Package 'curatedOvarianData'

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Title Clinically Annotated Data for the Ovarian Cancer Transcriptome
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Maintainer Levi Waldron < lwaldron.research@gmail.com>
Description The curatedOvarianData package provides data for gene expression analysis in patients with ovarian cancer.
Depends R ($>= 3.0.0$), Biobase
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
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Contents
curatedOvarianData-package

GSE12418 eset	7
-	10
	12
	15
99217701_9990111111111111111111111111111111	17
05217200_0000 1111111111111111111111111111111	21
-	23
	26
	28
	31
	33
000000000000000000000000000000000000000	35
00220712_0000 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	39
	39 41
00200101_0000	41
05202002.0120.00_0000 111111111111111111111111111111	
05262000_0000	48
	50
00213337_0000 111111111111111111111111111111111	53
	56
052000_656011111111111111111111111111111111	59
	62
	63
	66
	70
PMID17290060_eset	73
PMID19318476_eset	75
TCGA.mirna.8x15kv2_eset	78
TCGA.RNASeqV2_eset	82
TCGA_eset	86
	91

curatedOvarianData-package

Clinically Annotated Data for the Ovarian Cancer Transcriptome

Description

Index

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: curatedOvarianData

Type: Package
Version: 1.41.1
Date: 2023-10-31
License: Artistic-2.0
Depends: 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)

4 E.MTAB.386_eset

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

Department of Biostatistics and Computational Biology, Dana-Farber Cancer Institute, Harvard School of Public Health

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)</pre>
```

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

E.MTAB.386_eset 5

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 )
```

```
experimentData(eset):
Experiment data
 Experimenter name: Bentink S, Haibe-Kains B, Risch T, Fan J-B, Hirsch MS, Holton K, Rubio R, April C, Ch
 Laboratory: Bentink, Matulonis 2012
 Contact information:
 Title: Angiogenic mRNA and microRNA gene expression signature predicts a novel subtype of serous ovaria
 URL:
 PMIDs: 22348002
 Abstract: A 212 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      Illumina humanRef-8 v2.0 expression beadchip
   platform shorttitle:
      Illumina humanRef-8 v2.0
   platform_summary:
      illuminaHumanv2
   platform manufacturer:
      Illumina
   platform_distribution:
      commercial
   platform_accession:
      GPL6104
   platform_technology:
      in situ oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A2M A4GALT ... ZZEF1 (10357 total)
```

6 E.MTAB.386_eset

varLabels: probeset gene
varMetadata: labelDescription

Details

```
assayData: 10357 features, 129 samples
Platform type: illuminaHumanv2
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
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
  1 128
tumorstage:
 2 3 4
 1 109 19
substage:
  а
     b
          c NA's
  5 12 93 19
```

GSE12418_eset 7

```
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 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 )
```

SE12418_eset

Format

```
experimentData(eset):
Experiment data
 Experimenter name: Partheen K, Levan K, Osterberg L, Horvath G.Expression analysis of stage III serous
 Laboratory: Partheen, Horvath 2006
 Contact information:
 Title: Expression analysis of stage III serous ovarian adenocarcinoma distinguishes a sub-group of sur
 URL:
  PMIDs: 16996261
  Abstract: A 177 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      SWEGENE H_v2.1.1_27k
   platform_shorttitle:
      SWEGENE H_v2.1.1_27k
   platform_summary:
      PartheenMetaData
   platform_manufacturer:
      other
   platform_distribution:
      non-commercial
   platform_accession:
      GPL5886
   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

GSE12418_eset 9

```
alt_sample_name:
  Length
              Class
                         Mode
       54 character character
sample_type:
tumor
   54
histological_type:
ser
54
primarysite:
ον
54
summarystage:
late
 54
tumorstage:
3
54
substage:
b c
19 35
{\tt age\_at\_initial\_pathologic\_diagnosis:}
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                           Max.
  35.00 51.25 59.50
                          59.56 69.75
                                           84.00
pltx:
У
54
os_binary:
long short
   20
        34
debulking:
   optimal suboptimal
        13
                   41
uncurated_author_metadata:
  Length
              Class
                         Mode
       54 character character
```

10 GSE12470_eset

GSE12470_eset

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

Description

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

Usage

```
data( GSE12470_eset )
```

Format

notes:

```
experimentData(eset):
Experiment data

Experimenter name: Yoshihara K, Tajima A, Komata D, Yamamoto T, Kodama S, Fujiwara H, Suzuki M, Onishi Y, Laboratory: Yoshihara, Tanaka 2009
Contact information:
Title: Gene expression profiling of advanced-stage serous ovarian cancers distinguishes novel subclass URL:
PMIDs: 19486012

Abstract: A 253 word abstract is available. Use 'abstract' method.
Information is available on: preprocessing
```

GSE12470_eset 11

```
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: 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:
  Length Class
                       Mode
      53 character character
sample_type:
healthy tumor
    10
        43
histological_type:
 ser NA's
 43 10
primarysite:
٥٧
53
summarystage:
early late NA's
```

12 GSE13876_eset

```
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 )
```

GSE13876_eset 13

Format

```
experimentData(eset):
Experiment data
 Experimenter name: Crijns AP, Fehrmann RS, de Jong S, Gerbens F, Meersma GJ, Klip HG, Hollema H, Hofstra
 Laboratory: Crijns, van der Zee 2009
 Contact information:
 Title: Survival-related profile, pathways, and transcription factors in ovarian cancer.
 URL:
 PMIDs: 19192944
 Abstract: A 371 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      Operon human v3 ~35K 70-mer two-color oligonucleotide microarrays
   platform_shorttitle:
      Operon v3 two-color
   platform_summary:
      OperonHumanV3
   platform_manufacturer:
      other
   platform_distribution:
      non-commercial
   platform_accession:
      GPL7759
   platform_technology:
      spotted oligonucleotide
Preprocessing: default
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1CF ... ZZZ3 (20577 total)
  varLabels: probeset gene
  varMetadata: labelDescription
```

Details

14 GSE13876_eset

```
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:
157
primarysite:
ον
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

GSE14764_eset 15

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 )
```

platform_title:

```
experimentData(eset):
Experiment data
Experimenter name: Denkert C, Budczies J, Darb-Esfahani S, Gy??rffy B et al. A prognostic gene expressi
Laboratory: Denkert, Lage 2009
Contact information:
Title: A prognostic gene expression index in ovarian cancer - validation across different independent URL:
PMIDs: 19294737
Abstract: A 254 word abstract is available. Use 'abstract' method.
Information is available on: preprocessing
```

16 GSE14764_eset

```
[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
Details
   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
   _____
   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:

norecurrence recurrence NA's

50 26 4

days_to_death:

Min. 1st Qu. Median Mean 3rd Qu. Max.

210 660 1050 1011 1328 2190

vital_status:

deceased living

21 59

batch:

Length Class Mode

80 character character

uncurated_author_metadata:

Length Class Mode

80 character character

GSE17260_eset

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

Description

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

Usage

```
data( GSE17260_eset )
```

platform_manufacturer:

Agilent

```
experimentData(eset):
Experiment data
 Experimenter name: Yoshihara K, Tajima A, Yahata T, Kodama S, Fujiwara H, Suzuki M, Onishi Y, Hatae M, S
 Laboratory: Yoshihara, Tanaka 2010
 Contact information:
 Title: Gene expression profile for predicting survival in advanced-stage serous ovarian cancer across
 URL:
 PMIDs: 20300634
 Abstract: A 257 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
  platform_title:
     Agilent-012391 Whole Human Genome Oligo Microarray G4112A
  platform_shorttitle:
     Agilent G4112A
  platform_summary:
     hgug4112a
```

```
platform_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
   Available sample meta-data:
   alt_sample_name:
      Length Class
         110 character character
   sample_type:
   tumor
     110
   histological_type:
   ser
   110
   primarysite:
    οv
   110
   summarygrade:
   high low
     43
         67
```

```
summarystage:
late
110
tumorstage:
3 4
93 17
substage:
  a b c NA'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
                   76
        34
days_to_death:
  Min. 1st Qu. Median
                       Mean 3rd Qu.
                                      Max.
    30 660
              915
                        1086 1530
                                      2430
vital_status:
deceased living
        64
     46
debulking:
  optimal suboptimal
       57
uncurated_author_metadata:
  Length Class
     110 character character
```

GSE18520_eset 21

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, Yip D
 Laboratory: Mok, Birrer 2009
 Contact information:
 Title: A gene signature predictive for outcome in advanced ovarian cancer identifies a survival factor
 URL:
  PMIDs: 19962670
  Abstract: A 110 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
   platform_shorttitle:
      Affymetrix HG-U133Plus2
   platform_summary:
      hgu133plus2
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL570
   platform_technology:
```

22 GSE18520_eset

```
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

assayData: 19816 features, 63 samples
Platform type: hgu133plus2
Overall survival time-to-event summary (in years):
```

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:

Call: survfit(formula = Surv(time, cens) ~ -1)

histological_type: ser NA's 53 10

 ${\tt primary site:}$

ov 63

summarygrade:
high NA's

53 10

 $\verb"summary stage":$

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
                    630
                           1212
                                   1440
                                           4500
                                                      10
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:
   Length
             Class
                         Mode
       63 character character
```

GSE19829.GPL570_eset

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

Description

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

Usage

```
data( GSE19829.GPL570_eset )
```

```
experimentData(eset):
Experiment data
 Experimenter name: Konstantinopoulos PA, Spentzos D, Karlan BY, Taniguchi T et al. Gene expression pro-
 Laboratory: Konstantinopoulos, Cannistra 2010 hgu133plus2
 Contact information:
 Title: Gene expression profile of BRCAness that correlates with responsiveness to chemotherapy and wit
 URL:
 PMIDs: 20547991
  Abstract: A 241 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
   platform_shorttitle:
      Affymetrix HG-U133Plus2
   platform_summary:
      hgu133plus2
   platform_manufacturer:
      Affymetrix
   platform_distribution:
```

```
commercial
      platform_accession:
         GPL570
      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
   assayData: 19816 features, 28 samples
   Platform type: hgu133plus2
   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
             28.00 28.00 17.00
                                     3.95
     28.00
                                             2.71
   _____
   Available sample meta-data:
   alt_sample_name:
      Length
                 Class
                            Mode
          28 character character
   sample_type:
   tumor
      28
   primarysite:
   ον
   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 )
```

Format

```
experimentData(eset):
Experiment data
Experimenter name: Konstantinopoulos PA, Spentzos D, Karlan BY, Taniguchi T et al. Gene expression pro-
Laboratory: Konstantinopoulos, Cannistra 2010 hgu95
Contact information:
```

Title: Gene expression profile of BRCAness that correlates with responsiveness to chemotherapy and wit

```
URL:
     PMIDs: 20547991
     Abstract: A 241 word abstract is available. Use 'abstract' method.
     Information is available on: preprocessing
     notes:
      platform_title:
         [HG_U95Av2] Affymetrix Human Genome U95 Version 2 Array
      platform_shorttitle:
         Affymetrix HG_U95Av2
      platform_summary:
         hgu95av2
      platform_manufacturer:
         Affymetrix
      platform_distribution:
         commercial
      platform_accession:
         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
Details
   assayData: 8995 features, 42 samples
   Platform type: hgu95av2
   Overall survival time-to-event summary (in years):
   Call: survfit(formula = Surv(time, cens) ~ -1)
             n.max n.start events median 0.95LCL 0.95UCL
   records
     42.00 42.00 42.00 23.00 3.78 2.79
   _____
   Available sample meta-data:
   _____
   alt_sample_name:
      Length
               Class
                           Mode
          42 character character
   sample_type:
   tumor
```

28 GSE20565_eset

```
42
primarysite:
ον
42
days_to_death:
   Min. 1st Qu.
                 Median
                            Mean 3rd Qu.
                                             Max.
                          1089.0 1485.0
          727.5 1155.0
                                           2040.0
vital_status:
deceased
           living
      23
batch:
2001-09-14 2001-12-14 2002-08-20 2003-09-09 2003-09-18
                     4
                                           13
                               14
uncurated_author_metadata:
   Length
              Class
       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. GSE20565_eset 29

In all pairs, the result of the transcriptomic analysis was concordant with that of the genomic analysis. In patients with ovarian carcinoma and a previous history of breast cancer, SNP array analysis can be used to distinguish primary and secondary ovarian tumors. Transcriptomic analysis may be used when primary breast tissue specimen is not available.

Usage

```
data( GSE20565_eset )
```

Format

```
experimentData(eset):
Experiment data
 Experimenter name: Meyniel JP, Cottu PH, Decraene C, Stern MH, Couturier J, Lebigot I, Nicolas A, Weber
 Laboratory: Meyniel, Sastre-Garau 2010
 Contact information:
 Title: A genomic and transcriptomic approach for a differential diagnosis between primary and secondar
 URL:
 PMIDs: 20492709
 Abstract: A 277 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
   platform_shorttitle:
      Affymetrix HG-U133Plus2
   platform_summary:
      hgu133plus2
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL570
   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

assayData: 19816 features, 140 samples

30 GSE20565_eset

```
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
        ٥v
  44
        96
summarygrade:
high low NA's
 63 33 44
summarystage:
early late NA's
  27 67 46
tumorstage:
          3 4 NA's
  1
       2
 18
       9 52 15 46
substage:
         c NA's
  a b
 14 10 55 61
grade:
      2
  1
         3 NA's
  6 27 63 44
batch:
  Length
            Class
                      Mode
     140 character character
uncurated_author_metadata:
  Length
           Class
                      Mode
     140 character character
```

GSE2109_eset 31

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.intgen.org
  Laboratory: exp0, IGC 2005
  Contact information:
  Title: IGC EXpression Project for Oncology
  URL:
  PMIDs: PMID unknown
  Abstract: A 8 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
      [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
   platform_shorttitle:
      Affymetrix HG-U133Plus2
   platform_summary:
      hgu133plus2
   platform_manufacturer:
      Affymetrix
   platform_distribution:
      commercial
   platform_accession:
      GPL570
   platform_technology:
      in situ oligonucleotide
Preprocessing: frma
featureData(eset):
An object of class 'AnnotatedDataFrame'
  featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total)
  varLabels: probeset gene
```

32 GSE2109_eset

varMetadata: labelDescription

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
                        194
histological_type:
  Length Class
                     Mode
     204 character character
primarysite:
other ov NA's
  23 178 3
summarygrade:
high low NA's
 91 31 82
summarystage:
early late NA's
  37 87 80
tumorstage:
  1 2 3 4 NA's
 20 14 58 18 94
substage:
         c NA's
 a
     b
 17 22
         79 86
grade:
  1
      2
          3
               4 NA's
     20
         83
               8 82
{\tt age\_at\_initial\_pathologic\_diagnosis:}
  Min. 1st Qu. Median Mean 3rd Qu.
                                    Max.
```

GSE26193_eset 33

```
25.00
          45.00
                   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 )
```

```
experimentData(eset):
Experiment data
Experimenter name: Mateescu B, Batista L, Mariani O, Meyniel J, Cottu PH, Sastre-Garau X, Mechta-Grigor 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
```

34 GSE26193_eset

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
   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:
             Class
                           Mode
      Length
         107 character character
   sample_type:
   tumor
     107
   histological_type:
             endo mucinous
                                 other
   clearcell
                                               ser
                   8 8
                                                79
           6
                                       6
```

GSE26712_eset 35

```
high low
 67 40
summarystage:
early late
  31 76
tumorstage:
1 2 3 4
20 11 59 17
substage:
         c NA's
  a b
 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
        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:
  Length
         Class Mode
     107 character character
uncurated_author_metadata:
  Length
         Class
     107 character character
```

summarygrade:

36 GSE26712_eset

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 rapeutic 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, Ozbun L,
 Laboratory: Bonome, Birrer 2008
 Contact information:
 Title: A gene signature predicting for survival in suboptimally debulked patients with ovarian cancer.
 URL:
 PMIDs: 18593951
  Abstract: A 238 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
  platform_title:
      [HG-U133A] Affymetrix Human Genome U133A Array
  platform_shorttitle:
     Affymetrix HG-U133A
  platform_summary:
     hgu133a
```

GSE26712_eset 37

```
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
Details
   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:
    ον
   195
   summarygrade:
```

38 GSE26712_eset

```
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
         42
               153
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
vital_status:
deceased living
                 NA's
    129
             56
                    10
debulking:
  optimal suboptimal
                         NA's
       90
                 95
                           10
percent_normal_cells:
20-
195
percent_stromal_cells:
20-
195
percent_tumor_cells:
+08
195
```

batch:

GSE30009_eset 39

```
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 )
```

Format

experimentData(eset):

Experiment data

Experimenter name: Gillet JP, Calcagno AM, Varma S, Davidson B et al. Multidrug resistance-linked gene Laboratory: Gillet, Gottesman 2012

Contact information:

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

JIL.

PMIDs: 22492981

40 GSE30009_eset

```
Abstract: A 244 word abstract is available. Use 'abstract' method.
     Information is available on: preprocessing
     notes:
      platform_title:
         TaqMan qRT-PCR Homo sapiens Low-Density Array 380
      platform_shorttitle:
         TaqMan qRT-PCR
      platform_summary:
         NA
      platform_manufacturer:
         TaqMan
      platform_distribution:
         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)
   records n.max n.start events median 0.95LCL 0.95UCL
    103.00 103.00 103.00 57.00
                                      3.42
                                              2.92
                                                       5.34
   Available sample meta-data:
   alt_sample_name:
      Length
                 Class
                            Mode
         103 character character
   sample_type:
   borderline
                   tumor
                    102
   histological_type:
```

```
clearcell
                ser
                102
summarygrade:
high low NA's
 92
        9
summarystage:
late
103
tumorstage:
3 4
82 21
substage:
  b
       c NA's
   2
       60
            41
grade:
  1
        2
            3 NA's
        5
            92
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.
            598
                   1053
                           1156
                                   1568
                                           4748
vital_status:
deceased
           living
      57
               46
debulking:
  optimal suboptimal
        81
                   22
uncurated_author_metadata:
             Class
  Length
      103 character character
```

GSE30161_eset

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

Description

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

Usage

```
data( GSE30161_eset )
```

Format

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

```
GPL570
      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
   assayData: 19816 features, 58 samples
   Platform type: hgu133plus2
   Overall survival time-to-event summary (in years):
   Call: survfit(formula = Surv(time, cens) ~ -1)
   records n.max n.start events median 0.95LCL 0.95UCL
     58.00 58.00 58.00 36.00
                                    4.19
                                           2.70 6.17
   _____
   Available sample meta-data:
   _____
   alt_sample_name:
                          Mode
      Length
                Class
          58 character character
   sample_type:
   tumor
      58
   histological_type:
      Length
               Class
                          Mode
          58 character character
   summarygrade:
   high low NA's
     33 21
   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.
  12.0 255.2 386.0 742.1 768.2 4208.0
recurrence_status:
                          NA's
norecurrence recurrence
            48
days_to_death:
  Min. 1st Qu. Median Mean 3rd Qu.
  49.0 585.2 1010.0 1375.0 2131.0 4208.0
vital_status:
deceased living
     36
         22
debulking:
  optimal suboptimal NA's
         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 cancer base Laboratory: Yoshihara, Tanaka 2012

Contact information:

Title: High-risk ovarian cancer based on 126-gene expression signature is uniquely characterized by do URL:

PMIDs: 22241791

```
Abstract: A 255 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
   platform_title:
     Agilent-014850 Whole Human Genome Microarray 4x44K G4112F (Probe Name vers
ion)
   platform_shorttitle:
      Agilent G4112F
   platform_summary:
      hgug4112a
   platform_manufacturer:
      Agilent
   platform_distribution:
      commercial
   platform_accession:
      GPL6480
   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)
records
       n.max n.start events median 0.95LCL 0.95UCL
260.00 260.00 260.00 121.00
                              4.93 4.11 6.58
_____
Available sample meta-data:
_____
alt_sample_name:
  Length
                     Mode
           Class
     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 c NA'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
     260 character character
```

48 GSE32063_eset

GSE32063_eset

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

Description

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

Usage

```
data( GSE32063_eset )
```

Format

```
experimentData(eset):
Experiment data
    Experimenter name: Yoshihara K, Tsunoda T, Shigemizu D, Fujiwara H et al. High-risk ovarian cancer base
    Laboratory: Yoshihara, Tanaka 2012
    Contact information:
    Title: High-risk ovarian cancer based on 126-gene expression signature is uniquely characterized by do
    URL:
    PMIDs: 22241791

Abstract: A 255 word abstract is available. Use 'abstract' method.
    Information is available on: preprocessing
    notes:
        platform_title:
            Agilent-014850 Whole Human Genome Microarray 4x44K G4112F (Probe Name vers
ion)
```

GSE32063_eset 49

```
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, 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
    -----
   Available sample meta-data:
   -----
   alt_sample_name:
      Length
               Class
                           Mode
          40 character character
   sample_type:
   tumor
      40
   histological_type:
   ser
    40
   summarygrade:
   high low
```

50 *GSE44104_eset*

17 23 summarystage: late 40 tumorstage: 3 4 31 9 substage: b c NA's 3 28 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:

GSE44104_eset

Length

Class

40 character character

Mode

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

GSE44104_eset 51

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:
```

52 *GSE44104_eset*

```
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
   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
          60 character character
   sample_type:
   tumor
      60
   histological_type:
   clearcell
              endo mucinous
                                       ser
          12
                   11
                                        28
   summarystage:
   early late
      25
            35
   tumorstage:
    1 2 3 4
   17 8 30 5
```

GSE49997_eset 53

```
recurrence_status:
norecurrence
               recurrence
                        20
os_binary:
long short
   44
relapse_binary:
long short
   40
         20
batch:
2010-09-07 2010-09-08 2010-10-14 2010-12-10 2010-12-14
                     2
                               18
                                           16
uncurated_author_metadata:
   Length
              Class
       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.

54 GSE49997_eset

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 J, Bra
 Laboratory: Pils, Zeilinger 2012
 Contact information:
 Title: Validating the impact of a molecular subtype in ovarian cancer on outcomes: a study of the OVCAD
 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
```

Details

```
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
```

GSE49997_eset 55

records n.max n.start events median 0.95LCL 0.95UCL 194.00 194.00 194.00 57.00 NA 3.67 NA Available sample meta-data: _____ alt_sample_name: Length Class Mode 204 character character sample_type: tumor 204 histological_type: other ser NA's 23 171 10 summarygrade: high low NA's 143 50 11 summarystage: early late NA's 9 185 10 tumorstage: 2 3 4 NA's 9 154 31 10 grade: 2 3 NA's 50 143 11 age_at_initial_pathologic_diagnosis: Min. 1st Qu. Median Mean 3rd Qu. Max. NA's 26.00 50.00 57.00 57.66 67.00 85.00 10 days_to_tumor_recurrence: Min. 1st Qu. Median Mean 3rd Qu. Max. NA's 30.0 335.0 487.0 580.1 722.5 1461.0 10 recurrence_status: norecurrence recurrence NA's

10

124

70

days_to_death:

56 GSE51088_eset

```
Min. 1st Qu.
                 Median
                            Mean 3rd Qu.
                                                      NA's
                                              Max.
                   745.5
                           782.9 1027.0
   30.0
          517.0
                                           1491.0
                                                        10
vital_status:
deceased
           living
                       NA's
      57
              137
                         10
debulking:
   optimal suboptimal
                              NA's
       137
                    57
                                10
uncurated_author_metadata:
   Length
              Class
      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 the rapeutic 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 )
```

GSE51088_eset 57

Format

```
experimentData(eset):
Experiment data
 Experimenter name: Karlan BY, Dering J, Walsh C, Orsulic S, Lester J, Anderson LA, Ginther CL, Fejzo M,
 Laboratory: Karlan, Slamon 2014
  Contact information:
 Title: POSTN/TGFBI-associated stromal signature predicts poor prognosis in serous epithelial ovarian
  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
   platform_accession:
      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
```

58 GSE51088_eset

```
alt_sample_name:
  Length
           Class
                      Mode
     172 character character
sample_type:
   benign borderline healthy metastatic tumor 5 12 15 17 123
histological_type:
  Length
            Class
                      Mode
     172 character character
summarygrade:
high low NA's
119 30 23
summarystage:
early late NA's
  31 120
tumorstage:
  1 2 3 4 NA's
 22
       9 103 17 21
substage:
          c NA's
  а
     b
 17
      22 94 39
grade:
     1 2 3 NA's
       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:
norecurrence recurrence
                               NA's
                                25
                111
         36
days_to_death:
```

Available sample meta-data:

GSE6008_eset 59

```
Min. 1st Qu.
                  Median
                             Mean 3rd Qu.
                                                       NA's
                                              Max.
     30
            791
                                              7001
                    1491
                             1835
                                     2344
                                                         20
vital_status:
deceased
           living
                       NA's
     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 60 GSE6008_eset

that transcriptional events mediated by LPA in the tumor microenvironment influence tumor progression through modulation of cell adhesion molecules like claudin-1 and, for the first time, report an LPA-mediated expression signature in ovarian cancer that predicts a worse prognosis.

Usage

data(GSE6008_eset)

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

Details

assayData: 13104 features, 103 samples

Platform type: hgu133a

GSE6008_eset 61

```
Available sample meta-data:
alt_sample_name:
  Length Class
                    Mode
    103 character character
sample_type:
healthy tumor
          99
histological_type:
                                    NA's
clearcell endo mucinous ser
            37
                 13
                             41
                                      4
primarysite:
ον
103
summarygrade:
high low NA's
 38 36 29
summarystage:
early late NA's
  42 53 8
tumorstage:
 1 2 3 4 NA's
 35 11 44 9 4
substage:
        c d NA's
 a b
      2 54 1 27
 19
grade:
 1 2 3 NA's
 19 17 38 29
batch:
        Class
                  Mode
  Length
    103 character character
uncurated_author_metadata:
  Length Class Mode
    103 character character
```

62 *GSE6822_eset*

GSE6822_eset

Classification of ovarian tumor samples

Description

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

Usage

```
data( GSE6822_eset )
```

Format

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

Details

```
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
                   2
                            58
histological_type:
  Length
          Class
                       Mode
      66 character character
primarysite:
٥v
66
summarygrade:
high low NA's
 40
     15 11
grade:
       2
           3 NA's
  1
  1
     14 40 11
batch:
  Length
             Class
                       Mode
      66 character character
uncurated_author_metadata:
  Length
            Class
                       Mode
      66 character character
```

64 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 )
```

platform_accession:

Format

```
experimentData(eset):
Experiment data
 Experimenter name: Marchini S, Mariani P, Chiorino G, Marrazzo E, Bonomi R, Fruscio R, Clivio L, Garbi
 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
```

GSE8842_eset 65

```
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
Details
   assayData: 6536 features, 83 samples
   Platform type: hgug4100a
   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
              83
        83
                      83
                          15
                                     NA
                                            12
   _____
   Available sample meta-data:
   _____
   alt_sample_name:
                          Mode
      Length
                Class
          83 character character
   sample_type:
   borderline
                  tumor
                     68
   histological_type:
      Length
                Class
                          Mode
          83 character character
   primarysite:
   ον
   83
   summarygrade:
   high low NA's
     35 33 15
   summarystage:
   early
      83
```

```
tumorstage:
1
83
substage:
a b c
  5 53
25
grade:
   1
        2
             3 NA's
  13
       20
            35
                 15
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median
                           Mean 3rd Qu.
                                            Max.
  21.00
          43.00
                  50.00
                           51.25
                                 61.00
                                           87.00
recurrence_status:
norecurrence
               recurrence
days_to_death:
  Min. 1st Qu.
                            Mean 3rd Qu.
                 Median
                                            Max.
      0
           1192
                   2248
                            2273
                                    3048
                                             5824
vital_status:
deceased
           living
      15
uncurated_author_metadata:
  Length
              Class
       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 k-means 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 )
```

featureData(eset):

Format

```
experimentData(eset):
Experiment data
 Experimenter name: Tothill RW, Tinker AV, George J, Brown R, Fox SB, Lade S, Johnson DS, Trivett MK, Ete
 Laboratory: Tothill, Bowtell 2008
 Contact information:
 Title: Novel molecular subtypes of serous and endometrioid ovarian cancer linked to clinical outcome.
 URL:
 PMIDs: 18698038
 Abstract: A 243 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
  platform_title:
     [HG-U133_Plus_2] Affymetrix Human Genome U133 Plus 2.0 Array
  platform_shorttitle:
      Affymetrix HG-U133Plus2
  platform_summary:
     hgu133plus2
  platform_manufacturer:
     Affymetrix
  platform_distribution:
      commercial
  platform_accession:
     GPL570
  platform_technology:
     in situ oligonucleotide
Preprocessing: frma
```

An object of class 'AnnotatedDataFrame'

varLabels: probeset gene

featureNames: A1BG A1BG-AS1 ... ZZZ3 (19816 total)

```
varMetadata: labelDescription
Details
   assayData: 19816 features, 285 samples
   Platform type: hgu133plus2
   Overall survival time-to-event summary (in years):
   Call: survfit(formula = Surv(time, cens) ~ -1)
      7 observations deleted due to missingness
   records n.max n.start events median 0.95LCL 0.95UCL
    278.00 278.00 278.00 113.00
                                    3.95 3.53
   _____
   Available sample meta-data:
   alt_sample_name:
      Length
                Class
                           Mode
         285 character character
   sample_type:
   borderline
                  tumor
           18
                    267
   histological_type:
    endo other
      20
                264
            1
   primarysite:
      ft other
                 ΟV
       8
           34
                243
   arrayedsite:
      ft other
                 ٥v
       2
           83
                200
   summarygrade:
   high low NA's
    163 116
   summarystage:
   early late NA's
      42 240
```

```
1 2 3 4 NA's
 24 18 218 22 3
substage:
 a b c NA'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 y NA's
 39 243 3
tax:
n y NA's
 87 195 3
neo:
n y NA's
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 recurrence
                         NA's
        94
           188
                          3
days_to_death:
  Min. 1st Qu. Median Mean 3rd Qu. Max. NA's
   0.0 547.5 855.0 955.1 1252.0 6420.0
vital_status:
deceased living NA's 113 169 3
debulking:
  optimal suboptimal NA's 160 88 37
```

tumorstage:

PMID15897565_eset

```
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 29fold compared with early-stage cases). These results suggest that gene expression patterns underlie differences in outcome, and an examination of the genes that provide this discrimination reveals that many are implicated in processes that define the malignant phenotype. Differences in survival of advanced ovarian cancers are reflected by distinct patterns of gene expression. This biological distinction is further emphasized by the finding that early-stage cancers share expression patterns with the advanced stage long-term survivors, suggesting a shared favorable biology.

Usage

```
data( PMID15897565_eset )
```

Format

```
experimentData(eset):
Experiment data
Experimenter name: Berchuck A, Iversen ES, Lancaster JM, Pittman J, Luo J, Lee P, Murphy S, Dressman HK
Laboratory: Berchuck, Marks 2005
Contact information:
```

Title: Patterns of gene expression that characterize long-term survival in advanced stage serous ovari

```
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
     varMetadata: labelDescription
Details
   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:
οv
63
summarygrade:
high low NA's
 25 37 1
summarystage:
early late
  11 52
tumorstage:
1 2 3 4
7 4 48 4
grade:
       2 3 4 NA's
  1
  2 35 24
              1 1
{\tt 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
           11
  24 28
debulking:
  optimal suboptimal
                        NA's
       24
                 28
                           11
batch:
            Class
  Length
                       Mode
      63 character character
uncurated_author_metadata:
  Length
            Class
      63 character character
```

PMID17290060_eset 73

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 )
```

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

74 PMID17290060_eset

```
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
Details
   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:
    ον
   117
```

PMID19318476_eset 75

```
summarygrade:
high low NA's
 57 57
summarystage:
early late NA's
   1
      115
tumorstage:
   2
       3
            4 NA's
   1
       98
            17
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
                           1496
                  1020
                                   2220
                                           5550
vital_status:
deceased
          living
      67
               50
primary_therapy_outcome_success:
  completeresponse progressivedisease
                                   32
debulking:
   optimal suboptimal
        63
batch:
              Class
                         Mode
   Length
      117 character character
uncurated_author_metadata:
   Length
             Class
      117 character character
```

PMID19318476_eset

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

76 PMID19318476_eset

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 )
```

```
experimentData(eset):
Experiment data
 Experimenter name: Berchuck A, Iversen ES, Luo J, Clarke JP, Horne H, Levine DA, Boyd J, Alonso MA, Seco
 Laboratory: Berchuck, Lancaster 2009
 Contact information:
 Title: Microarray analysis of early stage serous ovarian cancers shows profiles predictive of favorabl
 URL:
 PMIDs: 19318476
 Abstract: A 241 word abstract is available. Use 'abstract' method.
  Information is available on: preprocessing
  notes:
  platform_title:
      [HG-U133A] Affymetrix Human Genome U133A Array
  platform_shorttitle:
     Affymetrix HG-U133A
  platform_summary:
     hgu133a
  platform_manufacturer:
     Affymetrix
  platform_distribution:
      commercial
  platform_accession:
```

```
GPL96
      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
     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)
             n.max n.start events median 0.95LCL 0.95UCL
   records
           42.00 42.00 22.00
                                     2.79
     42.00
                                          2.30
   _____
   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
   summarystage:
   early late NA's
       2
          39
                 1
   tumorstage:
```

```
3
                  4 NA's
            29
       1
                 10
substage:
  а
             c NA's
  1
       1
            29
                 11
grade:
   1
       2
             3 NA's
   2
       15
            24
age_at_initial_pathologic_diagnosis:
  Min. 1st Qu. Median
                                                   NA's
                           Mean 3rd Qu.
                                           Max.
 33.00 55.00
                62.00
                          61.46
                                70.00
                                          81.00
                                                      1
recurrence_status:
norecurrence
             recurrence
days_to_death:
  Min. 1st Qu. Median
                           Mean 3rd Qu.
                                           Max.
   30.0 367.5
                825.0 1105.0 1050.0 3420.0
vital_status:
deceased
           living
     22
               20
debulking:
  optimal suboptimal
                            NA's
       20
                   21
                               1
batch:
  Length
              Class
                         Mode
       42 character character
uncurated_author_metadata:
  Length
              Class
       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 )
```

```
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:
   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
_____
Available sample meta-data:
-----
alt_sample_name:
  Length
           Class
                      Mode
     554 character character
unique_patient_ID:
  Length
           Class
     554 character character
sample_type:
tumor
 554
histological_type:
ser
554
primarysite:
other
       ٥v
       550
summarygrade:
high low NA's
     68 12
474
summarystage:
early late NA's
  39 511
```

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 y NA's
 19 478 57
tax:
  n y NA's
 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
                   286
       268
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
    286
        261
site_of_tumor_first_recurrence:
             locoregional locoregional_plus_metastatic
                     145
                                                3
               metastasis
                                              NA's
```

268

primary_therapy_ completerespon 3 NA 1	success: tialresponse progressive 63			edisease 41	stabledisease 30	
debulking:						
optimal suboptimal		NA's				
359	137	58	3			
percent_normal_c		Maaa	2	Marr	NIA I -	
Min. 1st Qu.			=			
0.000 0.000	0.000	2.375	0.000	55.000	10	
percent_stromal_	cells:					
Min. 1st Qu.	Median	Mean	3rd Qu.	Max.	NA's	
0.00 5.00	10.00	12.78	19.00	70.00	16	
percent_tumor_ce Min. 1st Qu. 0.00 75.00	Median		=	Max. 100.00	NA's 13	
uncurated_author Length Cl 554 charac	ass	Mode				

138

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.
 URL:
 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
   platform_distribution:
      sequencing
   platform_accession:
   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:
  Length Class
                      Mode
     261 character character
unique_patient_ID:
  Length
           Class
                      Mode
     261 character character
sample_type:
tumor
 261
histological_type:
ser
261
primarysite:
other ov
   1
       260
summarygrade:
high low NA's
226 29
summarystage:
early late NA's
  18 242 1
tumorstage:
  2 3 4 NA's
 18 209 33 1
substage:
  b c NA'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.
               58.00
  34.00 51.00
                        58.84 66.00
                                        87.00
pltx:
  n
       y NA's
 17 215 29
tax:
       y NA's
  n
 17 215 29
neo:
  n NA's
 232 29
days_to_tumor_recurrence:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                                 NA's
                                         Max.
   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.
                                                 NA's
                                         Max.
   9.0 341.8 878.0 1018.0 1446.0 5480.0
vital_status:
deceased
                     NA's
          living
    143
             114
site_of_tumor_first_recurrence:
                                NA's
locoregional metastasis
         82
                      56
                                 123
primary_therapy_outcome_success:
  completeresponse
                     partialresponse progressivedisease
                                                           stabledisease
              147
                                 30
                                                   15
                                                                      15
             NA's
               54
debulking:
  optimal suboptimal
                          NA's
      171
                  60
                            30
percent_normal_cells:
  Min. 1st Qu. Median
                         Mean 3rd Qu.
                                                 NA's
                                         Max.
  0.000 0.000 0.000
                        2.066 0.000 55.000
                                                   5
```

```
percent_stromal_cells:
   Min. 1st Qu.
                            Mean 3rd Qu.
                                                      NA's
                 Median
                                             Max.
   0.00
           5.00
                  10.00
                           11.43
                                    15.00
                                            70.00
percent_tumor_cells:
   Min. 1st Qu.
                            Mean 3rd Qu.
                                                      NA's
                 Median
                                             Max.
   0.00
          77.00
                  85.00
                                    90.00
                           82.07
                                           100.00
                                                         4
uncurated_author_metadata:
   Length
              Class
                          Mode
      261 character character
```

TCGA_eset

Integrated genomic analyses of ovarian carcinoma.

Description

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

Usage

```
data( TCGA_eset )
```

```
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
            n.max n.start events median 0.95LCL 0.95UCL
    557.00 557.00 557.00 290.00
                                      3.73
                                               3.45
                                                       4.06
   Available sample meta-data:
   alt_sample_name:
      Length Class
                            Mode
```

```
578 character character
unique_patient_ID:
  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 c NA'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 y NA's 19 492 67

n y NA's

tax:

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 8 349 881 1010 1446 5480 21

vital_status:

deceased living NA's 290 270 18

site_of_tumor_first_recurrence:

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. Max. NA's 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 0.00 5.00 10.00 12.85 20.00 70.00 25

percent_tumor_cells:

Min. 1st Qu. Median Mean 3rd Qu. Max. NA's

0.00 75.00 85.00 80.64 90.00 100.00 22

batch:

Min. 1st Qu. Median Mean 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

Index

* datasets	GSE19829.GPL570_eset			
E.MTAB.386_eset, 4	(GSE19829.GPL570_eset), 23			
GSE12418_eset, 7	GSE19829.GPL8300_eset			
GSE12470_eset, 10	(GSE19829.GPL8300_eset), 26			
GSE13876_eset, 12	GSE20565_eset (GSE20565_eset), 28			
GSE14764_eset, 15	GSE2109_eset (GSE2109_eset), 31			
GSE17260_eset, 17	GSE26193_eset (GSE26193_eset), 33			
GSE18520_eset, 21	GSE26712_eset (GSE26712_eset), 36			
GSE19829.GPL570_eset, 23	GSE30009_eset (GSE30009_eset), 39			
GSE19829.GPL8300_eset, 26	GSE30161_eset (GSE30161_eset),41			
GSE20565_eset, 28	GSE32062.GPL6480_eset			
GSE2109_eset, 31	(GSE32062.GPL6480_eset), 45			
GSE26193_eset, 33	GSE32063_eset (GSE32063_eset), 48			
GSE26712_eset, 36	GSE44104_eset (GSE44104_eset), 50			
GSE30009_eset, 39	GSE49997_eset (GSE49997_eset), 53			
GSE30161_eset, 41	GSE51088_eset (GSE51088_eset), 56			
GSE32062.GPL6480_eset, 45	GSE6008_eset (GSE6008_eset), 59			
GSE32063_eset, 48	GSE6822_eset (GSE6822_eset), 62			
GSE44104_eset, 50	GSE8842_eset (GSE8842_eset), 63			
GSE49997_eset, 53	GSE9891_eset (GSE9891_eset), 66			
GSE51088_eset, 56	PMID15897565_eset			
GSE6008_eset, 59	(PMID15897565_eset), 70			
GSE6822_eset, 62	PMID17290060_eset			
GSE8842_eset, 63	(PMID17290060_eset), 73			
GSE9891_eset, 66	PMID19318476_eset			
PMID15897565_eset, 70	(PMID19318476_eset), 75			
PMID17290060_eset, 73	TCGA.mirna.8x15kv2_eset			
PMID19318476_eset, 75	(TCGA.mirna.8x15kv2_eset), 78			
TCGA.mirna.8x15kv2_eset, 78	TCGA.RNASeqV2_eset			
TCGA.RNASeqV2_eset, 82	(TCGA.RNASeqV2_eset), 82			
TCGA_eset, 86	TCGA_eset (TCGA_eset), 86			
rcda_eset, 80				
E MTAD 200 (E MTAD 200) 4	curatedOvarianData			
E.MTAB.386_eset (E.MTAB.386_eset), 4	(curatedOvarianData-package), 2			
GSE12418_eset (GSE12418_eset), 7	curatedOvarianData-package, 2			
GSE12470_eset (GSE12470_eset), 10				
GSE13876_eset (GSE13876_eset), 12	E.MTAB.386_eset, 4			
GSE14764_eset (GSE14764_eset), 15	20542442			
GSE17260_eset (GSE17260_eset), 17	GSE12418_eset, 7			
GSE18520_eset (GSE18520_eset), 21	GSE12470_eset, 10			

92 INDEX

```
GSE13876_eset, 12
GSE14764_eset, 15
GSE17260_eset, 17
GSE18520_eset, 21
GSE19829.GPL570_eset, 23
GSE19829.GPL8300_eset, 26
GSE20565_eset, 28
GSE2109_eset, 31
GSE26193_eset, 33
GSE26712_eset, 35
GSE30009_eset, 39
GSE30161_eset, 41
GSE32062.GPL6480_eset, 45
GSE32063_eset, 48
GSE44104_eset, 50
GSE49997_eset, 53
GSE51088_eset, 56
GSE6008_eset, 59
GSE6822_eset, 62
GSE8842_eset, 63
GSE9891_eset, 66
PMID15897565_eset, 70
PMID17290060_eset, 73
PMID19318476_eset, 75
TCGA.mirna.8x15kv2_eset, 78
TCGA.RNASeqV2_eset, 82
TCGA_eset, 86
```