



Sciences Economiques et Sociales de la Santé  
& Traitement de l'Information Médicale

[sesstim.univ-amu.fr](http://sesstim.univ-amu.fr)

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**Enjeux des entrepôts de données de santé**

**janvier 2018**



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Hôpital européen Georges-Pompidou

ASSISTANCE  
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DE PARIS

Necker   
ENFANTS MALADES  
HÔPITAL UNIVERSITAIRE

# Enjeux des entrepôts de données de santé

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Webinar SESSTIM 19 Jan 2018



# Plan

- Éléments de contexte
- La recherche sur les EDS
- Les EDS en support/complément de la recherche clinique
- Les EDS en support de la décision clinique

# Hôpital Européen Georges Pompidou

Ouverture en **2000**

Hôpital de l'AP-HP

700 lits

Spécialisation en **oncologie**, **maladies cardio-vasculaires** et **polytraumatisés**



Très impliqué dans le numérique:

- Electronic Health Record depuis 2000
- Prescription de médicaments (CPOE)
- Examens de laboratoire (robot)
- EDS i2b2 depuis 2008

**HIMMS level 6**

(<http://www.himss.eu/node/1116>)

# European EMR Adoption Model

2013



Stage	Cumulative Capabilities
Stage 7	Complete EMR; CCD transactions to share data; Data warehousing feeding outcomes reports, quality assurance, and business intelligence; Data continuity with ED, ambulatory, OP.
Stage 6	Physician documentation interaction with full CDSS (structured templates related to clinical protocols trigger variance & compliance alerts) and Closed loop medication administration.
Stage 5	Full complement of PACS displaces all film-based images.
Stage 4	CPOE in at least one clinical service area and/or for medication (i.e. e-Prescribing); may have Clinical Decision Support based on clinical protocols.
Stage 3	Nursing/clinical documentation (flow sheets); may have Clinical Decision Support for error checking during order entry and/or PACS available outside Radiology.
Stage 2	Clinical Data Repository (CDR) / Electronic Patient Record; may have Controlled Medical Vocabulary, Clinical Decision Support (CDS) for rudimentary conflict checking, Document Imaging and health information exchange (HIE) capability.
Stage 1	Ancillaries – Lab, Radiology, Pharmacy – All Installed OR processing LIS, RIS, PHIS data output online from external service providers.
Stage 0	All Three Ancillaries (LIS, RIS, PHIS) Not Installed OR Not processing Lab, Radiology, Pharmacy data output online from external service providers.

# Hôpital Necker et Institut Imagine

adulte AP-HP association cardiologie chirurgie  
développement durable hôpital Droit école  
éducation thérapeutique enfant Espace Plein  
Ciel greffe de rein halte garderie Hôpital  
Necker Laboratoires loisir  
maladies rares maternité mécénat  
**Necker** Necker-Enfants malades  
néphrologie Noël Paris pédiatrie Plan d'accès  
radiologie rein soin transplantation

« hôpital translationnel »



Institut Imagine  
Maladies génétiques  
410 personnes  
25 laboratoires

# La recherche sur les EDS



© 2011 Thomas G. Murphy, MD.

Tiré de Thomas McMurphy, repris dans la thèse de N. Garcelon 7





Published in final edited form as:

*Clin Pharmacol Ther.* 2011 July ; 90(1): 133–142. doi:10.1038/clpt.2011.83.

### **Detecting Drug Interactions From Adverse-Event Reports: Interaction Between Paroxetine and Pravastatin Increases Blood Glucose Levels**

**NP Tatonetti<sup>1,2,3</sup>, JC Denny<sup>4,5</sup>, SN Murphy<sup>6,7</sup>, GH Fernald<sup>1,2,3</sup>, G Krishnan<sup>8</sup>, V Castro<sup>6</sup>, P Yue<sup>9</sup>, PS Tsau<sup>9</sup>, I Kohane<sup>7,10,11</sup>, DM Roden<sup>5</sup>, and RB Altman<sup>2,3</sup>**

<sup>1</sup>Biomedical Informatics Training Program, Stanford University, Stanford, California, USA

<sup>2</sup>Department of Bioengineering, Stanford University, Stanford, California, USA

<sup>3</sup>Department of Genetics, Stanford University, Stanford, California, USA

<sup>4</sup>Department of Biomedical Informatics, Vanderbilt University, Nashville, Tennessee, USA

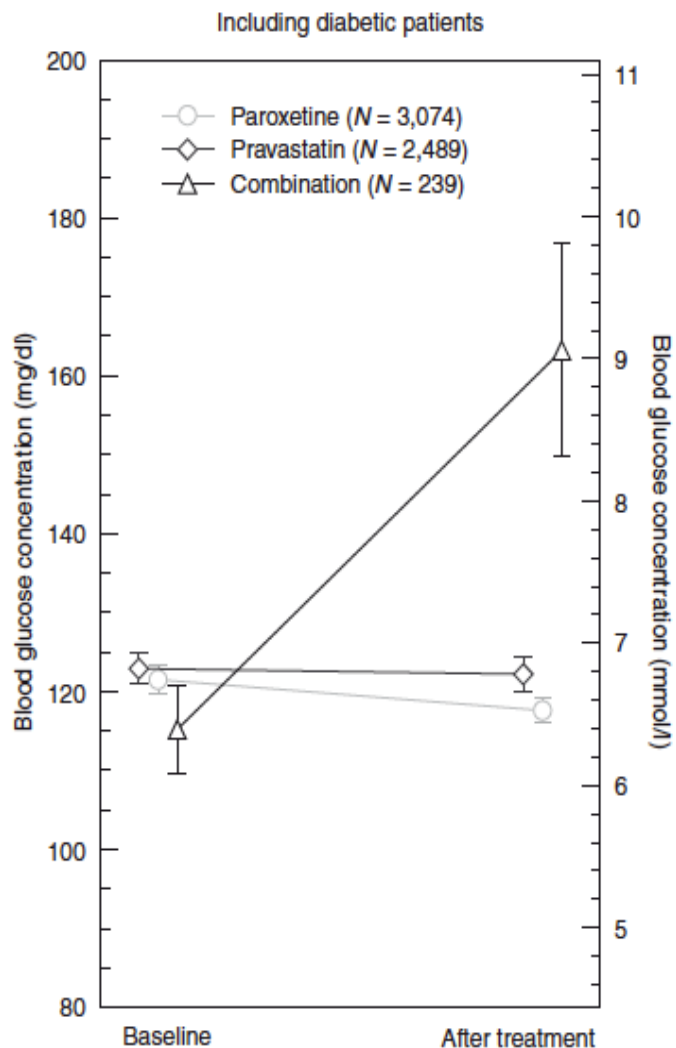
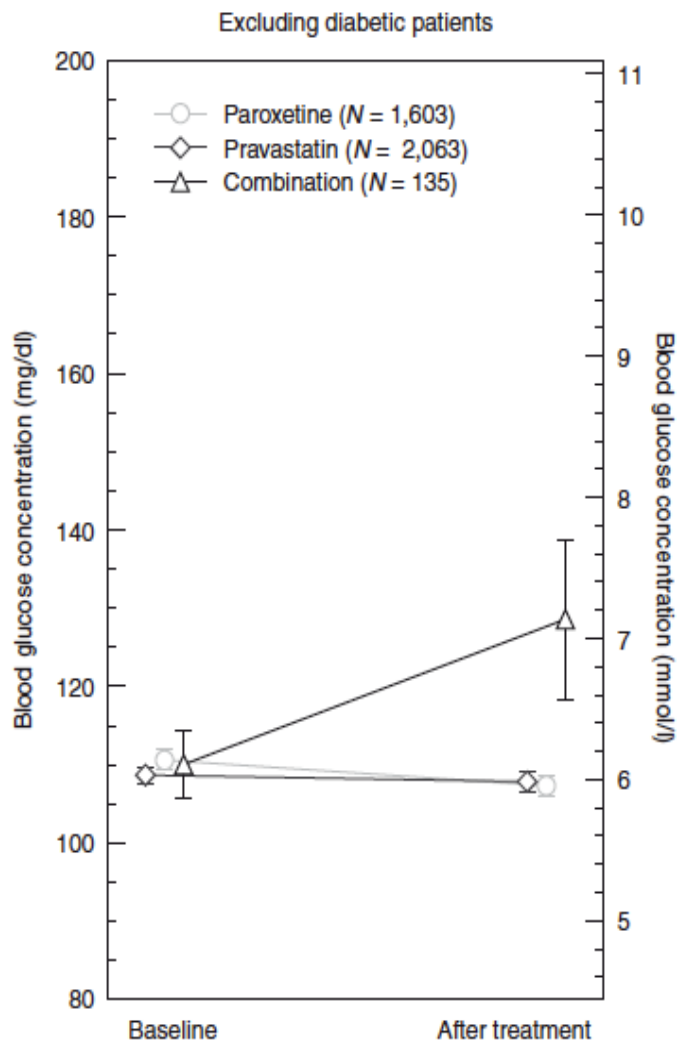
<sup>5</sup>Department of Medicine, Vanderbilt University, Nashville, Tennessee, USA

<sup>6</sup>Laboratory of Computer Science, Massachusetts General Hospital, Boston, Massachusetts, USA

<sup>7</sup>Harvard Medical School, Boston, Massachusetts, USA

<sup>8</sup>Stanford Center for Clinical Informatics, Stanford University, Stanford, California, USA

- Data mining FDA's Adverse Event Reporting System (FAERS) pour les effets adverses sur la glycémie
- Detection d'un signal avec l'association pravastatine et paroxetine.
- Dossiers patients de 3 hôpitaux
- Entrepôt i2b2
- Recherche l'augmentation de la glycémie sous paroxetine + pravastatine



Augmentation de 19 mg/dl (1.0 mmol/l)  
 Augmentation de 48 mg/dl (2.7 mmol/l) chez les diabétiques  
 Pas d'effet pour chaque médicament pris isolément  
 Pas d'effet connu pour les associations SSRI – statine

## The Georges Pompidou University Hospital Clinical Data Warehouse: A 8-years follow-up experience



Anne-Sophie Jannot<sup>a,b,c,\*</sup>, Eric Zapletal<sup>c</sup>, Paul Avillach<sup>d</sup>, Marie-France Mamzer<sup>a,e</sup>,  
Anita Burgun<sup>a,b,c</sup>, Patrice Degoulet<sup>a,b,c</sup>

- 74 projets de recherche approuvés par le Comité d’Ethique ( Institutional Review Board) entre Janvier 2011 et Décembre 2015.
- 25 nouveaux projets in 2017
- 54% épidémiologie clinique
- 41% health service research
- 5% support à la recherche clinique
- 2 ANRs (pharmacogénétique, coeliaque), 2 PREPS et 4 projets soumis
- Nouveaux projets collaboratifs en radiothérapie et radiomique
- 1 EU Flag-ERA : ITfoC
- 1 projet AP-HP sur la transition dans les maladies chroniques

- Planche, V., et al. (2014). Etiologies and diagnostic workup of extreme macrocytosis defined by an erythrocyte mean corpuscular volume over 130° fL: A study of 109 patients. *American journal of hematology*, 89(6), 665–666.
- Sahut, D. I., et al. (2012). Risk factors and clinical outcome of unsuspected pulmonary embolism in cancer patients: a case-control study. *Journal of thrombosis and haemostasis: JTH*, 10(10), 2032–2038.
- Neuraz, A., et al. (2013). **Phenome-wide association studies on a quantitative trait: application to TPMT enzyme activity and thiopurine therapy in pharmacogenomics.** *PLoS Comput Biol*, 9(12), e1003405.
- Girardeau, Y., et al. (2015). Detection of Drug–Drug Interactions Inducing Acute Kidney Injury by Electronic Health Records Mining. *Drug safety*, 38(9), 799–809.
- Pallet, N., et al. (2014). Urinary retinol binding protein is a marker of the extent of interstitial kidney fibrosis. *PloS one*, 9(1), e84708.
- Pham, A. D., et al. (2014). Natural language processing of radiology reports for the detection of thromboembolic diseases and clinically relevant incidental findings. *BMC bioinformatics*, 15(1), 1.
- Pettenati, C., et al. (2016). Prostate Cancer Characteristics and Outcome in Renal Transplant Recipients: Results from a Contemporary Monocentric Study. *Clinical transplantation*.
- Escudié, J. B., et al. (2015). Reviewing 741 patients records in two hours with FASTVISU. In *AMIA Annual Symposium Proceedings (Vol. 2015, p. 553)*. American Medical Informatics Association.
- Deborde, T., et al. (2016). [OP. 2A. 07] Trends in patients attending a tertiary hypertension unit: patient’s characteristics and treatment strategies strongly evolved over the last 15 years. *Journal of Hypertension*, 34, e17.
- Cheminet, G., et al. (2016). Causes associées aux hypomagnésiémies chez 5000 patients hospitalisés: étude cas-témoins. *La Revue de Médecine Interne*, 37, A105.
- Faucon, A. L., et al. (2016). Infarctus rénal: analyse rétrospective des causes d’une série de 186 cas sur 15 ans. *Néphrologie & Thérapeutique*, 12(5), 356.
- Jannot AS, Burgun A, Thervet E, Pallet N. **The Diagnosis-Wide Landscape of Hospital-Acquired AKI.** *Clin J Am Soc Nephrol*. 2017 May 11. pii: CJN.10981016. doi: 10.2215/CJN.10981016

**Deborde et al. Sex differences in antihypertensive treatment in France among 17 856 patients in a tertiary hypertension unit. J Hypertens. 2018 Jan 4**

- Recommandations nationales et internationales pour le traitement de l'hypertension.
- Centre de référence HTA
- Quelles sont les pratiques?
  
- 17856 patients à la 1<sup>ère</sup> consultation
- Les femmes ont plus souvent que les hommes des prescriptions de diurétiques de l'anse ou thiazidiques, des antagonistes du récepteur de l'aldostérone et des bêta-bloquants
- Sans bénéfice sur le contrôle de la tension (après ajustement)

# Reviewing Translational Research Platforms

Briefings in Bioinformatics Advance Access published March 7, 2014

BRIEFINGS IN BIOINFORMATICS. page 1 of 11

doi:10.1093/bib/bbu006

## **Translational research platforms integrating clinical and omics data: a review of publicly available solutions**

*Vincent Canuel\*, Bastien Rance\*, Paul Avillach, Patrice Degoulet and Anita Burgun*

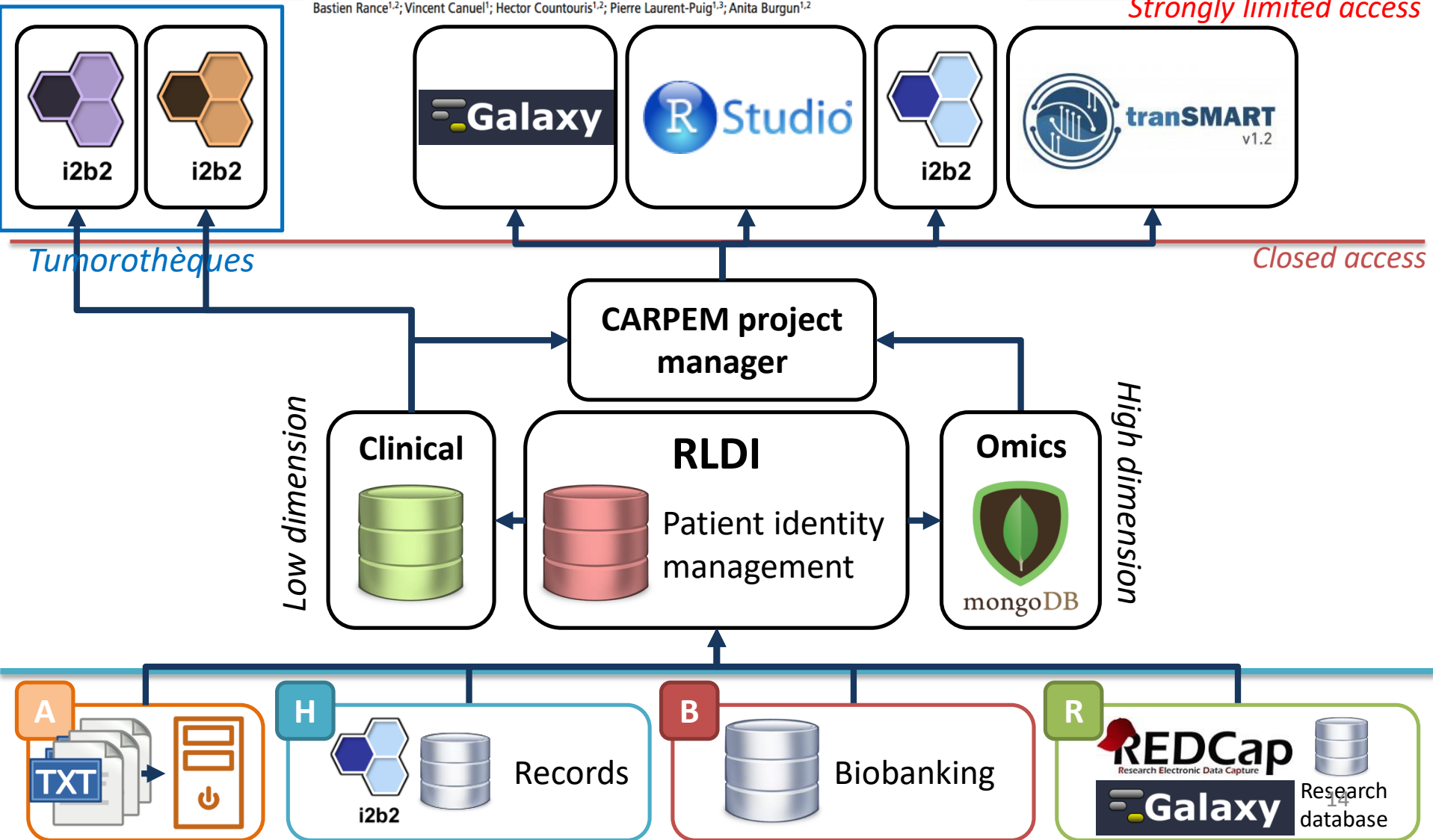
Submitted: 11th November 2013; Received (in revised form): 3rd February 2014

BRISK, caTRIP, **cBio Cancer Portal**,  
**G-DOC**, iCOD, iDASH, **tranSMART**

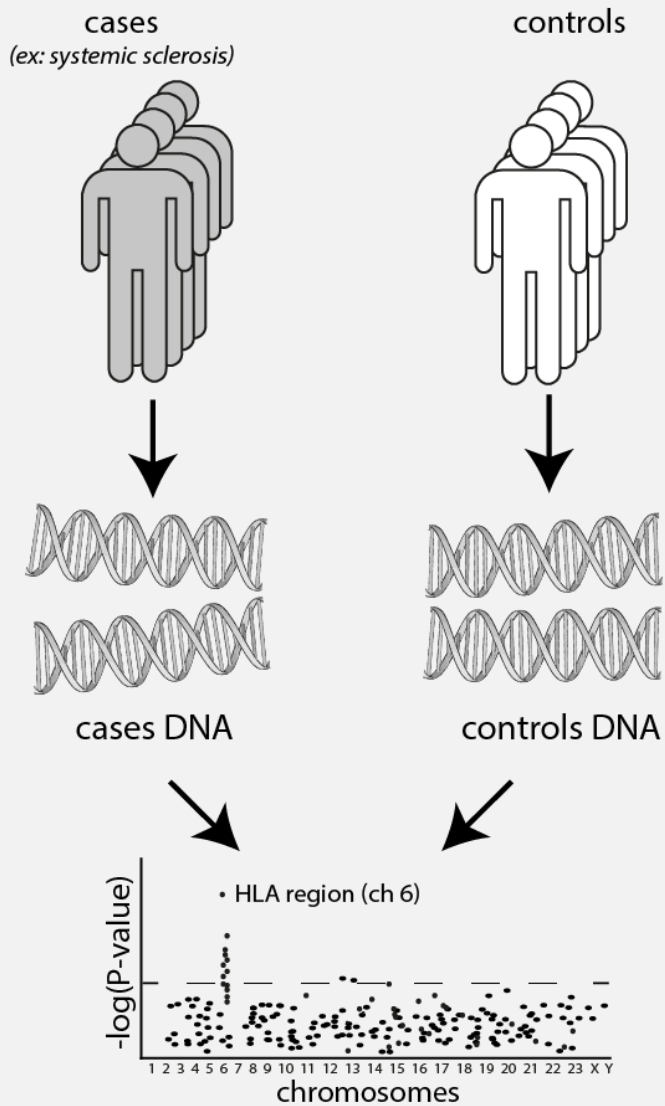
# Integrating Heterogeneous Bio-medical Data for Cancer Research: the CARPEM infrastructure

Bastien Rance<sup>1,2</sup>; Vincent Canuel<sup>1</sup>; Hector Countouris<sup>1,2</sup>; Pierre Laurent-Puig<sup>1,3</sup>; Anita Burgun<sup>1,2</sup>

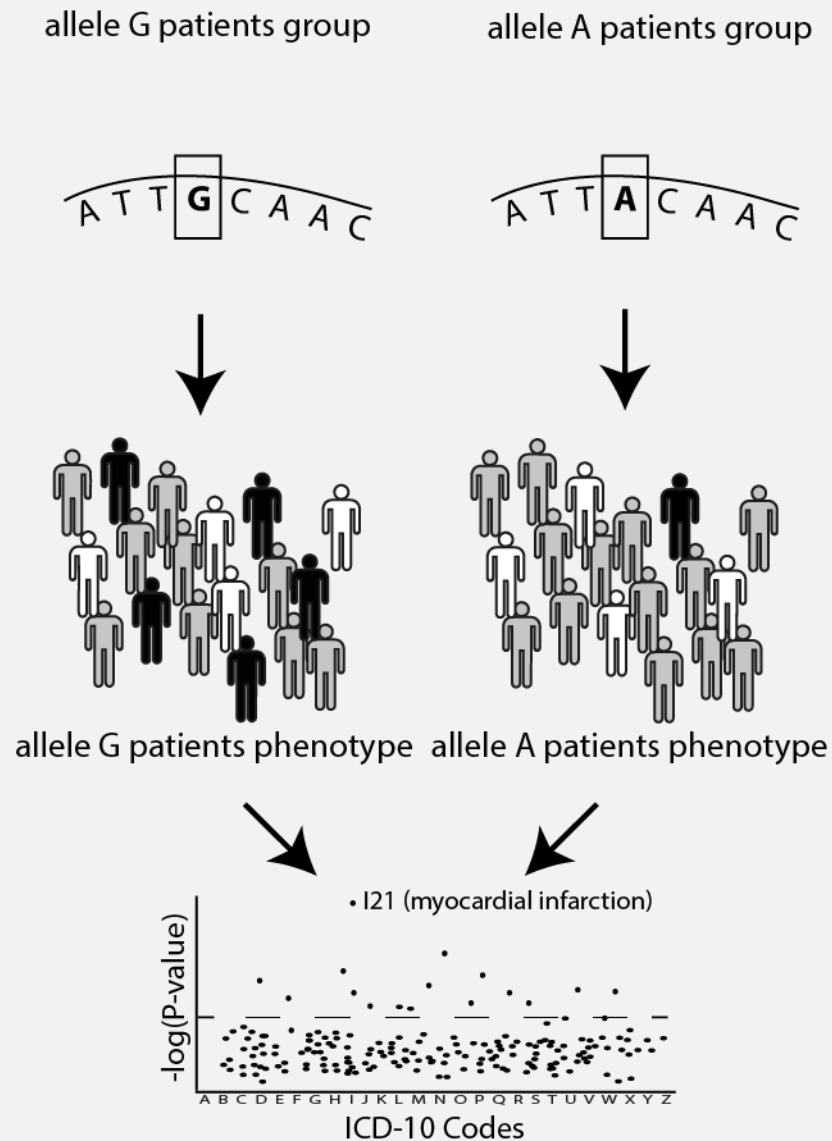
*Strongly limited access*



## Genome Wide Association Study (GWAS)



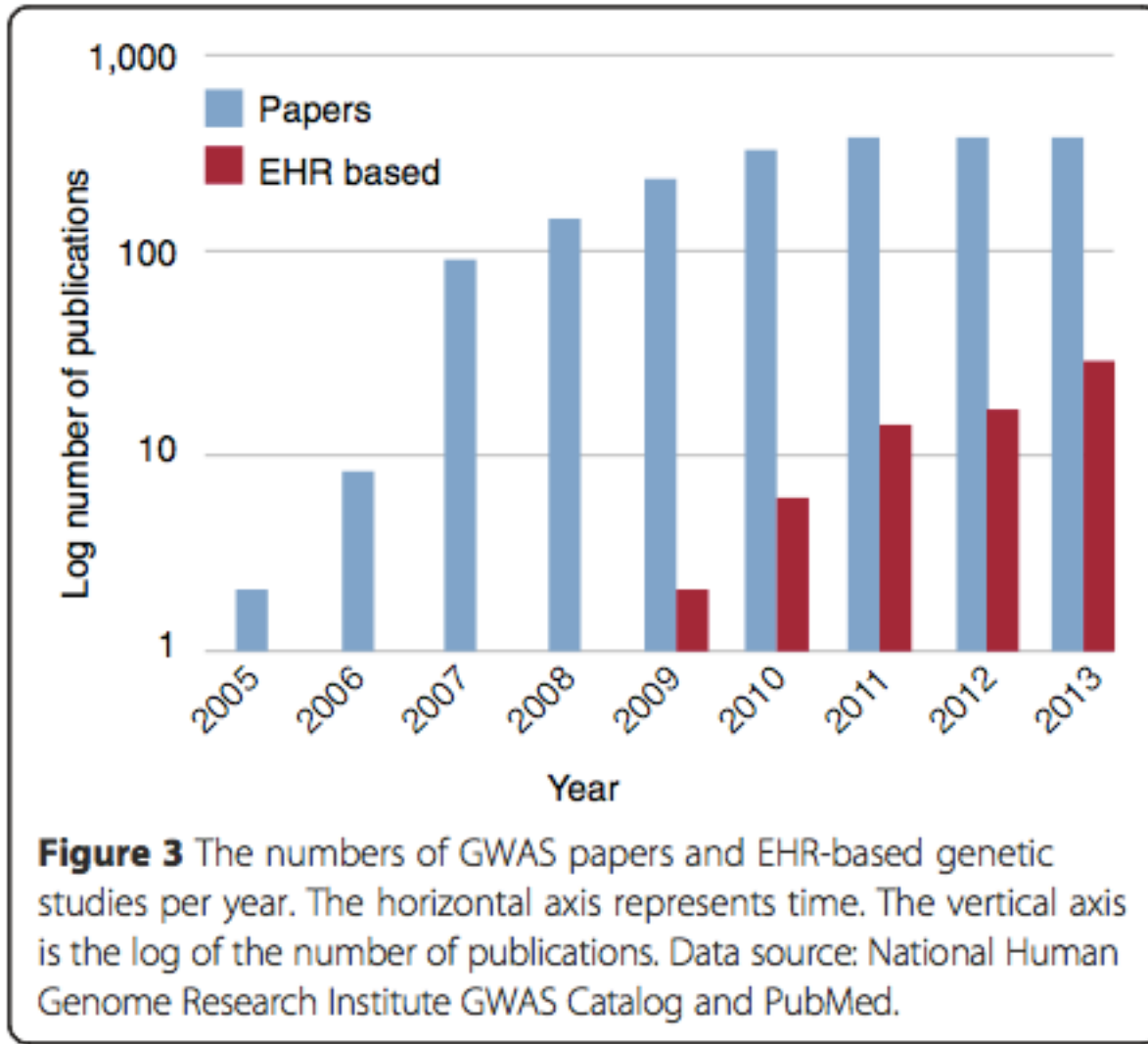
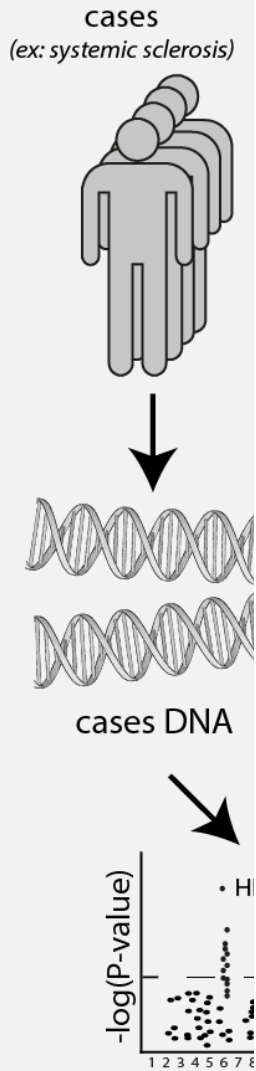
## Phenome Wide Association Study (PheWAS)





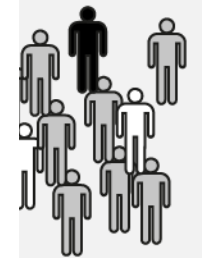
# Genome Wide Association Study (GWAS)

# Phenome Wide Association Study (PheWAS)



Wei WQ et al Genome Medicine, 2015

patients group



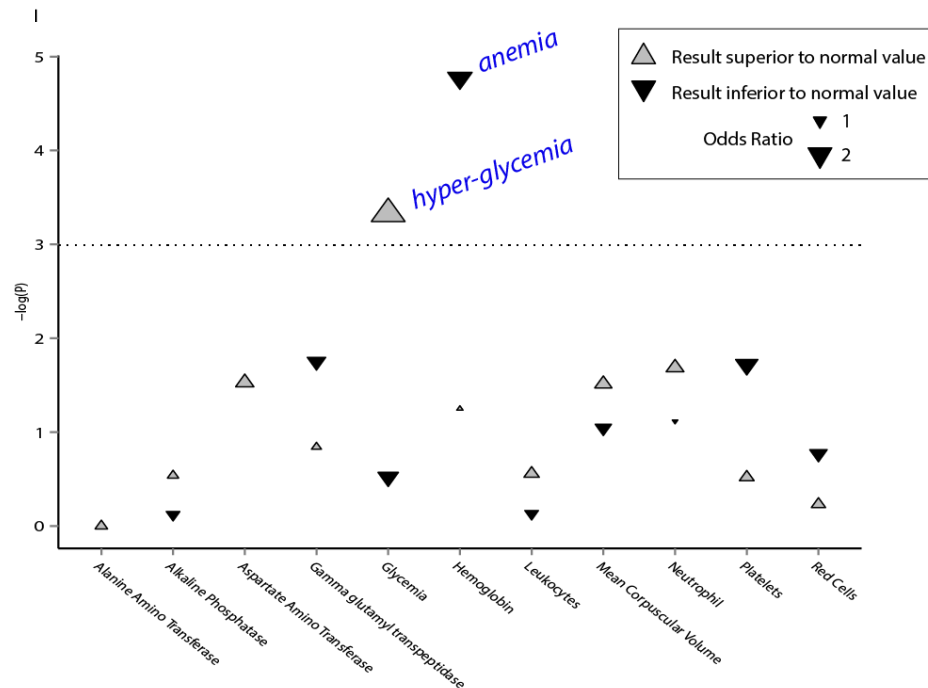
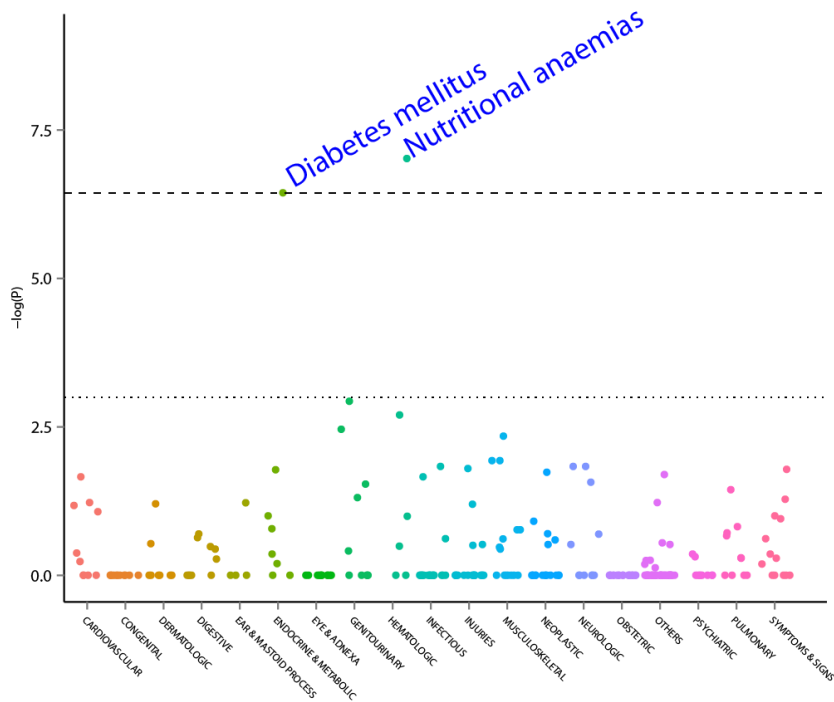
ion)



A B C D E F G H I J K L M N O P Q R S T U V W X Y Z  
ICD-10 Codes

# Phenome-Wide Association Studies on a Quantitative Trait: Application to TPMT Enzyme Activity and Thiopurine Therapy in Pharmacogenomics

Antoine Neuraz<sup>1,2</sup>, Laurent Chouchana<sup>3</sup>, Georgia Malamut<sup>4</sup>, Christine Le Beller<sup>5</sup>, Denis Roche<sup>6</sup>, Philippe Beaune<sup>3,6</sup>, Patrice Degoulet<sup>1,2</sup>, Anita Burgun<sup>1,2</sup>, Marie-Anne Lorient<sup>3,6</sup>, Paul Avillach<sup>1,2\*</sup>



Phenotype

Thiopurine

Low activity

10 % dose

Intermediate activity

30 – 70 % dose

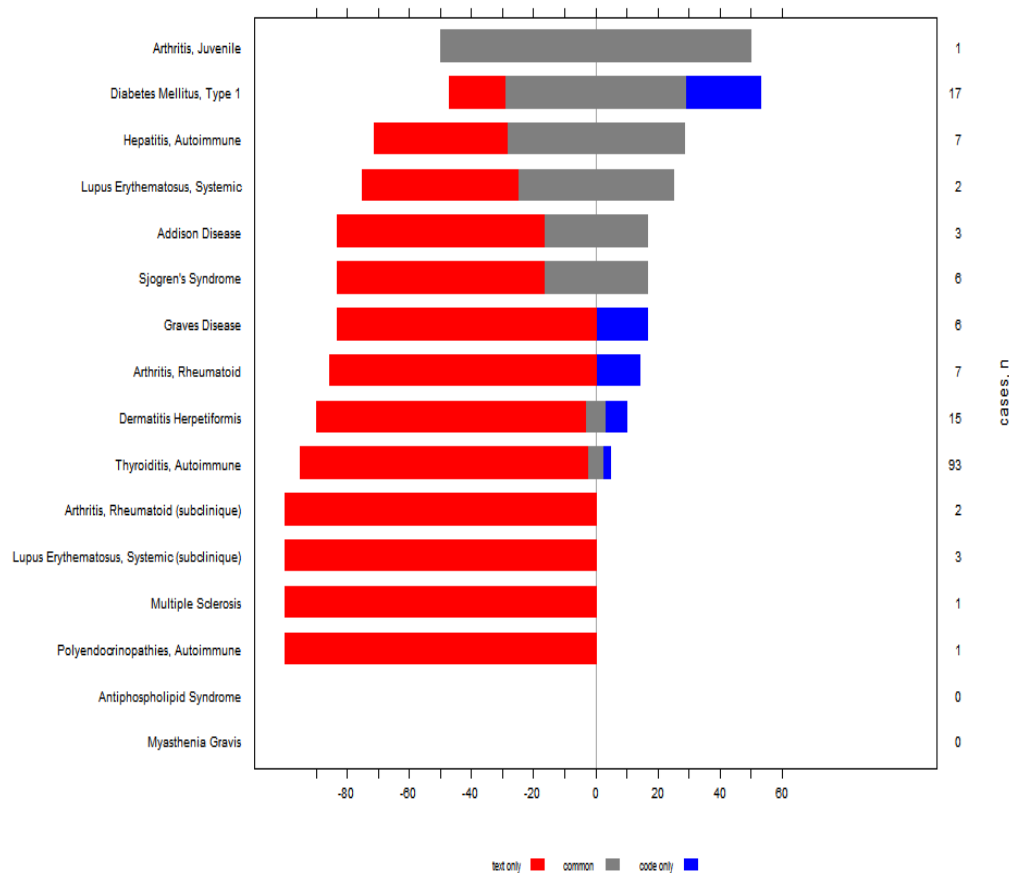
Normal activity

100 % dose

Very High activity

> 100 % Dose ?

# Case study: Autoimmune comorbidities of the Celiac Disease



81%

de l'information présente  
dans les textes uniquement  
(et absente des données  
structurées)

*Escudié JB, Rance B, Malamut G, Khater S, Burgun A, Cellier C, Jannot AS. A novel data-driven workflow combining literature and electronic health records to estimate comorbidities burden for a specific disease: a case study on autoimmune comorbidities in patients with celiac disease. BMC Med Inform Decis Mak. 2017 Sep 29;17(1):140.*

*Escudié JB, Jannot AS, Zapletal E, Cohen S, Malamut G, Burgun A, Rance B. Reviewing 741 patients records in two hours with FASTVISU. AMIA Annu Symp Proc.2015 Nov 5;2015:553-9.*

# Phenotyping with Dr Warehouse (Garcelon et al. JBI 3017)

Dr WareHouse ©Imagine  
Entrepôt de données

Accueil | Moteur de recherche | Mes requêtes | Mes cohortes | Outils | Contact | Patient Nom / ICPP | Admin |

## Rechercher des patients

Sur tout l'entrepôt

mecp2 274/274

**MECP2**

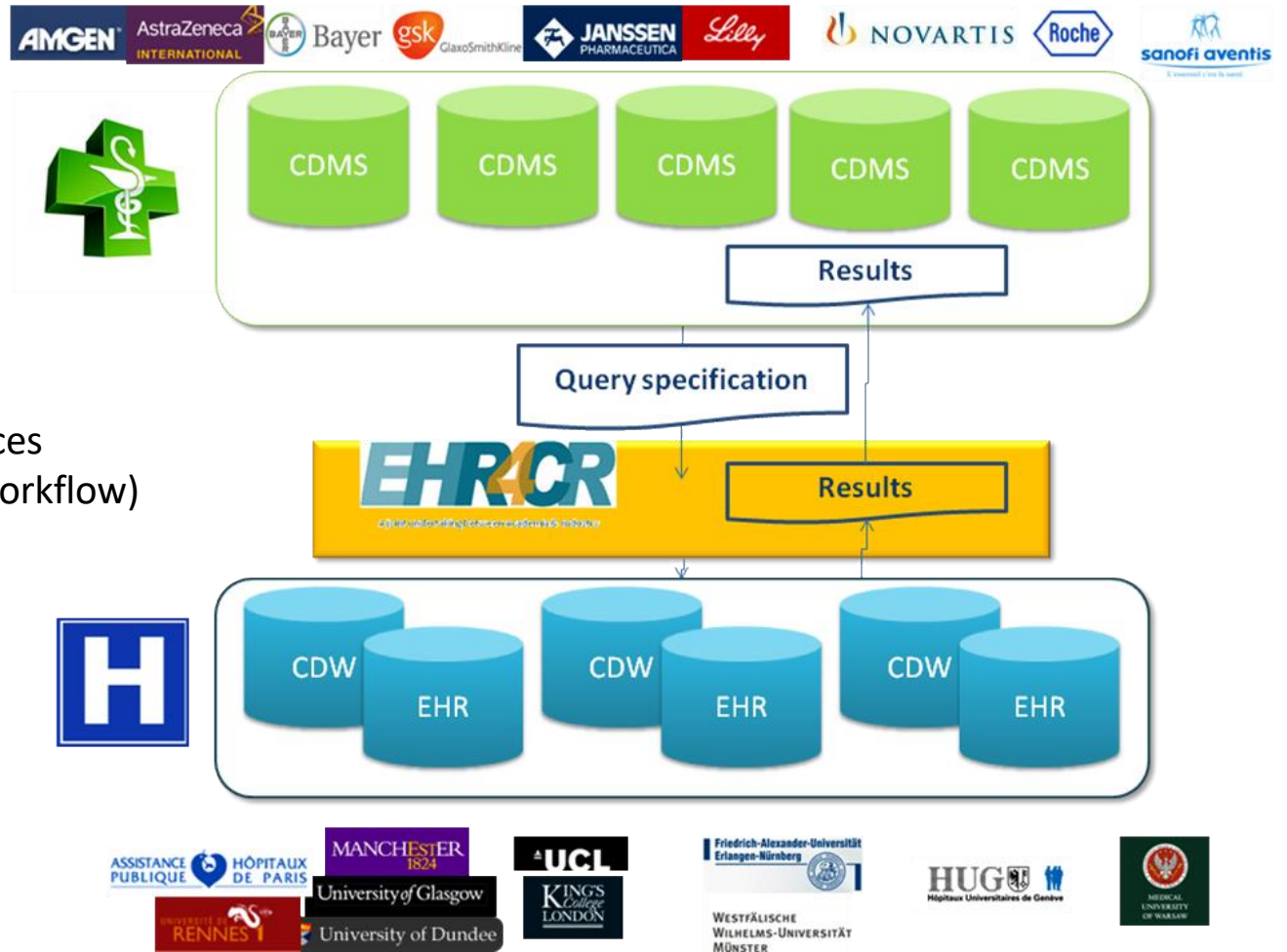
- + Ajouter un filtre Full text
- + Ajouter un filtre structuré
- + Filtre patient

**LANCER LA RECHERCHE** ?

Sur tout l'hôpital :  
274 Patients  
603 Documents  
Sur les documents trouvés - Sur tous les documents des patients trouvés  
Profondeur : 10

endormissement  
REFLUX GASTRO-OESOPHAGIEN  
encephalopathie  
troubles du sommeil  
Syndrome de Rett  
Crise  
EPILEPSIE  
Scoliose  
Agitation  
Surdité  
Microcephalie  
crises toniques  
spasmes  
Osteoporose  
baisse cdg  
RETARD DE DEVELOPPEMENT  
Bruxisme  
Syndrome pyramidal  
Retard psychomoteur  
endormissement  
REFLUX GASTRO-OESOPHAGIEN  
Cyphose  
troubles du sommeil  
infection  
Constipation  
Atrophie  
hypertonie  
Fievre  
Incontinence  
Hypothyroïdie

# Support / complément à la recherche clinique



Protocol feasibility  
 Patient recruitment services  
 Clinical trial execution (workflow)  
 Pharmacovigilance

Search by Subject | **Navigate Terms** | Across Trials

Comparison | **Advanced Workflow** | Results/Analysis | Grid View | Data Export | Export Jobs

Analysis ▾

**transSMART**

# Kaplan-Meier estimator

— 0 ≤ CATEGORY < 20  
— 20 ≤ CATEGORY ≤ 76

**p = 0,04**

Overall survival (%)

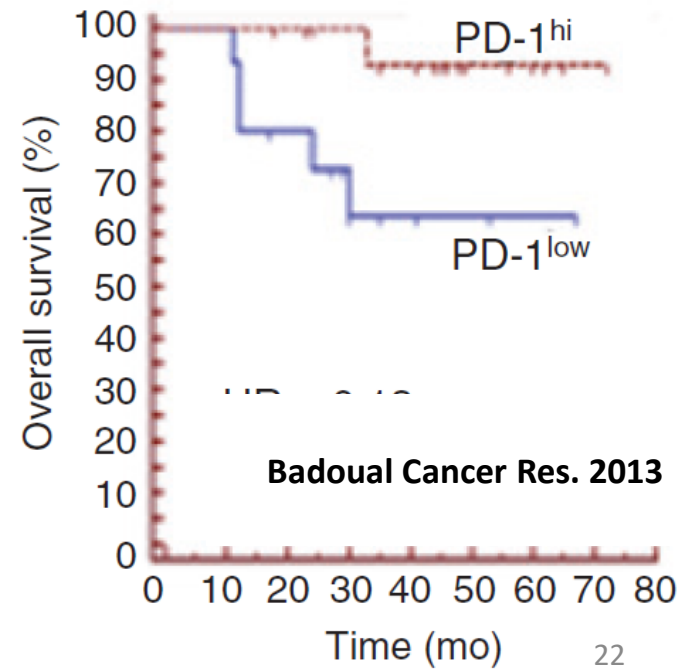
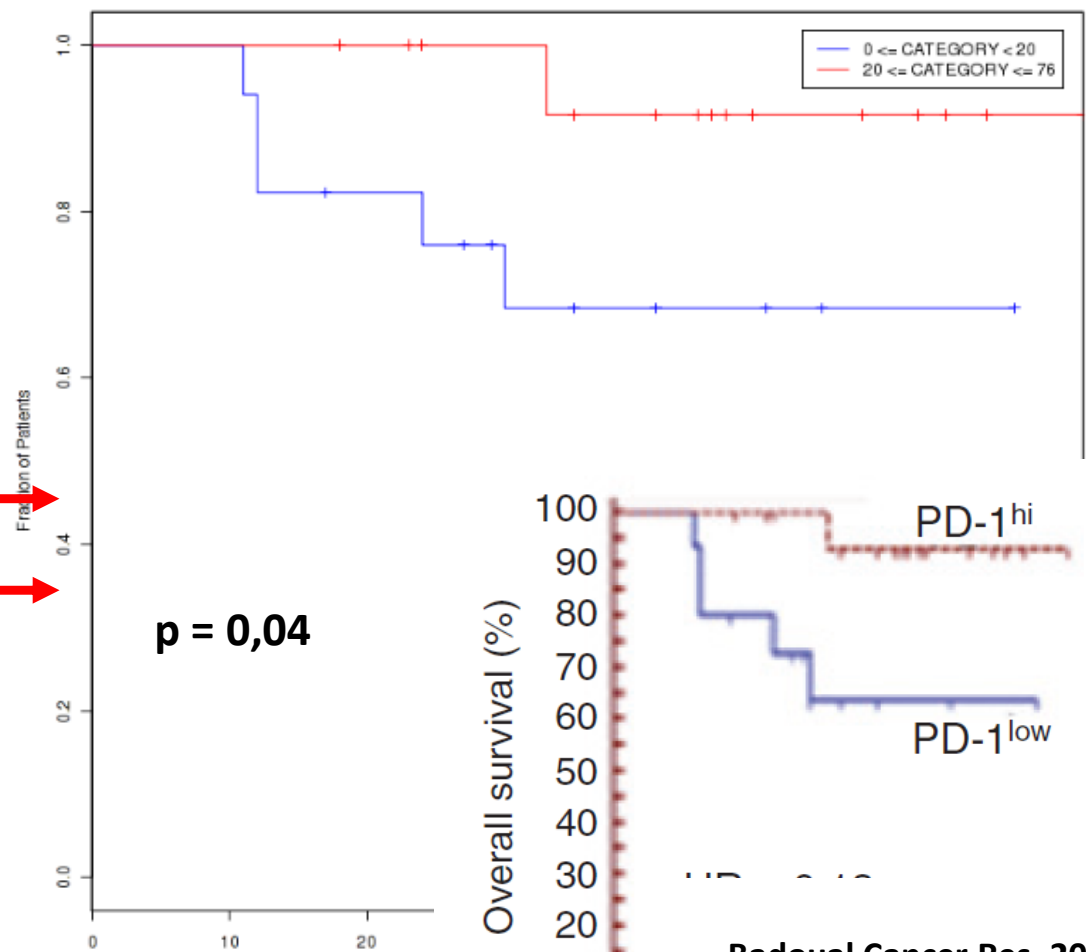
PD-1<sup>hi</sup>  
PD-1<sup>low</sup>

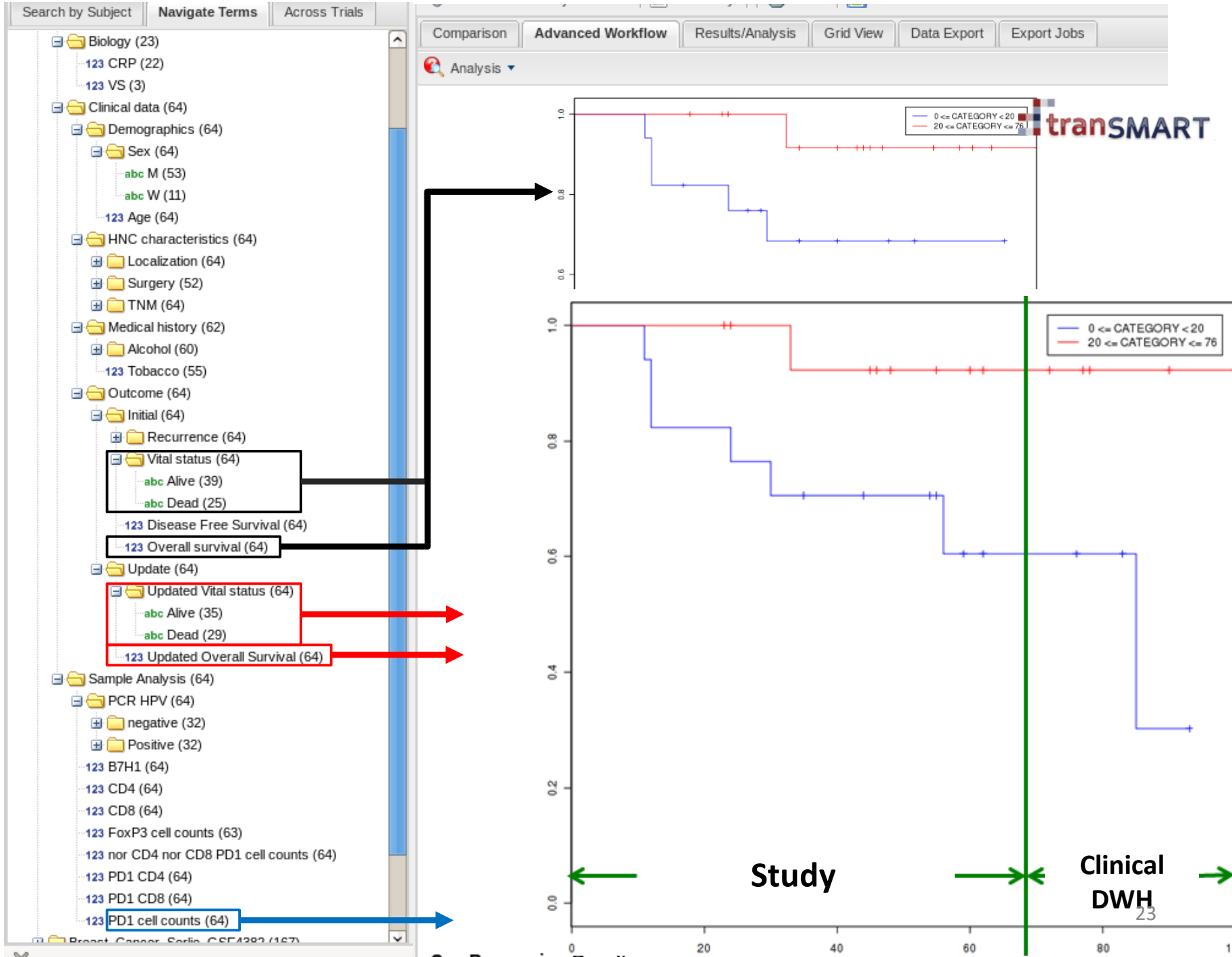
Badoual Cancer Res. 2013

Time (mo)

22

- Biology (23)
    - 123 CRP (22)
    - 123 VS (3)
  - Clinical data (64)
    - Demographics (64)
      - Sex (64)
        - abc M (53)
        - abc W (11)
      - 123 Age (64)
    - HNC characteristics (64)
      - Localization (64)
      - Surgery (52)
      - TNM (64)
    - Medical history (62)
      - Alcohol (60)
      - 123 Tobacco (55)
    - Outcome (64)
      - Initial (64)
        - Recurrence (64)
        - Vital status (64)
          - abc Alive (39)
          - abc Dead (25)
        - 123 Disease Free Survival (64)
        - 123 Overall survival (64)
      - Update (64)
        - Updated Vital status (64)
          - abc Alive (35)
          - abc Dead (29)
        - 123 Updated Overall Survival (64)
    - Sample Analysis (64)
      - PCR HPV (64)
        - negative (32)
        - Positive (32)
      - 123 B7H1 (64)
      - 123 CD4 (64)
      - 123 CD8 (64)
      - 123 FoxP3 cell counts (63)
      - 123 nor CD4 nor CD8 PD1 cell counts (64)
      - 123 PD1 CD4 (64)
      - 123 PD1 CD8 (64)
      - 123 PD1 cell counts (64)

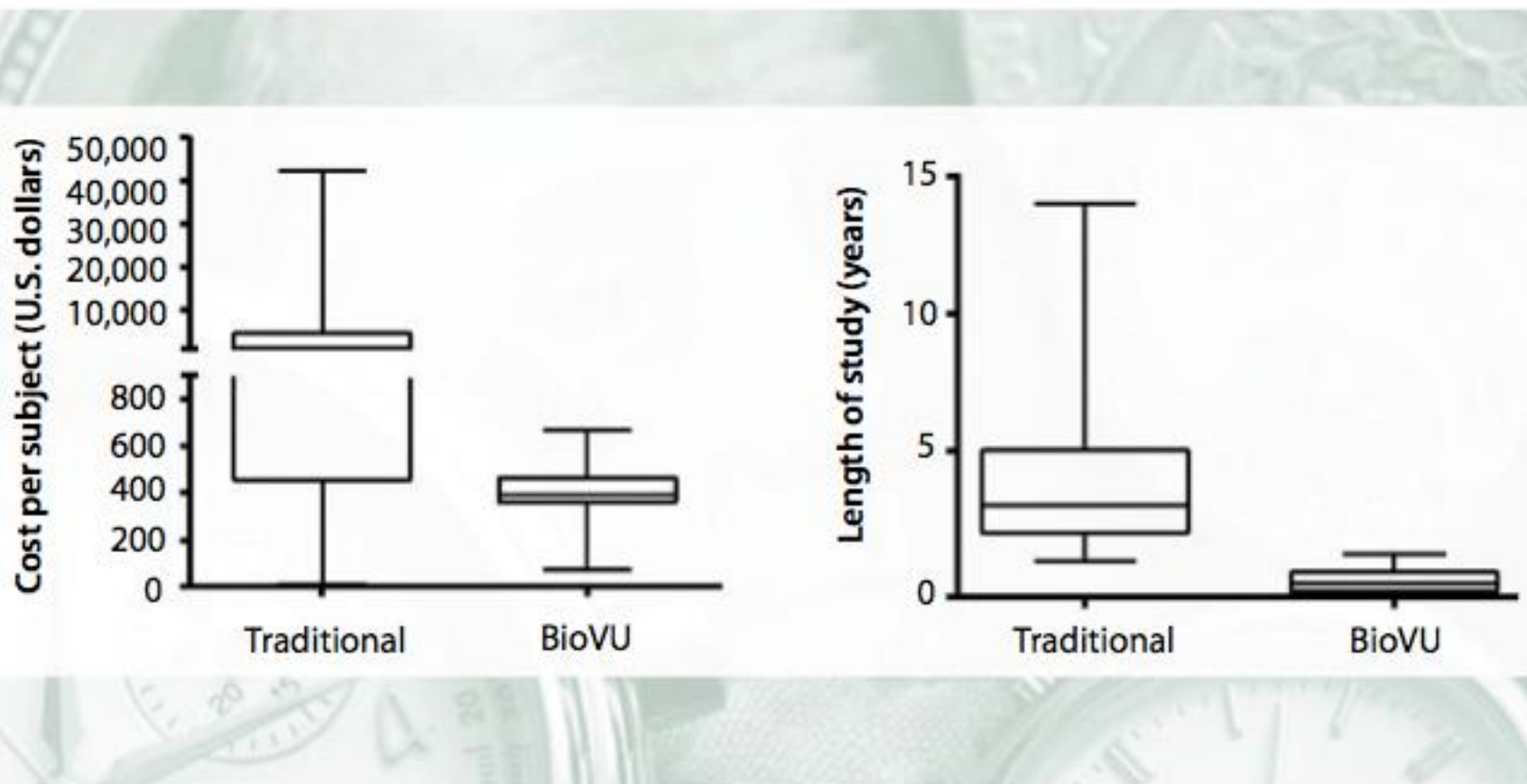






## Biobanks and Electronic Medical Records: Enabling Cost-Effective Research

Erica Bowton,<sup>1\*</sup> Julie R. Field,<sup>1</sup> Sunny Wang,<sup>1</sup> Jonathan S. Schildcrout,<sup>2</sup> Sara L. Van Driest,<sup>3</sup> Jessica T. Delaney,<sup>4</sup> James Cowan,<sup>1</sup> Peter Weeke,<sup>4</sup> Jonathan D. Mosley,<sup>4</sup> Quinn S. Wells,<sup>4</sup> Jason H. Karnes,<sup>4</sup> Christian Shaffer,<sup>4</sup> Josh F. Peterson,<sup>4,5</sup> Joshua C. Denny,<sup>4,5</sup> Dan M. Roden,<sup>4,6</sup> Jill M. Pulley<sup>7</sup>



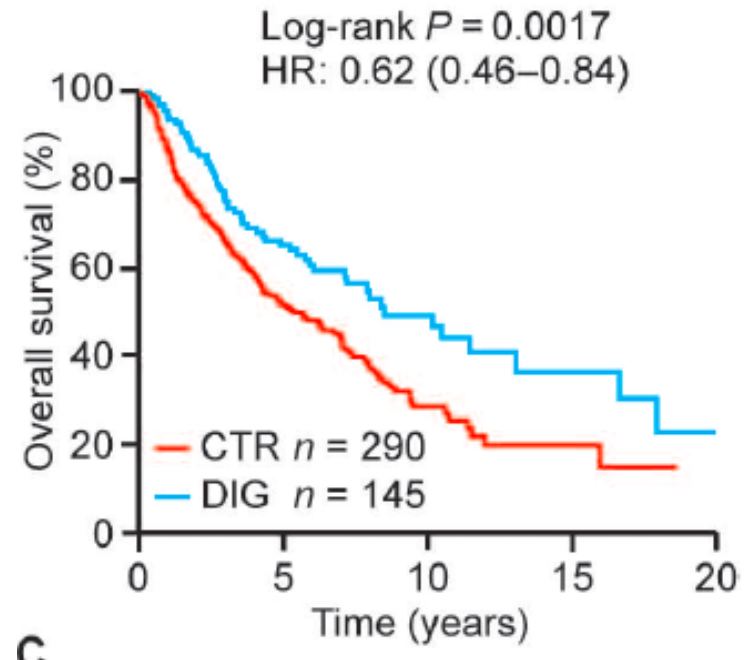
**Fig. 1. Time is money.** Comparison of traditional NIH-funded pharmacogenomic studies versus EMR/biobank studies (BioVU). (**Left**) Median cost of study per subject. (**Right**) Median length of study in years.

## CANCER

# Cardiac Glycosides Exert Anticancer Effects by Inducing Immunogenic Cell Death

Laurie Menger,<sup>1,2,3</sup> Erika Vacchelli,<sup>1,2,3</sup> Sandy Adjemian,<sup>1,2,3</sup> Isabelle Martins,<sup>1,2,3</sup> Yuting Ma,<sup>1,2,3</sup> Shensi Shen,<sup>1,2,3</sup> Takahiro Yamazaki,<sup>2,3,4</sup> Abdul Qader Sukkurwala,<sup>1,2,3</sup> Mickaël Michaud,<sup>1,2,3</sup> Grégoire Mignot,<sup>5,6</sup> Frederic Schlemmer,<sup>1,2,3</sup> Eric Sulpice,<sup>7</sup> Clara Locher,<sup>2,3,4</sup> Xavier Gidrol,<sup>7</sup> François Ghiringhelli,<sup>5,6</sup> Nazanine Modjtahedi,<sup>1,2,3</sup> Lorenzo Galluzzi,<sup>2,8</sup> Fabrice André,<sup>2,9</sup> Laurence Zitvogel,<sup>2,3,4</sup> Oliver Kepp,<sup>1,2,3,\*†</sup> Guido Kroemer<sup>1,8,10,11,12,\*†</sup>

- Cell -> mice -> retrospective data
- text-based research algorithm to identify all carcinoma patients who, between 1981 and 2009, received digitalin during conventional carcinoma therapies
- compared the overall survival of 145 patients treated with CGs with that of 290 patients who did not receive CGs.



Martin K, et al. . Differences between clinical trials and postmarketing use.  
Br J Clin Pharmacol.

- Essais cliniques : populations sélectionnées et conditions standardisées
- Vie réelle : comorbidités et comportements des patients
- 3 populations 3 molécules : tacrine, simvastatin et celecoxib
  - Cible : critères d'inclusion/exclusion
  - Effets adverses
  - Toute la population

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**Table 2**

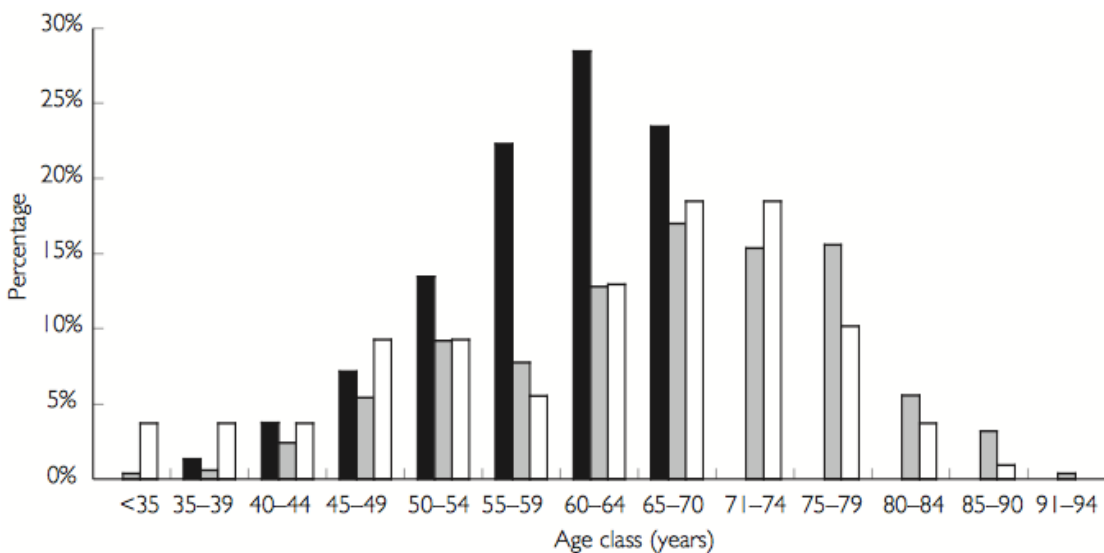
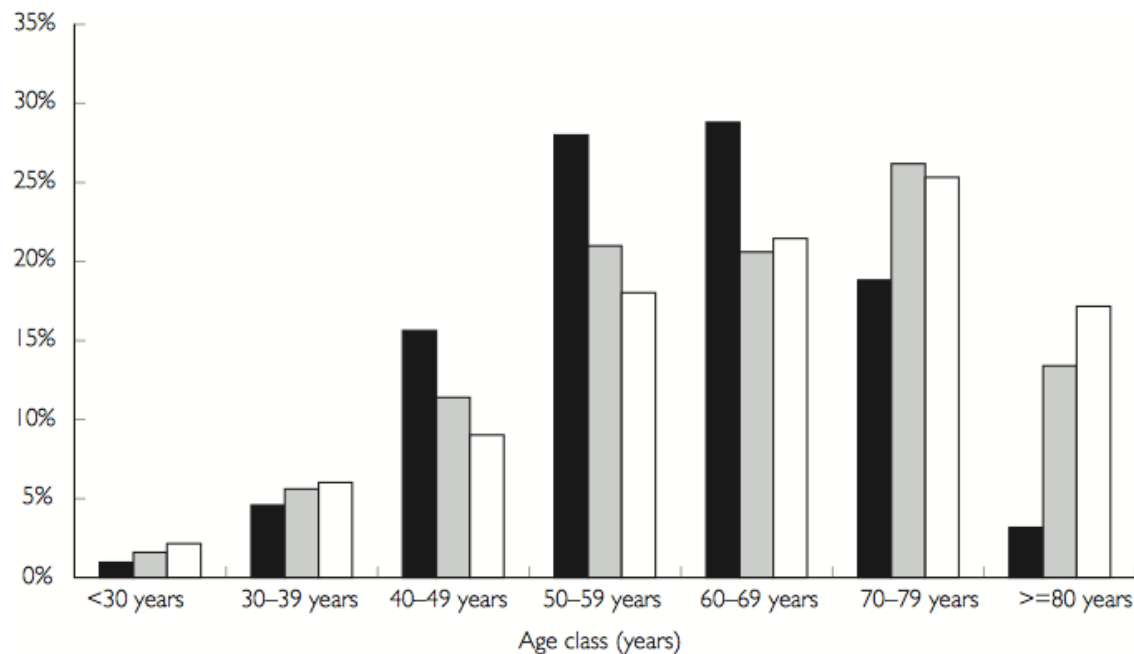
Comparison of male : female ratios between the three populations

	Target	Population Joint	Injured
Tacrine	0.9	0.6	0.5
Simvastatin	4.6	1	1.5
Celecoxib	0.4	0.5	0.5

**Figure 1**

Age distribution for the three populations for celecoxib. Target population (■), joint population (■) and injured population (□)

Martin K, et al. . Differences between clinical trials and postmarketing use. *Br J Clin Pharmacol* tacrine, simvastatin et celecoxib



**Figure 2**

Age distribution for the three populations for simvastatin. Target population (■), joint population (■) and injured population (□)

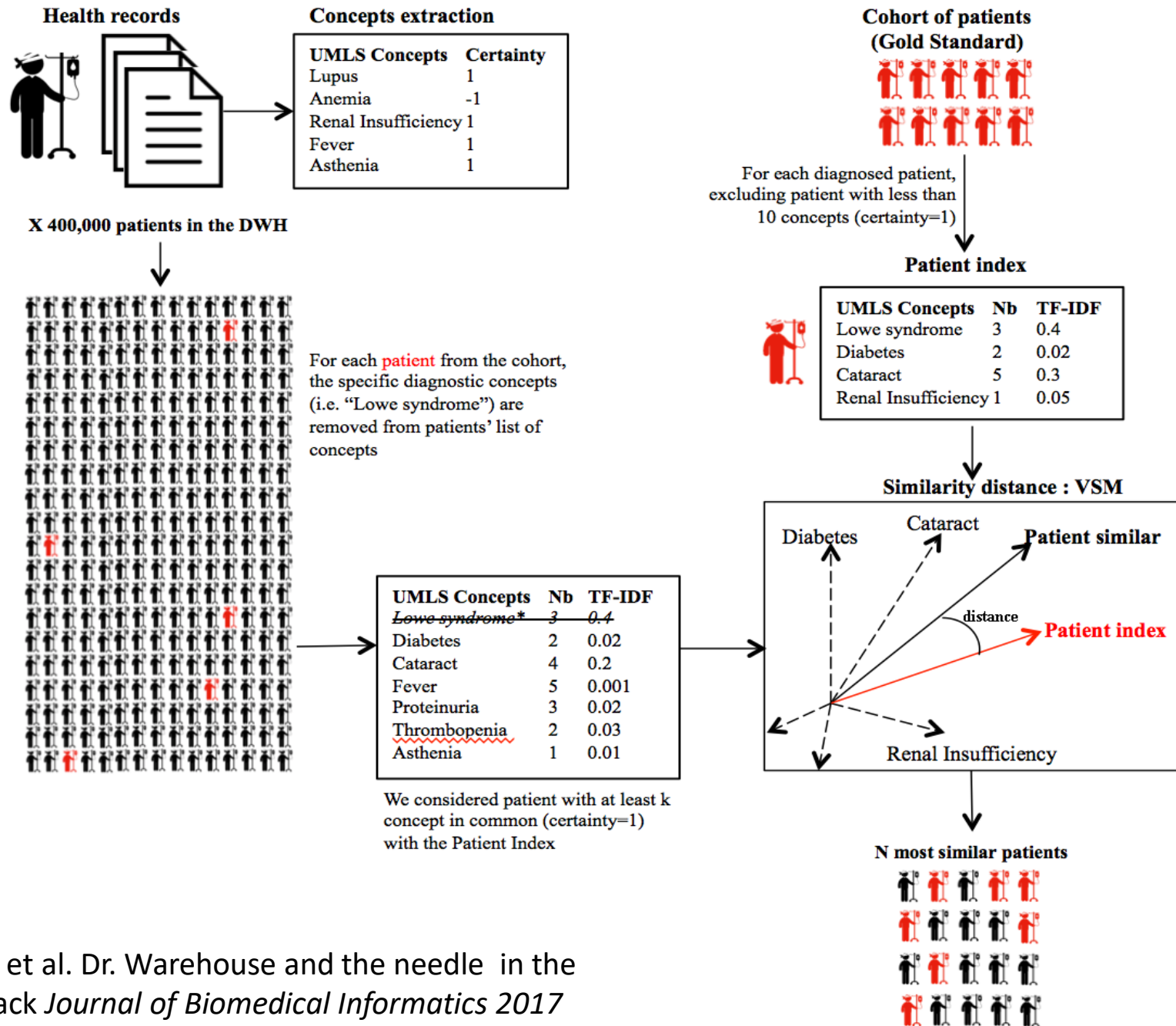
# Aide à la décision dans le cadre du soin




**Evidence-Based Medicine in the EMR Era**  
 J.Frankovich, C.A. Longhurst, S M. Sutherland.  
 Stanford, publication NEJM 9 Nov 2011

**Results of Electronic Search of Patient Medical Records (for a Cohort of 98 Pediatric Patients with Lupus) Focused on Risk Factors for Thrombosis Relevant to Our 13-Year-Old Patient with Systemic Lupus Erythematosus.\***

Outcome or Risk Factor	Keywords Used to Conduct Expedited Electronic Search	Prevalence of Thrombosis <i>no./total no (%)</i>	Relative Risk (95% CI)
Outcome — thrombosis	“Thrombus,” “Thrombosis,” “Blood clot”	10/98 (10)	Not applicable
Thrombosis risk factor			
Heavy proteinuria (>2.5 g per deciliter)			
Present at any time	“Nephrosis,” “Nephrotic,” “Proteinuria”	8/36 (22)	7.8 (1.7–50)
Present >60 days	“Urine protein”	7/23 (30)	14.7 (3.3–96)
Pancreatitis	“Pancreatitis,” “Lipase”	5/8 (63)	11.8 (3.8–27)
Antiphospholipid antibodies	“Aspirin”	6/51 (12)	1.0 (0.3–3.7)

**« we made the decision on the basis of the best data available »  
 « in the light of experience as guided by intelligence. »**



Patient	Similarité	Concepts communs	Concepts patient
Patient 1 (statut cohorte Syndrome de lowe : 1 )  <a href="#">Afficher le comparateur</a>	54	Acidose, Albuminurie, CATARACTE CONGENITALE, Crise, Deshydratation, Retard de croissance, Syndrome de Fanconi, Hemolyse, saignement, Hypercalciurie, nephropathie, Cyphose, Lithiase, HYPOTONIE, Nephrocalcinose, Proteinurie, Rachitisme, Scoliose, renal, Agitation, CRISTALLURIE, Tolerance, Selles liquides, Crises convulsives, Tension, Petite taille, Retard psychomoteur, Microalbuminurie, anomalies du transport des acides amines, agitation, Demineralisation, CATARACTE, Insuffisance renale	Glaucome, gastroenterite, Handicap, EPILEPSIE, rougeur oculaire, Acidose metabolique, Diarrhee, Diarrhee chronique, temps, Lithiases renales, ocr1, Paleur, Pale, <b>Syndrome de Lowe</b> , RETARD DE DEVELOPPEMENT, maladies metaboliques, Masse musculaire, Fievre, Hypothermie, Sursauts, Gammaglobulinemie, Glycosurie, ANEMIE MICROCYTAIRE, Hypertonie oculaire, glaucome, Epilepsie occipitale, Malaise, Hematurie, Polypnee, Hematurie macroscopique, Prise de poids, Bronchiolite, Photophobie, diarrhees, t3 normale, Syndrome nephrotique, alimentation, Selles molles, rhinite, Anorexie, vomissement, Troubles du transit, Eruption, Urgences, Fatigue, Constipation, Cetonurie, Sedation, Keratite, opacite corneenne, Hypoglycemie, lithiase renale, Crises generalisees, HYPOKINESIE, hypertension, Gaz
Patient 2 (statut cohorte Syndrome de lowe : 1 )  <a href="#">Afficher le comparateur</a>	51	Acidose, CATARACTE CONGENITALE, Kyste, Deshydratation, Retard de croissance, saignement, Hypercalciurie, allergie, nephropathie, Cyphose, Lithiase, HYPOTONIE, Nephrocalcinose, Proteinurie, renal, CRISTALLURIE, Tolerance, Tension, Amino-acidurie, Malformation, Petite taille, Masse, Microalbuminurie, anomalies du transport des acides amines, Demineralisation, CATARACTE	Prise de poids, Eruption, Retard staturο-ponderal, hypovolemie, Amblyopie, Gastrostomie, Peau seche, urate, Genu valgum, vomissement, ulceration, Diarrhee, <b>Syndrome de Lowe</b> , Infections, Perte de poids, Cytolyse hepatique, Syndrome nephrotique, lithiase renale, Constipation, ocr1, Urgences, Dysmorphie, Fente palatine, Masse musculaire, temps, retrognathisme, malnutrition, Nerveux, Stress, cryptorchidie bilaterale, reflexe nauseeux, Anorexie, Gastroenterite aigue, luxation congenitale de hanche, Glaucome, hypotonie neonatale, alimentation, Bouton, hypertrophie, Acidose metabolique, glaucome congenital
Patient 3 (statut cohorte Syndrome de lowe : 1 )  <a href="#">Afficher le comparateur</a>	51	vesicule, Crise, Deshydratation, Retard de croissance, pieds plats, saignement, hepatite b, Hypercalciurie, allergie, Lithiase, HYPOTONIE, Nephrocalcinose, osteomalacie, Proteinurie, Scoliose, thrombopenie, renal, Agitation, CRISTALLURIE, Tension, Dysmorphie faciale, Malformation, Retard psychomoteur, troubles de l'hemostase, anomalies du transport des acides amines, agitation, Demineralisation, CATARACTE	Gaz, marche ataxique, Douleurs abdominales, incontinence fecale, Sursauts, Chute, Inquietude, Tumefaction, Fecalome, Protides totaux, azotemie, Gastrite, RETARD DE DEVELOPPEMENT, Colique, depots calciques, Tartre, ocr1, Acidose metabolique, Alopecie, alimentation, Lordose, Puberte, <b>Syndrome de Lowe</b> , Gynecomastie, parodontopathies, Intervention chirurgicale, lithiase renale, Constipation chronique, EPILEPSIE, Carie, perte de sommeil, cataracte, Plaquettes normales, Cecite, distension, Hepatite C, Constipation, Douleur, Glaucome, crises d'epilepsie, retention d'urine, Varicelle, Gingivite, Stress, perte de connaissance, Observation, cecite des deux yeux, Constipation opiniatre, Douleurs abdominales, Anorexie, temps, glaucome



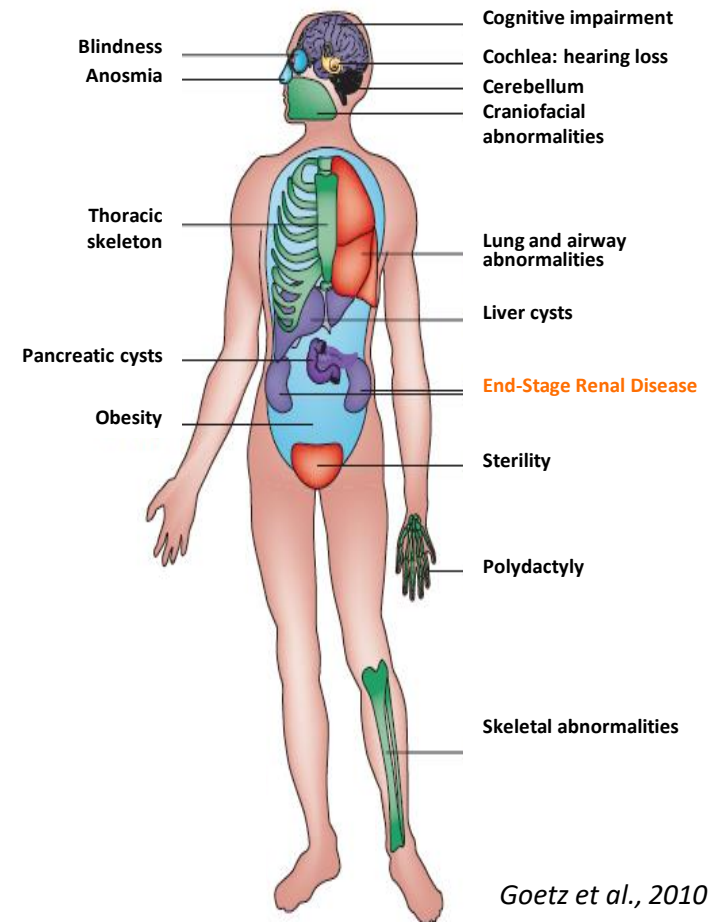
# Une redéfinition des maladies: l'exemple des ciliopathies

Source : **C'IL-LICO Consortium**

# The C'IL-LICO Project: Next-Generation Medicine For Renal Ciliopathies

C'IL-LICO is based on similarities between ciliopathy signaling pathways, offering an opportunity to identify common therapeutic option(s).

- › **Rare and severe genetic diseases** caused by **ciliary dysfunction**
- › **Individually rare, collectively frequent** : **1/2000 birth**
- › **End-stage renal disease (ESRD)**: main cause of morbidity and mortality
- › **25% ESRD patients are undiagnosed**: part of them could have a ciliopathy
- › **Common pathways in ciliopathies**: opportunity for **common treatment of different patients**

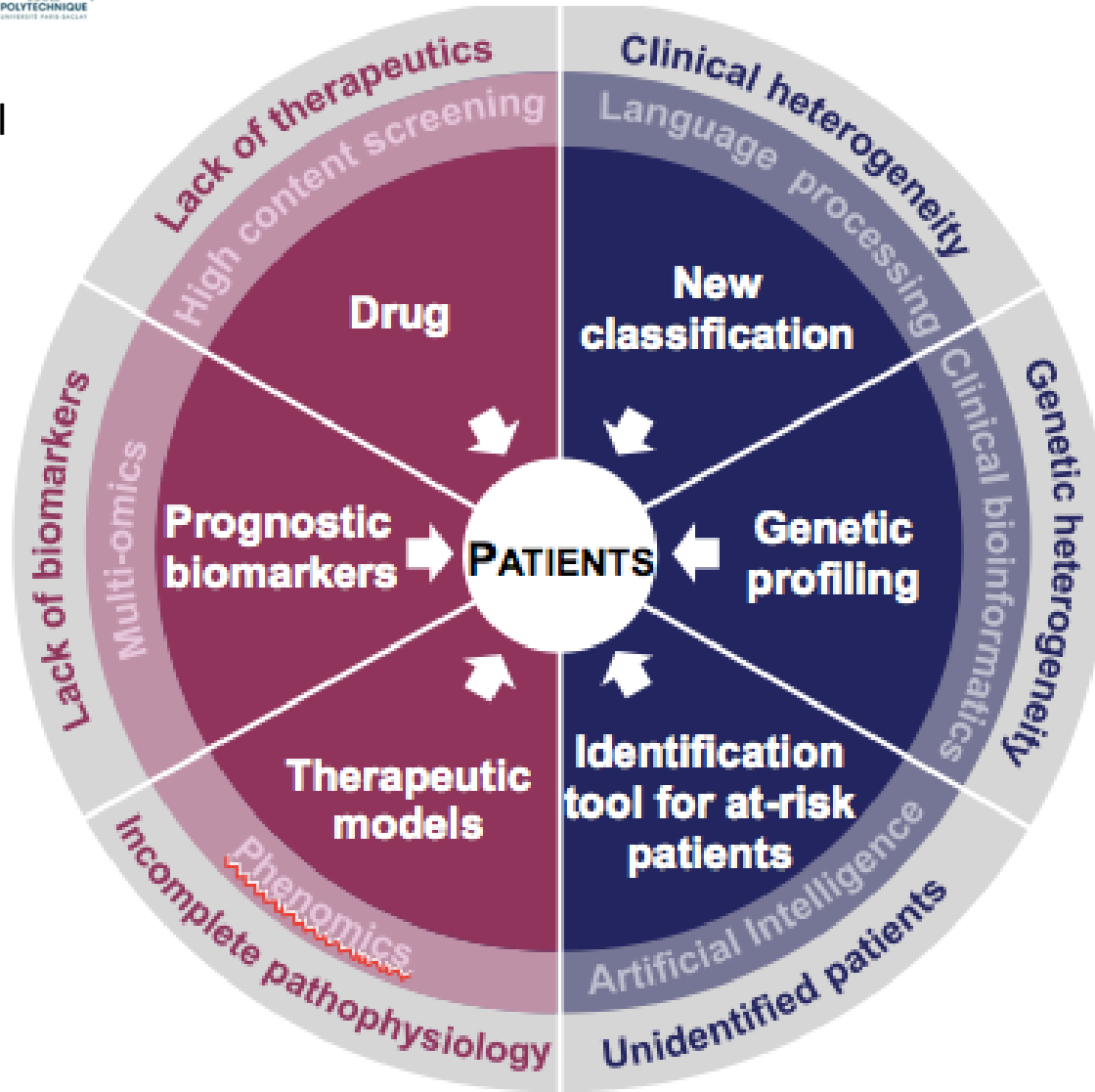


Goetz et al., 2010

Source : **C'IL-LICO Consortium**

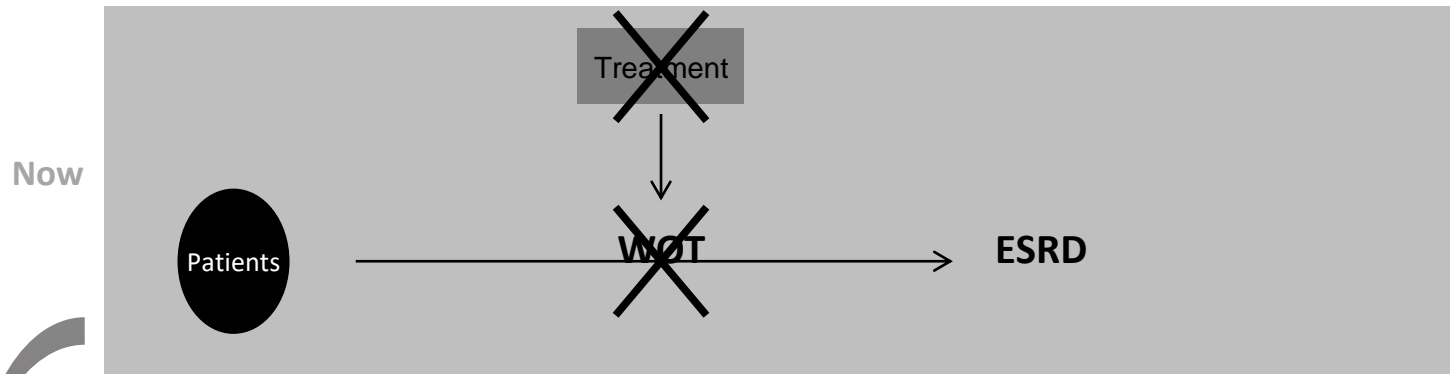
- › **DRUG** ready to enter clinical trial (active file of 25 patients to be recruited)

- › **CIL-BIOM:** Prognostic biomarker-based kit to predict patients at-risk for renal impairment and eligible for treatments

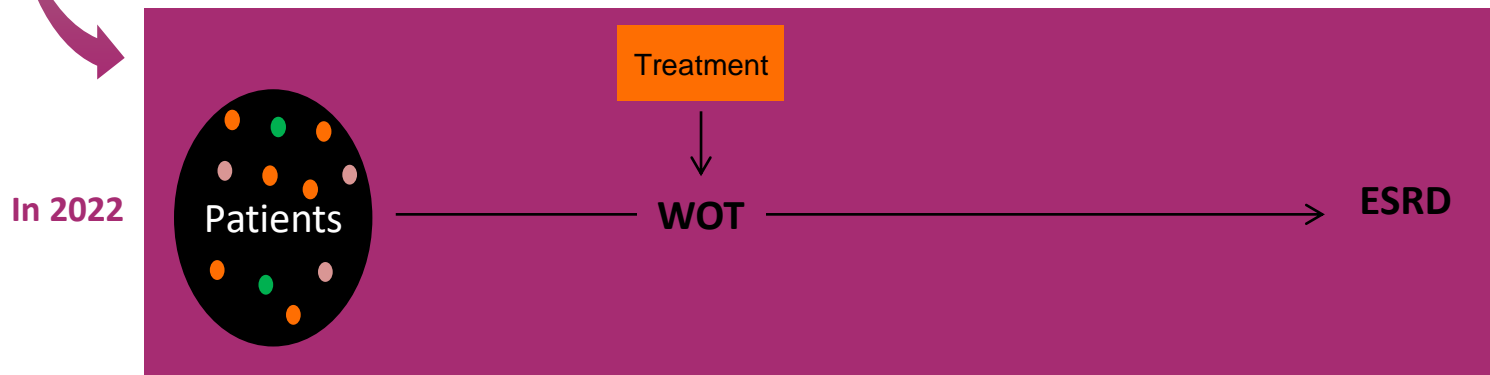


- › **CIL-DIAG2:**
- › Genetic diagnostic validation tool in clinical settings
- › **CIL-SMART:**
- › AI-based diagnostic decision-support tool

Providing the right treatment, to the right patient, at the right time:  
C'IL-LICO outstanding opportunity for ciliopathy patients.



- › **Detect undiagnosed patients**
- › Prognosticate the occurrence of comorbidities to **define the Window Of Therapeutics**
- › Develop **first-in-class** pathway-specific treatment and identify **patients eligible** for it



Source : **C'IL-LICO Consortium**

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