                             NA's
        20
                                 1
                    21
batch:
   Length
              Class
                          Mode
       42 character character
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
   platform_technology:
      in situ oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: ebv-miR-BART10 ebv-miR-BART10* ... 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
    544.00 544.00 544.00 286.00 3.71 3.42 4.03
```

73

```
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 NA's
474 68 12
summarystage:
early late NA's
  39 511
          4
tumorstage:
 1 2 3 4 NA's
 16 23 426 85 4
substage:
 b c NA's
 31 434 89
grade:
 1 2 3 4 NA's
  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 yNA's
 19 478 57
```

74

\_\_\_\_\_

```
tax:
     y NA's
  n
 43 454 57
neo:
  n NA's
497 57
days_to_tumor_recurrence:
  Min. 1st Qu. Median Mean 3rd Qu. Max. NA's
   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.
                                             NA's
   8.0 346.0 867.5 997.7 1446.0 5480.0
                                              10
vital_status:
deceased
        living
                   NA's
                    7
    286
            261
site_of_tumor_first_recurrence:
              locoregional locoregional_plus_metastatic
                      145
                                                  3
                                               NA's
                metastasis
                                                268
                      138
primary_therapy_outcome_success:
 completeresponse partialresponse progressivedisease
                                                     stabledisease
             308
                               63
                                                41
                                                                  30
            NA's
             112
debulking:
  optimal suboptimal
                         NA's
      359
              137
                         58
percent_normal_cells:
  Min. 1st Qu. Median Mean 3rd Qu. Max.
                                             NA's
 0.000 0.000 0.000 2.375 0.000 55.000
                                              10
percent_stromal_cells:
                                              NA's
  Min. 1st Qu. Median
                      Mean 3rd Qu.
                                      Max.
  0.00 5.00 10.00 12.78 19.00 70.00
                                              16
percent_tumor_cells:
                                             NA's
  Min. 1st Qu. Median
                      Mean 3rd Qu.
                                      Max.
  0.00 75.00 85.00 80.72 90.00 100.00
                                              13
```

```
uncurated_author_metadata:
Length Class Mode
554 character character
```

TCGA. RNASeqV2\_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.RNASeqV2\_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.
 URI ·
 PMIDs: 21720365
 Abstract: A 179 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [RNASeqV2] Illumina HiSeq RNA sequencing
   platform_shorttitle:
      Illumina HiSeq RNA sequencing
   platform_summary:
      NA
   platform_manufacturer:
      Illumina
```

## TCGA.RNASeqV2\_eset

```
platform_distribution:
    sequencing
platform_accession:
    NA
platform_technology:
    RNA sequencing
```

```
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
featureNames: ? A1BG ... ZZZ3 (20502 total)
varLabels: probeset gene
varMetadata: labelDescription
```

## Details

```
assayData: 20502 features, 261 samples
Platform type: NA
Overall survival time-to-event summary (in years):
Call: survfit(formula = Surv(time, cens) ~ -1)
  5 observations deleted due to missingness
records n.max n.start events median 0.95LCL 0.95UCL
256.00 256.00 256.00 143.00 3.62 3.19 4.03
_____
Available sample meta-data:
_____
alt_sample_name:
         Class Mode
  Length
     261 character character
unique_patient_ID:
  Length
          Class
                      Mode
     261 character character
sample_type:
tumor
 261
histological_type:
ser
261
primarysite:
other
       ov
       260
   1
summarygrade:
high low NA's
226 29
         6
```

```
summarystage:
early late NA's
 18 242 1
tumorstage:
 2 3 4 NA's
 18 209 33 1
substage:
 b cNA's
 16 211 34
grade:
 1 2 3 4 NA's
  1 28 225 1 6
age_at_initial_pathologic_diagnosis:
 Min. 1st Qu. Median Mean 3rd Qu. Max.
 34.00 51.00 58.00 58.84 66.00 87.00
pltx:
 n yNA's
 17 215 29
tax:
 n yNA's
 17 215 29
neo:
 n NA's
232 29
days_to_tumor_recurrence:
  Min. 1st Qu. Median Mean 3rd Qu. Max. NA's
   9.0 225.0 426.5 585.3 755.0 5480.0 19
recurrence_status:
norecurrence recurrence
      123
            138
days_to_death:
  Min. 1st Qu. Median Mean 3rd Qu. Max. NA's
   9.0 341.8 878.0 1018.0 1446.0 5480.0 5
vital_status:
deceased living NA's
143 114 4
site_of_tumor_first_recurrence:
locoregional metastasis NA's
       82
            56
                          123
```

primary_therapy_						
completerespon		rtialres			stabledisease	
147 30					15	15
NA						
	54					
dobulling						
debulking:	+imal					
optimal subop						
171	60	30	)			
<pre>percent_normal_c</pre>	ماام					
Min. 1st Qu.		Maan	3rd Ou	Max	NA's	
0.000 0.000			-			
0.000 0.000	0.000	2.000	0.000	33.000	5	
<pre>percent_stromal_</pre>	cells:					
Min. 1st Qu.		Mean	3rd Ou.	Max.	NA's	
	10.00		15.00		4	
					·	
<pre>percent_tumor_ce</pre>	lls:					
Min. 1st Qu.	Median	Mean	3rd Qu.	Max.	NA's	
0.00 77.00	85.00	82.07	90.00	100.00	4	
uncurated_author	_metadata	a:				
Length Cl	ass	Mode				
261 charac	ter chara	acter				

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 )

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
   platform_technology:
      in situ oligonucleotide
   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'
  featureNames: A1CF A2M ... ZZZ3 (13104 total)
  varLabels: probeset gene
 varMetadata: labelDescription
```

# Details

```
assayData: 13104 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

\_\_\_\_\_

80

```
Available sample meta-data:
_____
alt_sample_name:
  Length Class
                  Mode
    578 character character
unique_patient_ID:
  Length Class
                  Mode
    578 character character
sample_type:
healthy tumor
   8 570
histological_type:
ser NA's
568 10
primarysite:
other ov NA's
  4 564 10
summarygrade:
high low NA's
480 75 23
summarystage:
early late NA's
 43 520 15
tumorstage:
 1 2 3 4 NA's
 16 27 436 84 15
substage:
 b cNA's
 31 448 99
grade:
  1 2 3 4 NA's
  6 69 479 1 23
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median Mean 3rd Qu. Max. NA's
 26.00 51.00 59.00 59.70 68.25 89.00 10
pltx:
 n yNA's
 19 492 67
```

```
n
       y NA's
  43 468 67
neo:
  n NA's
 511 67
days_to_tumor_recurrence:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                         Max.
                                                NA's
   8.0 238.2 443.5 623.7 812.0 5480.0
                                                  56
recurrence_status:
norecurrence recurrence
        279
                     299
days_to_death:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                         Max.
                                                 NA's
           349
                  881
                                         5480
     8
                         1010 1446
                                                  21
vital_status:
deceased
         living
                    NA's
             270
    290
                      18
site_of_tumor_first_recurrence:
               locoregional locoregional_plus_metastatic
                        153
                                                     3
                                                  NA's
                 metastasis
                       143
                                                   279
primary_therapy_outcome_success:
  completeresponse
                    partialresponse progressivedisease
                                                           stabledisease
              318
                                 65
                                                   41
                                                                      30
             NA's
              124
debulking:
   optimal suboptimal
                          NA's
      367
                140
                            71
percent_normal_cells:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                                NA's
                                         Max.
  0.000 0.000 0.000
                        2.385 0.000 55.000
                                                 19
percent_stromal_cells:
  Min. 1st Qu. Median
                        Mean 3rd Qu.
                                         Max.
                                                 NA's
        5.00 10.00
   0.00
                        12.85 20.00
                                        70.00
                                                  25
percent_tumor_cells:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                                 NA's
                                         Max.
   0.00 75.00 85.00 80.64 90.00 100.00
                                                  22
```

82

```
batch:
```

Min.	1st Qu.	Median	Mean 3	3rd Qu.	Max.	NA's
9.00	13.00	17.00	18.55	22.00	40.00	1

# uncurated\_author\_metadata:

Length Class Mode

578 character character

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